

## DEFENSE THREAT REDUCTION AGENCY 8725 JOHN J. KINGMAN ROAD, STOP 6201 FORT BELVOIR, VA 22060-6201

February 17, 2022

Shannon Murray U.S. Right to Know

Re: FOIA Case No.: 22-008

Dear Ms. Murray:

This is our final response to your Freedom of Information Act (FOIA) request perfected on October 6, 2021, and assigned FOIA case number 22-008 by the Defense Threat Reduction Agency (DTRA). You requested a copy of grant proposals, scientific and technical interim reports, reviewer response documents, policy-related documents, and email communications between DTRA staff and EcoHealth Alliance staff (email to or from the domain @ecohealthalliance.org, including attachments) about the following grants awarded by DTRA to EcoHealth Alliance, since January 2019: HDTRA12010026, HDTRA12010029, and HDTRA12110023.

Enclosed is a copy of the documents totaling 311 pages. These records are being released to you in part. Some information is being withheld under FOIA Exemption 6. Exemption 6 applies to information, which, if released, would constitute a clearly unwarranted invasion of the personal privacy of an individual. No fees are due as the assessable cost total \$25.00 or less.

Determinations for this interim release were made by the Initial Denial Authority (IDA), Mr. Earl Washington, Chief, Records Management, FOIA, and Privacy Act Division / DTRA Records Officer, Information Management and Technology Directorate, on behalf of DTRA. If you consider this decision to be an adverse determination, you may file a written appeal that is postmarked no later than 90 calendar days after the date of this letter to the Deputy Director, Defense Threat Reduction Agency, Information Management and Technology Directorate, ATTN: FOIA/PA Office, 8725 John J. Kingman Road, MSC 6201, Fort Belvoir, Virginia 22060. The appeal should reference the FOIA/Privacy Act case number, contain a concise statement of the grounds upon which the appeal is brought, and a description of the relief sought. A copy of this letter should also accompany your appeal. Both the envelope and your letter should clearly identify that a Freedom of Information Act and/or a Privacy Act Appeal is being made.

Should you have additional questions or concerns regarding this case, you may seek dispute resolution services from the DTRA FOIA Public Liaison or the Office of Government Information Services (OGIS). The DTRA FOIA Public Liaison, Mr. Mario Vizcarra, may be contacted by phone at (703)767-1792 or by email at dtrafoiaprivacy@mail.mil. The contact information for OGIS can be found at www.archives.gov/ogis.

Sincerely,

Eugene McGirt

Eugene McGirt FOIA/Privacy Act Specialist Freedom of Information/Privacy Act Office

Enclosure(s): As stated

From: Aleksei Chmura To: (b)(6) Cc: (b)(6) DTRA Ft Belvoir SI Mailbox SI-FMKA SPS Contracts; onr boston@navy.mil; Joe Riccardi Subject: [Non-DoD Source] Re: HDTRA1-20-1-0026 Award Date: Friday, September 25, 2020 8:59:49 AM All active links contained in this email were disabled. Please verify the identity of the sender, and confirm the authenticity of all links contained within the message prior to copying and pasting the address to a Web browser. Dear(b)(6) I confirm receipt. Many thanks most, Sincerely, -Aleksei Aleksei Chmura, PhD Chief of Staff EcoHealth Alliance 520 Eighth Avenue, Suite 1200 New York, NY 10018-4182  $+1.212.380.4473 \le \text{tel}: 1.212.380.4469 \ge \text{(office)}$ (b)(6) (mobile) Caution-www.ecohealthalliance.org < Caution-http://www.ecohealthalliance.org > EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation. On Sep 25, 2020, at 08:04 (b)(6) Caution-mailtd(b)(6) > wrote: Attached you will find award documentation for grant HDTRA1-20-1-0026. Please confirm once you have received this email. Thank you! V/rContractor

Office Phone: (571) 616-4277

Contract Specialist • Broadleaf

Defense Threat Reduction Agency (DTRA)

(b)(6)

<EHA India SOW.pdf><Terms and Conditions (DTRA-specific)\_14Sep2020 FINAL.pdf><HDTRA1-20-1-0026 Award.pdf>

From:	Aleksei Chmura (b)(6)	
To:	Dr. Jon Epstein; Ava Sullivan;(b)(6)	Joe Riccardi; (b)(6)
Cc:	(b)(6) Alison Andre; (b)(6)	JOS NICCATOL, [ * * * * * * * * * * * * * * * * * *
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justified and	t our India project budget is \$4,956,064.36. This is less than our original verified budget costs were slightly reduced compared to what was origin tp://Grants.gov > .	
Many thanks	!	
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Aleksei Chm Chief of Staff		
-	lliance Avenue, Suite 1200 IY 10018-4182	
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EcoHealth Al	lliance develops science-based solutions to prevent pandemics and prome	ote conservation.
On Sep Caution- <u>mail</u>	16, 2020, at 10:04 (b)(6)	
	re a question, while reviewing your budget I see a total budget of \$4,956, ignal. For clarification, what is the total proposed amount?	064.36 and \$4,964,011.70,
From: A	ginal Message Aleksei Chmura <chmura@ecohealthalliance.org <="" caution-<u="">mailto:chmur</chmura@ecohealthalliance.org>	ra@ecohealthalliance.org > >
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	Ion Epstein <epstein@ecohealthalliance.org <="" caution-<u="">mailto:epstein@ecohealthal livan@ecohealthalliance.org &lt; Caution-<u>mailto:sullivan@ecohealthalliance.org</u> &gt; &gt;:</epstein@ecohealthalliance.org>	
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Dear <sup>(b)</sup>	6)	
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Thonk	ou for your continued patience with us and our partners as we gathered the requeste	ad daaumantatiar
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Attache	d are the following files:	
	India KVASU budget justification v01.docx	
- DTR	. India KVASU supporting docs.pdf	
- DTR/	India HMJF budget justification v01.docx	
- DTR	India HMJF supporting docs.pdf	
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- DTR	India KGMU supporting docs.pdf	
- DIK	India KGMU budget justification v01.docx	
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- DTR	India EHA budget v01.xlsx	
- DTR	. India EHA Clarifications v01.xlsx	
- DTR	India EHA Questions and Responses v01.docx	
Note th	it we have included (last file listed above) a word document version of our response	es to the question
	iously, but these also are included in our Clarifications (xlsx) file.	•
Please	all anytime, if you have questions, require additional details, or would like us to rap	oidly direct you to
	documentation in our attached files.	nary ancer you a
iy specific	documentation in our attached mes.	
Many t	anks:	
-Alekse		

Aleksei Chmura, PhD Chief of Staff & Authorized Organizational Representative

EcoHealth Alliance 520 Eighth Avenue, Suite 1200 New York, NY 10018-4182 +1.212.380.4473 < tel: 1.212.380.4469 < tel: 1.212.380.4469 > > (office)(b)(6) > (mobile) Caution-Caution-www.ecohealthalliance.org < Caution-http://Caution-Caution-www.ecohealthalliance.org > < Caution-Caution-http://www.ecohealthalliance.org/ < Caution-Caution-http://www.ecohealthalliance.org/ > > EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation. On Sep 1, 2020, at 14:52, Ava Sullivan < sullivan@ecohealthalliance.org < Caution-mailto:sullivan@ecohealthalliance.org > <Caution-mailto:sullivan@ecohealthalliance.org < Caution-mailto:sullivan@ecohealthalliance.org >>> wrote: Thank you (b)(6) We will submit the full clarification by OOB tomorrow. We apologize for the late submission, we have run into a few obstacles related to COVID, among other delays. Despite these delays, we will be able to submit very shortly and look forward to sharing the information with this group. Thanks, Ava Sullivan On Sep 1, 2020, at 2:30 PM, Lyles, Latrice D CTR DTRA AL (USA) Caution-mailto(b)(6) Caution-mailto Good Afternoon, I will be picking up for Terrie. I know it was mentioned that the clarification questions/documents would be submitted Friday August 28, 2020. I wanted to follow up on that and make sure we get the documents in so we can proceed with awarding this grant. I hope to hear back from you soon. Thanks! -----Original Message-----From: Ava Sullivan <sullivan@ecohealthalliance.org < Caution-mailto:sullivan@ecohealthalliance.org > Sent: Wednesday, August 26, 2020 9:15 AM To (b)(6) Caution Caution Cc <Cautio Latrice <Cautio CIV DI Caution >>; Joe Riccardi <riccardi@ecohealthalliance.org < Caution-mailt Caution-mailto:riccardi@ecohealthalliance.org > < Caution-mailto:riccardi@ecohealthalliance.org <

Caution-mailto:riccardi@ecohealthalliance.org >>>; Nwani, Nkechiyere G CTR (USA)

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Aleksei MacDurian <chmura@ecohealthalliance.org <="" caution-mailto:chmura@ecohealthalliance.org=""></chmura@ecohealthalliance.org>
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Yes, I will do that.
Best wishes for the time ahead,
Ava Sullivan
Project Manager and Research Assistant
EcoHealth Alliance
520 Eighth Avenue – Suite 1200
New York, NY 10018
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EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation
On Aug 25, 2020, at 8:26 PM, Rodriguez, Terrie M CIV DTRA AL (USA) (b)(6)
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Good Evening,
Thank you very much for your e-mail and for the update. Kindly request that you please include all DTRA personnel contained on the "cc" line of this e-mail for all clarification responses, as I will be out on maternity leave beginning Monday 8/31.
Very Respectfully,
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Contract Specialist Defense Threat Reduction Agency
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Original Message
From: Ava Sullivan <sullivan@ecohealthalliance.org <="" caution-mailto:sullivan@ecohealthalliance.org=""></sullivan@ecohealthalliance.org>
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Sent: Tuesday, August 25, 2020 8:21 PM -{b)(6)
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Cc: Jon Epstein <epstein@ecohealthalliance.org <="" caution-mailto:epstein@ecohealthalliance.org=""></epstein@ecohealthalliance.org>
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authenticity of all links contained within the message prior to copying and pasting the address to a Web browser.
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Helld (b)(6)
Forgive the late response. We were expecting to submit our full packet of clarifications today. However, due to
unforeseen circumstances including issues with COVID-19 in-country, we will require a few extra days. I hope it is
acceptable that we submit this on Friday, the 28th.
Thank you for your understanding, and we look forward to providing all the requested materials, Ava

Caution-

On Thu, Aug 20, 2020 at 1:38 PM (b)(6)

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> wrote:	
Hi Ava,	
Thanks so much for your e-mail. We look forward to receiving the responses. Have	a great day!
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A kind hello,
l am jumping on this thread on behalf of EcoHealth Alliance. We expect to submit the clarifications by Tuesday, August 25. I do hope this is agreeable.  Thank you,  Ava
Ava Sullivan
Project Manager and Research Assistant
EcoHealth Alliance
520 Eighth Avenue – Suite 1200
New York, NY 10018
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EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation

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Emerson CIV DTRA COOP THRT REDUCT (USA)" (b)(6)

Ca Ca

Greetings Dr. Chmura,

I just wanted to touch base to see if it was possible to receive a status update on the outstanding clarifications. We were hoping to reach resolution of these clarifications by mid-month in order to meet a prospective award date of 31 August however, we completely understand that EHA has its own competing priorities and that EHA may be waiting for responses from its own prospective subcontractors for this effort. Any update you might be able to share will help us shape expectations within our organization on a revised prospective award date.

Thank you very much and we look forward to continuing to work with you and your team! Have a wonderful day!

Very Respectfully,

(b)(6)

Contract Specialist
Defense Threat Reduction Agency

(b)(6)

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Original Message	
From: Aleksei Chmura <chmura@ecohealthalliance.org <<="" td=""><td></td></chmura@ecohealthalliance.org>	
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Subject: Re: [Non-DoD Source] FRBAA14-6-2-0454
All active links contained in this email were disabled. Please verify the identity of the sender, and
confirm the authenticity of all links contained within the message prior to copying and pasting the address to a Web
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Dear(b)(6)
I confirm receipt. Thank you for this notice and the additional clarification requests. We are working
through the requested details and will get them back to as swiftly as possible.
through the requested details and with get them back to as swiftly as possible.
Many thanks!
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Aleksei Chmura
Chief of Staff &
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EcoHealth Alliance
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wrote:	
	Good Morning Dr. Chmura.

I just wanted to follow up to confirm that you did receive the below e-mail. I hope you had a nice weekend. Thanks in advance and have a wonderful day!

Very Respectfully, Contract Specialist Defense Threat Reduction Agency <Cautio (b)(6) Caution Caution Caution < Cautio Caution ----Original Message-----From: (b)(6) Sent: Monday, August 3, 2020 3:15 PM To: 'Aleksei Chmura' <chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org > <Caution-mailto:chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org >> < Caution-Caution-mailto:chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org > > < Caution-Caution-Caution-mailto:chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org > < Caution-Caution-mailto:chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org > > > < Caution-Caution-Caution-Caution-mailto:chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org > < Caution-Caution-mailto;chmura@ecohealthalliance.org < Caution-mailto;chmura@ecohealthalliance.org > > < Caution-Caution-Caution-mailto:chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org > < Caution-Caution-mailto:chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org > > > > < Caution-Caution-Caution-Caution-mailto:chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org > < Caution-Caution-mailto:chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org > > < Caution-Caution-Caution-mailto:chmura@ecohealthalliance.org < Caution-mailto:chmura@ecohealthalliance.org > < Caution-

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Subject: RE: [Non-DoD Source] Re: FRBAA14-6-2-0454

Importance: High

Good Afternoon Dr. Chmura,

I hope that you enjoyed a nice weekend. We have some additional clarification question to add to our initial clarification request. Please let me know if you have any questions or concerns. Thank you in advance!

- 1. The proposal contains a 36.8% Indirect Rate for EHA however the latest NICRA for FY21 dated April 1, 2020 has 35.84% from 7/1/20 to 6/30/21. Please provide some additional information to support the use of 36.8% or update the proposal, if needed.
- 2. Please provide quotes, website link, or basis of estimate to substantiate cost proposed for Luminex machine in Y1 (EHA) and freezer in Y1 (KGMU)?

- 3. For all proposed travel which includes a conference registration cost, please provide the name of conference and the cost of registration.
- 4. For all proposed materials and supplies for which the budget justification refers to quotes or previous purchases, please provide any documentation you have available to support the proposed costs or to provide additional detail into the basis of estimates for the items.
- 5. If possible, request that KGMU and KVASU provide a copies of its institutional salary grades to support its proposed labor?
- 6. Please provide a breakout of all of HJF's proposed travel. (# of travelers, locations, per diem, dates, transportation, etc.)
  - 7. Please provide documentation to support HJF's proposed cost for the AKTA service agreement.
- 8. If possible, request that HJF please provide directly to DTRA an updated copy of its provisional indirect cost and fringe benefit rates for FY20, if available?
- 9. Please provide documentation/basis of estimate to support KVASU's proposed cost \$,1000 per year is to contribute to organizing the ongoing One Health stakeholder engagement meeting for venue and hosting costs. (ex. Breakout to support \$1,000, what is included in the cost, etc.)

We are hoping to reach resolution of these clarifications (and those previously submitted) by midmonth, if possible, in order to meet a prospective award date of 31 August. Should your team run into any issues, please let us know. Thank you very much and we look forward to continuing to work with you and your team! Have a wonderful afternoon!

Very Respectfully, (b)(6) Contract Specialist Defense Threat Reduction Agency <Caution(b)(6) Caution-Caution-Caution-< Cautio Caution-Caution-Caution-Caution-Caution-Caution-Caution-Caution-Caution-Caution-Caution-Caution-Caution-Caution-

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----Original Message-----
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Subject: [Non-DoD Source] Re: FRBAA14-6-2-0454 Importance: High

All active links contained in this email were disabled. Please verify the identity of the sender, and confirm the authenticity of all links contained within the message prior to copying and pasting the address to a Web browser.

Dear<sup>(b)(6)</sup>

Apologies for my slow reply. We are delighted to read your email!

I have cc'ed our Finance Manager Joe Riccardi on this email as well.

We will work through your requests this week and get responses back to you and Latrice as rapidly as possible.

I hope this email finds you and yours safe and healthy most,

Sincerely!

-Aleksei

Aleksei Chmura Chief of Staff & Authorized Organizational Representative

EcoHealth Alliance 520 Eighth Avenue, Suite 1200 New York, NY 10018-4182

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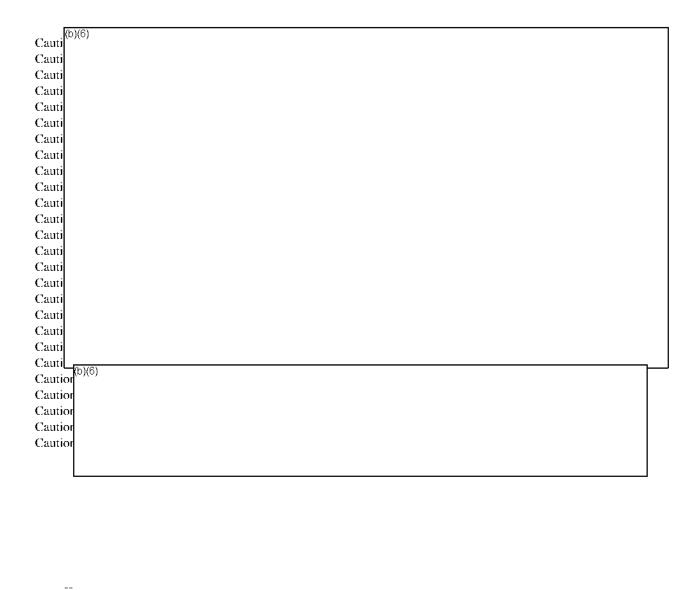
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	EcoHealth Alliance develops science-based solutions to prevent pandemics and promote
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wrote:
Greetings Dr. Epstein and Dr. Chmura,
CONCRATULATIONS V DTDA Dood Accordances
CONGRATULATIONS! Your proposal, submitted under DTRA Broad Agency Announcement HDTRA1-14-24-FRCWMD, Proposal Number: FRBAA14-6-2-0454, has been selected as one that the Defense
Threat Reduction Agency (DTRA) expects to award. PLEASE DO NOT BEGIN WORK as any award is subject to
(1) successful negotiations between our contracting division and your organization and (2) the availability of funds.
This notification must not be construed as an obligation on the part of the Government; only duly authorized
procurement personnel may commit resources. Also, this notification must not be used as a basis for accruing costs
to the Government prior to award.
Myself and (b)(6) will be your POCs until the award has been made. We will work
directly with your office to discuss any clarifications required to support your proposal. Below are some initial
clarifications for EHA's review and response.
1. Does EHA have its proposed budget available in MS Excel format and if so, would EHA be
willing to share an executable file with DTRA for ease of review?
2. Does EHA have quotations available to support any of the proposed costs (ex. Proposed
equipment or materials, sub-award)? If so, please provide that information to DTRA.  3. The proposal contains the procurement of a Luminex 200 Machine in Year I for use
throughout the project. Please provide additional details as to how EHA proposes to mitigate the expected foreign
tax burden for the local purchase, or importation of this piece of equipment.
4. The proposal contains the procurement of three laptops for this project, however it was noted
in the budget justification that only one will be used to perform Luminex reads. Please provide additional
justification to support the procurement of the additional two laptops.  The proposal contains a sub-award to LHE, which includes intermediated Places provides
5. The proposal contains a sub-award to HJF, which includes international travel. Please provide additional details which addresses the specific nature and frequency of international travel and how it relates to the
requirements for project execution.
6. The proposal narrative states that Dr. Linfa Wang of Duke-NUS will serve as a Technical
Advisor for this project, yet his role is not adequately characterized. There is no sub-award associated with Duke-
NUS, nor are Dr. Wang's responsibilities, projected time, or salary for his role captured in the technical proposal,
Statement of Work, or budget proposal. DTRA requests clarification on the role of Duke-NUS in the project, and
subsequent submission of any relevant sub-award documentation, if appropriate.
Please let us know if you have any questions. Thank you and we look forward to working with you.
Very Respectfully,
(b)(6)
Contract Specialist
Defense Threat Reduction Agency
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Contract Specialist
Defense Threat Reduction Agency

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Ava Sullivan
PREDICT-2 Operations Coordinator
Country Liaison, India

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(b)(6) (mobile)

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EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation

Ava Sullivan

Project Manager and Research Assistant

EcoHealth Alliance 520 Eighth Avenue – Suite 1200 New York, NY 10018

EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation

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						31 Jan 2021	31 Jan 2027	31 Jan 2023	31 .an 2024	31 Jan 2025	
MONTH	r-5					VI	Y2	YJ	OAT	OY2	4%
Y2 Y3	ÜYI	L OY2	BASE	A. SENIOR KEY PERSONNEL							
3 3		.4	\$ 170,887,50	Epstein, Ion	Principal Investigator	5 42,721.88	5 44,/17.14	\$ 45,764.74	\$ 47,866.51	5 49 024,84	3 279 094,60
•				Fringe, Epsieir		5 15,721,65	\$ 16,271.01	5 16,841.42	\$ 17,450,87	5 18 040 95	5 34 306 81
2 2	2	2	\$ 00,730,03	Ross, Noam	Date enalyst	5 15,289,34				5 [7 545 44	
				Fringe, Ross		5 5,626 66		5 6,027,42	5 6,238.38		5 30,172.77
		Į.		TOTAL SENIOR KEY PERSONNEL	ļ	\$ 79,260,02	5 92,137,62	5 95,012,44	5 97.987.89	\$ 91,267,45	\$ 429,565,41
	L	l		B OTHER PERSONNEL	l						
12 12	12	12	\$ 71,460,60	TBC	Post Dot Research Asst		\$ 75,899.90	\$ 76,485.47	\$ 79 [62.46	5 81,958,14	5 582 880 06
				Forige, TBD		5 26,275 20	5 27.194.83			5 30,151.40	5 140,899.86
12 12	12	12	5 55,109.75	T6C	MSc Program Assistant					5 67,829.80	
				Fringe, TBD		\$ 21,752,39	\$ 22,513,72	\$ 23,301,70	\$ 24,117,26	\$ 24,961,37	\$ 116,646,44
				TOTAL OTHER PERSONNEL		5 175,537,54	\$ 184,786.14	\$ 191,753.66	\$ 197,947.54	\$ 204 B75.70	3 957 400.38
				Total Salary		5 188,521.46			5 209,017.12		5 1.010,919.91
			26,80%	Total Fringe		\$ 69,275,90					\$ 372,025,89
				Total Salary + Fringe	ļ.	\$ 257,897,36	\$ 266,923,77	\$ 276,266,10	\$ 295,935,41	\$ 295,943,15	\$ 1,382.965.80
				C FQUIPMENT							
				Luminez Bio-Plex* 200 with wash etc		5 55,719.50		5 .	5 .	5 -	5 55 710 50
				TOTAL EQUIPMENT		5 55,719 50	5 -	5 .	5 -	5 -	5 55,719.50
				D. FRAVET							
				1 Domestic		\$ 11,981.90	\$ 11,561.90	\$ 11,581.90	\$ 11,581.90	3 11,581,90	3 37 909,50
				2 International S		5 34,515.40					5 172 577 00
				TOTAL TRAVEL		5 46,097,30	5 46,097,30	5 46,097,30	5 46,097.30	5 46,097.30	5 230,486.50
				F. PARTICIPANT/TRAINER SUPPORT COSTS							
				1 Tuition/Fees/Hearn Insurance		5 ·	\$ .	5 .	5 .	5 .	5 -
				2 Supenda		2 .	5 .		5 .	5 .	5 -
				J Trave		S .	5 -	s .	5 -	5	5 -
				4 Subustence		ş	S	5	S	>	\$
				5 Other		5	5	5	5	5	3
				Total Participant/Trainee Support Costs		5 .	5 .	5 .	5 .	5	5 -
				F. OTHER DIRECT COSTS							
				1 Materials and Supplies							
				Lagtops for Luminex Reads & data analy	2	\$ 4,799.00		5	S	5	\$ 4,798.00
				Harp Irap		5 4,433 00					3 4 33 3,000
				Cryn-Britis		5 1,358.00	5 .	5 .	5 .		5 1,358,00
				Dry Shipper	2	5 4,753 00	5 -	s .	5 -	5	5 4,753.00
				Needles, Synnges, Tubes		\$ 1,866,96	\$ 1,966,96	\$ 1,966,96	\$ 1,966,96	\$ 1,966,96	\$ 9,324,80
				Micro Centrifuge	2	\$ 1,686,06			\$ 1,686,06		\$ 2,372,12
				Total Materials and Supplies		\$ 18,795.07	5 1,861: 91:	\$ 1,866:96	\$ 3,953.07	5 1 Rec. 96	3 27,948,92
				2 Publication Costs		5 ·	5 .	5 .	\$ [0.760,00	5 (0.760.00	5 21,520,00
				1 Conscitant Services							
S 6	6		\$ 48,240,60	Bhatia, Rajesh	Car sultant 1	\$ 20,100,00		\$ 24,120,00	\$ 24,120,00	\$ 24,120,00	\$ 112,560,00
5 6	b	b	\$ 48,740,00	Zarnanan, Arun	Consultant 2	\$ 70,100,00	5 20,100,00	\$ 74,120,00	\$ 74,120,00	\$ 74,120,00	3 112 560.00
-				Lotal Consultant Services		\$ 40,700,00	\$ 40,700,00	\$ 45,740,00	\$ 45,740,00	\$ 48,740,00	\$ 275,120,00
				4 ADP/Computer Services		5 .	5 .	5 .	5	5 .	5 .
				5 Subawards/Consortium/Contractual C		5 398,167.58	5 509,772 13	5 490,965,93	5 459,623.57	5 450,220,46	5 2,308,749.67
				6 Equipment or Facility Rental/Oser Fee		ş	S	S	5	5	ś
				7. Alterations and Bonovations		5	IS	-	١٢ -	3	3
				8 Onnei		5 .	5 -	5 .	5 .	5 -	5 -
				Total Other Direct Costs		5 457,162,60	\$ 551,859.09	\$ 541,072,80	5 522 176 59	5 511 087 42	5 2,589 398 50
				G. OIRECT & MODIFIED DIRECT COSTS							
				Direct Costs		\$ 810,876,76		\$ 863,436,29	\$ 854,209,30		\$ 4,252,510,39
					I	\$ 862,589.68	\$ 895,088.03	\$ 372,470,36	\$ 394,585,73	3 402 907.41	3 1,888 041.27
				Modified Direct Costs							
				H INDIRECT COSTS							
			IDC Rate>	H INDIRECT COSTS 35.54%		5 130.095 50				5 144,402.02	5 676,673.97
			IDC Rate >	H INDIRECT COSTS  35.84%  35.84%		5 21,842.70	5 3,922.70	5 1.114.61	5 -	5 -	5 26,880.00
				H INDIRECT COSTS  35.54%  35.54%  Total Indirect Costs				5 1.114.61		5 144,402.02 5 \$ 144,402.02	
				H INDIRECT COSTS  35.54%  35.54%  Total ling cot Costs  L Total DIRECT AND INDIRECT COSTS		S 23,842.70 S 151,939,20	5 3.922.70 5 131.186.25	5 1.114.61 5 134.607.99	S 141,419,S3	5 144.402.02	5 26,880.00 \$ 702,553,97
				H INDIRECT COSTS  35.54% 35.54% 35.54% Total Indirect Costs LIGITAL DIRECT AND INDIRECT COSTS Direct - Indirect		5 21,842.70	5 3,922.70	5 1.114.61 5 134.607.99	S 141,419,S3	5 144.402.02	5 26,880.00
				H INDIRECT COSTS  35.54%  35.54%  Total ling cot Costs  L Total DIRECT AND INDIRECT COSTS		S 23,842.70 S 151,939,20	5 3.922.70 5 131.186.25	5 1.114.61 5 134.607.99	S 141,419,S3	5 144.402.02	5 26,880.00 \$ 702,553,97
				H INDIRECT COSTS  35.84%  35.84%  Total lina rest Costs L (GIAL DIRECT AND INDIRECT COSTS  0.460 + Indirect 1 (TES)		S 23,842.70 S 151,939,20	5 3.922.70 5 131.186.25	5 1.114.61 5 134.607.99	S 141,419,S3	5 144.402.02	5 26,880.00 \$ 702,553,97
				H INDIRECT COSTS  35.54%  35.54%  Total indirect Costs Fried-1-Winect 1 TEES Free Free Free Free Free Free Free Fre		\$ 23,842,70 \$ 153,939,20 \$ 068,814,06	\$ 3,922.70 \$ 131,186,25 \$ 006,246.41	\$ 1,114,61 \$ 134,607,99 \$ 938,344.27	5 141,419,53 5 935,626,83	5 144,402,02 5 297,520,89	5 26,880.00 \$ 702,553.97 5 4,956,064.56
				H INDIRECT COSTS  35.84%  35.84%  Total lina rest Costs L (GIAL DIRECT AND INDIRECT COSTS  0.460 + Indirect 1 (TES)		S 23,842.70 S 151,939,20	\$ 3,922.70 \$ 131,186,25 \$ 096,046.41	5 1.114.61 5 134.607.99	S 141,419,S3	5 144,402,02 5 297,520,89	5 26,880.00 \$ 702,553,97
				H INDIRECT COSTS  5.5 64%  5.5 64%  Total Indirect Costs  L Idial Direct And In Direct Costs  E-vice - Indirect  1 FEES  Five  K. TOTAL COSTS AND FEES  Intal Costs  Anaport to Scientes gov		\$ 23,842,70 \$ 153,939,20 \$ 068,814,06	\$ 3,922,70 \$ 131,186,25 \$ 006,046,41 \$ 996,046,41	\$ 1,114,61 \$ 134,607,99 \$ 938,344.27	\$ 141,419,53 \$ 205,626,83 \$ 995,626,83	5 144,402,02 5 297,520,89 5 997,529,89	5 26,880.00 \$ 702,553.97 5 4,956,064.56
				H INDIRECT COSTS  35.564%  35.544%  Total and set Costs L I COLAT DIRECT AND INDIRECT COSTS E-490 - Indirect J FEES Fue K TOTAL COSTS AND FEES I I COSTS COSTS AND FEES		\$ 21,842,70 \$ 151,939,20 \$ 068,814,06 \$ 968,814,96	\$ 3,922,70 \$ 131,186,25 \$ 006,046,41 \$ 996,046,41	\$ 1,119,61 \$ 134,607,99 \$ 938,244.27 \$ 938,044,27	\$ 141,419,53 \$ 205,626,83 \$ 995,626,83	5 144,402,02 5 297,520,89 5 997,529,89	5 26,880.00 \$ 702,523,97 5 4,956,064,56 5 4,956,064,46
				H INDIRECT COSTS  5.5 64%  5.5 64%  Total Indirect Costs  L Idial Direct And In Direct Costs  E-vice - Indirect  1 FEES  Five  K. TOTAL COSTS AND FEES  Intal Costs  Anaport to Scientes gov		\$ 21,842,70 \$ 151,939,20 \$ 068,814,06 \$ 968,814,96 \$ 970,566,84	\$ 3.922.70 \$ 131.186.25 \$ 036.046.41 \$ 946,046.41 \$ 595.332.63	\$ 1.114.61 \$ 134.607.99 \$ 936.344.27 \$ 946,044.27 \$ 996.645.5	\$ 141,419,53 \$ 205,626,83 \$ 995,626,83	5 144,402,02 5 297,520,89 5 997,529,89	5 26,880.00 \$ 702,523,97 5 4,956,064,56 5 4,956,064,46
				H INDIRECT COSTS  35: 564%  35: 564%  Total Indirect Costs L I CITAL DIRECT AND INDIRECT COM15 E-9ct - Indirect J FEES Fee K, TOTAL COSTS AND FEES Indirect Seeson Significant Costs Seeson Significant Costs Seeson Significant Significa		\$ 23,842,70 \$ 151,839,20 \$ 068,814,06 \$ 968,814,96 \$ 570,566,84 2	\$ 3.922.70 \$ 131.186.25 \$ 036.046.41 \$ 946,046.41 \$ 595.332.63	\$ 1.114.61 \$ 134.607.99 \$ 938.344.27 \$ 946,044.27 \$ 939.665.56	\$ 141,419,53 \$ 995,628,63 \$ 995,628,63 \$ 995,628,64 \$ 996,965,85	5 997 520 88 5 997 520 88 5 997 529 88 5 997 529 88	5 26,880.00 \$ 702,523,97 5 4,956,364,56 \$ 4,956,364,56 \$ 4,955,527,86
				H. INDIRECT COSTS  5.5 64%  5.5 64%  Total Ind. ver. Costs  L. ICITAL DIRECT AND IN DIRECT COSTS  1. TOTAL COSTS AND FEES  FOR  T. TOTAL COSTS AND FEES  Intol Costs  SUPPLYMENT OF SUPP		\$ 21,842.70 \$ 151,839.20 \$ 058,814.06 \$ 968,814.96 \$ 570,566.84 \$ 2 \$ 211,609.16	\$ 3.922.70 \$ 131.180.25 \$ 006.046.41 \$ 996,046.41 \$ 996,046.41 \$ 995.392.63 \$ 248.014.35 \$ 248.014.35	\$ 1.114.61 \$ 134.607.99 \$ 938.344.27 \$ 945,044.27 \$ 995.66 \$6 \$ 229.208.15	\$ 141,419,53 \$ 995,618,63 \$ 995,628,63 \$ 995,628,65 \$ 998,965,86 \$ 210,401,94	5 144,402,02 5 997,520,89 3 997,529,89 5 998,985,95 5 200,998,84	5 26,880.00 \$ 702,523,97 5 4,956,364,56 \$ 4,966,364,56 \$ 4,965,527,86 \$ 1,100,221,44
				H. INDIRECT COSTS  5.5 64%  5.5 64%  Total Ind. ver. Costs  L. CILIAL DIRECT AND IN DIRECT COSTS  L. CILIAL DIRECT AND IN DIRECT COSTS  Free C. Indirect  J. TEES  Free  I. TOTAL COSTS AND FEES  India Costs  SUPPLYMENT SO		\$ 23,842,70 \$ 153,939,20 \$ 058,814,06 \$ 968,814,96 \$ 570,566,84 2 213,699,16 \$ 175,644,40 \$ 175,644,40	\$ 3,922.70 \$ 131,180,25 \$ 006,046.41 \$ 006,046.41 \$ 996,046.41 \$ 595,392.63 \$ 248,014.35 \$ 750,817.76 \$ 10,945.07	\$ 1.114.61 \$ 124.607.99 \$ 936.244.27 \$ 995,044.27 \$ 990.666.56 \$ 226.209.15 \$ 750,814.76 \$ 10,445.07	5 141,419,53 5 141,419,53 5 995,618,63 5 995,628,65 5 210,401,94 5 26,776,60 5 10,445,07	5 144.402.02 5 997.520.80 3 997.520.80 5 998.985.95 5 206.998.84 5 48.776.60 5 10.945.07	5 26,860,00 \$ 702,533,97 5 4,956,564,56 \$ 4,956,064,56 \$ 4,956,527,85 \$ 1,106,221,42 \$ 1,106,221,42 \$ 1,106,221,42 \$ 1,106,221,42 \$ 1,106,221,42
				H INDIRECT COSTS  35.84%  35.84%  Total Indirect Costs L ICITAL DIRECT AND INDIRECT CEATS E-regs - Indirect Free Linding JEES Free K. TOTAL COSTS AND FEES LINDIN CEATS APPOUND GRANTS GOV SURMANNESS KSEYU HAND		\$ 23,842.70 \$ 151,939.20 \$ 068,814.06 \$ 968,814.96 \$ 770,568.84 2 \$ 211,609.10 \$ 10,844.00 \$ 380,67.56	\$ 3,922.70 \$ 131,186.25 \$ 036,046.41 \$ 996,046.41 \$ 993,322.63 \$ 248,014.35 \$ 10,444.02 \$ 50,772.13	5 1.114.61 5 134.607.99 5 936.344.27 5 936.344.27 5 995.66 56 5 299.208.15 6 200.817.76 6 10,446.07 5 400.856 53	\$ 141,419,53 \$ 295,626,63 \$ 295,626,63 \$ 295,626,64 \$ 296,265,65 \$ 210,401,94 \$ 26,276,60	5 144,402,02 5 297,520,83 5 297,520,83 5 297,520,83 5 298,885,25 5 200,998,84 5 287,74,60 5 110,944,00 5 150,200,45	5 26,880,00 5 702,533,97 5 4,956,084,56 5 4,956,084,56 5 4,956,527,85 5 1,106,221,44 5 1,114,793,17 5 39,271,47 5 1,268,740,67
				H. INDIRECT COSTS  5.5 64%  5.5 64%  Total Ind. ver. Costs  L. CILIAL DIRECT AND IN DIRECT COSTS  L. CILIAL DIRECT AND IN DIRECT COSTS  Free C. Indirect  J. TEES  Free  I. TOTAL COSTS AND FEES  India Costs  SUPPLYMENT SO		\$ 23,842,70 \$ 153,939,20 \$ 058,814,06 \$ 968,814,96 \$ 570,566,84 2 213,699,16 \$ 175,644,40 \$ 175,644,40	\$ 3,922.70 \$ 133,186,25 \$ 936,548.41 \$ 996,348.41 \$ 995,332.63 \$ 248,014,35 \$ 248,014,35 \$ 10,444.02 \$ 505,772.13	5 1.114.61 5 134.607.99 5 238.344.37 5 938.344.37 5 939.565.55 5 239.208.15 5 240.815.45 5 10.345.07 5 40.365.23 71	5 141,413,53 5 235,518,83 5 235,518,83 5 298,965,85 5 210,401,94 5 2,67,6,67 5 10,445,07 5 450,513,57 ONI	5 144,402,02 5 297,512,83 5 297,512,83 5 298,985,25 5 200,998,84 5 202,796,63 5 103,445,07 5 455,212,45 072	5 26,860,00 \$ 702,533,97 5 4,956,564,56 \$ 4,956,064,56 \$ 4,956,527,85 \$ 1,106,221,42 \$ 1,106,221,42 \$ 1,106,221,42 \$ 1,106,221,42 \$ 1,106,221,42

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	Т	71	П	72	_	73	_	011	Г	012	TOTAL
. H4€	ļs.	862,989,68	5	355,088,03	١,	377,470.36	s	394,585.73	\$	402 907.41	3-1,888-041.27
HM IF	5	175,614.35	5	250,812.76	5	250,812,76	5	238,276,60	5	238 276 60	5 L155 793 12
KGMU	S	211,608 16	5	248.014.35	5	229,208.15	S	210,401.54	S	200,998.84	5 1.100,231.44
KVASU	Ş	10.945.02	Ş	10,949 02	ş	10,945.02	Ş	10,945.02	Ş	10.945.02	\$ 54,725,11
EllA in Country	ş	289,740,50	ş	221,626 14	ş	234,337.31	Ş	251,618.03	Ş	254,925,84	\$ 1,252,277.82

				COLAT	W/	LLOCATIONS (	[5]				
		71	Г	72		73		011	Г	OY2	TOTAL
JS	5	238,624.77	5	374,055 R4	b	378,727.01	. \$	371,025.51	S	376 029,38	3 1,738 467.57
NDIA	15	361,130.83	١	528,277.16	١,	527,177.12	٠ş	520,601.63	ls.	514 516.64	3,7,646,617.57
TOTAL	5	750,775 10	5	902,278.00	5	300,854 L5	5	801,627,14	5	800 545 72	5, 4,385,080,09
	-										

		40UN*1	RY ALLOCATIONS (5	vi		
:	Y1	177	14	071	(197	101AI
J5	30%	41%	42%	42%	42%	4,3%
NOIA	70%	59%	58%	53%	58%	63%
TOTAL	1 00	1.00	1 00	100	1.00	1.00

	18	TERN	ATIONAL TRAVE	L						
	Y1	П	-3		-)	Г	0-1	Г	()-1	TOTAL
NYC ↔ INDIA (DEL->EKO) - Meetings						Г		Г		
Hotel Max Per Green 3	29580	5	190(8)	5	196.00	5	190480	5	197.00	
Fertil Hats, 13	2,345,00	\$	2,345.00	\$	2,548,00	5	2,345,00	5	2,548,00	
Meals and Indicental Expenses Per Diem 18	74 69	Ś	74 69	Ś	74 00	5	74 60	Ś	74.90	
Total Mifs ()	845.00	9	\$99.00	5	494.00	5	499.00	5	994.00	
Flight (At male, (NYCk-2)(EL4-5(AO) - 2	1,414	5	1,414	5	[4]1	s	1,414	5	1 4   4	
port io/from Airbari (\$71/NYC, \$1.60 LKD)   6	-45 20	S	.45.20	S	145.20	5	.45.20	S	145.20	
Daily Transport in (kG (51 80) _ )	1.70	5	1.70	5	155	5	1.70	5	1.50	
No Days	ju		t)		j.d	L	Li		Li	
No. Nights	13		19		13		13		13	
Not Tups par Yes:					- 1	Г	4	Г	i,	
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Teta é	30 eSp 40	S	30 e5p 40	S	33juše 43	5	30 e55 40	Ś	34,63e.44	\$153,282
International Conference										1
Hotel Max Per Dien: 1	13700	5	15700	5	88.05	1	157103	5	519.00	
Total Hai- 5	101789	1.	10100	ĺ.	1,017.00	t	LCLUS	ĺ.	1,017.00	
Meals and Image et al Expenses Per Green 12	172 60	I.	173.00	I.	177.00	š	173 60	ĸ	177.00	
Tota MIEs !	20,000	4	20,000	5	7.40 0.4	14	70,000	1	7,40,00	
E gl : Est mat- 3	1.90	5	1.3%	5	1,350	t	1.250	ĺ.	1.00	
sport te/from Airport IS/1/NYC, \$75 GVA; 1	292.63	ŝ	292 03	ŝ	292.00	ŝ	292.00	ķ	292.00	
Daily Transpart in GVA	370	1	N/A	1	976	Ė	N/A	۲	N/A	
R-gorratus y	985	5	580	5	500	5	380	5	500	
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No. Nights	- 1	-	- 1	-	1	Г	٠,	Т	- 1	
Not Europe per Year			1			Г	- 1	Т	- 1	
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Tota !	535700	1/	5,457,000	5	1,855,00	17	535710	5	1,855.00	5 16,265

		DOMEST	IC TI	RAVEL							
	Г	ΥL	Г	43	Г	γ)	Г	0-1	_	1072	TOTA
NYC to DC Meesings											
Hat- Mai Pri D- ii	5	256	5	2)11	2	256	5	20	5	236	
Total motel	5	912	\$	512	7	212	5	512	s	5 (2	
Micais and Incidental Expenses Fer Diem	s	71	Ś	71	Ξ	71	Ś	71	5	71	
Tala MIF.	5	1.78	5	179	7	1.76	5	175	١.	179	
Oligat Transportation (stimoty)	5	127	\$	127	7	127	5	127	s	127	
al Transpari talmom Airport (\$71,00/NYC, \$98,90/WAS).	s	283	Ś	76 ii	1	280	s	287	5	250	
Daily Transpart or DC (\$14.65)	5	ar.	5	79	7	96	5	K5.	3	- 19	
Ker Cays		3		3		3	ı	3	:	1 [	
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hual Domestic Presentation/Conf. for Pl or 1 Sr. Personni											
Dell-, Man Per Di-, m	5	256	\$	276	7	256	5	250	s	236	
Tail: Hotel	5	1026	5	1,024	>	1.026	5	110/4	4	1,004	
Mice's And to cliente. Expenses Per Deilin	5	71	5	71	7	71	5	- 4	١.	- 11	
Tala MI(s	s	178	\$	128	10	178	s	178		178	
Fight/Train Transportation Estimate	5	197	5	177	5	197	5	177	4	107	
al Transport to/from Algaet (57) 00/NYC, \$68 50/T60).	5	290	5	2.40	7	290	5	260	. 5	248	
Daily Transport in TBD (514-65)	5	147	\$	147	3	147	5	141	s	117	
Registration	5	403	5	900	>	2000	5	50.0	4	5.0	
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[TOTAL DOMESTIC TRAVS. [\$ 11,581.90 [\$ 12,581.90 [\$11,581.90 [\$11,581.90 [\$ 57,999.50 ]

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27 17	17	17	-17	v 950000	4-6.2	finguini arease sa nazagahn (Are Assar, B	57,077,00	99,037,00	49,000,00	49 102 10	49.000.00	95 8000
					*OFA, SI NOME (* HESOKRA) SI OTHER PERSONNEL		\$-,377,30	\$4,077,00	\$9.077.00	\$9.077.00	29.0% 0.	\$ 15 320 11.
0.07	17	17	17	5 1 1,100,00	THE STREET PERSONANDE	250 Student of Res Trillows	(8,1.81.20	37,000,00	9150000	57, 550 (6)	35.050 01	329470.07
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27 17	22	17	17	y NUMBER SC	100:	Annal Lonekant Jacks Paties	55,708,00	V5.70F.00	V. ACE OC	W. ZUNDO	45-22E BU	5/6/14/00
27 , 17	-17	12	12	£ 5705.00	'mp	A cold principally to day relega-	\$4,700,00	\$5,705.00	55 At 5 JK	25.718.00	25.7,60.	Sycamor,
2 0	12	15	15	5 5 265 Ye	Tul.	April Tolor ve it (t. 3c. tellsc)	55,265.30	55,765 Yo	55 Y.S.Y.	25,575,00	25.508.00	526 min ne
0 0	17	17	17	3 2000 200	700	WithorWorlde Asset 1s	33,277.2n	30,072.00	37 072 20	57 072 00	35072.01	345 2000
$\frac{9}{5} \cdot \frac{9}{9}$	17	17	17	5 1899-00 6 1899-00	- m:	Pur an adapted formal rate (Art. Artes, 1) (Altreation Frahmers in the Property)	57,676,00	\$7,60% DC	\$7,876,00	\$1800 PC (7.845.00)	\$788-0. (28-0.0)	\$19.420.00 \$19.450.00
5 - 6	17	17	17	5 71/8-20		abovery fermion 1-to Associat	V. I/E 20	37,170-70	37.00-00	37.030-100	37030-07	3 Pr 450 m
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25 12	17	12	12	5 1 5 P - 30		after of a set 100 s. As on Ta	\$7,575,00	\$7,50% (0)	\$7.800.00	(7.8-4.10)	(2 m-4, n)	\$ 19 420 0.
5 1 6	- 12	12	2	3 3 5m. Y.	Tul.	Rideby to the triple is the or	57,59 - 30	53,5m, Ye	55 mm. Y.	77 ×50.00	37 ×50, 07	5 Pri 4/9 mile*
					TOTAL OTHER PRISONEL		\$102,437.20	\$102,449.90	\$102,410.00	\$172,450.00	5172450.07	5552,470.07
					1014, 561407		9100 5 77 00	9101377-00	910154700	910156730	\$10196210	5 67 75000
					1 EGUPMENT							
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					D. TRAVEL		33,535.20	50.00	50.00	52.00	30.01	39.35.0
					1.0 ms/s		\$40,519.79	\$42,500.00	\$42,500.00	\$10,500.00	\$10,000.00	\$1-1646.0-
					2 chranen		37.20	30.00	30.00	32.00	30.07	30.07
					1014, 10491		\$47.573.39	100000	\$40,549.09	\$10,509,90	\$12579.51	\$157,040,95
					ELPART CIPANT/TRAINER SUPPORT COSTS							
					I Contract of Attachbuts a con-	'	97.30	51.70	51%	5216	27.00	30.00
					2 Augustal		37.20	30.00	30.00	52.00	50.01	30.07
					i flevel		5.00	19.00	10.00	1516	45.00	50 0.
					4 subsections:		\$, 30 37.20	\$0.00 30.00	\$0.00 30.00	\$3.00 32.00	200. 3200	300
					Total Parkinganos' i, men Angipan, costs		5.30	10.00	10.00	10100	500.	300.
					F. OTHER O RECT COSTS		1	172	- 22	1510	1011.	11.1.2
					1 Natrods a Pagnes			'		'		
					LatingConscionation in \$1.571.90,	1	31,001.0	30.00	30.00	52.00	50.00	\$5,075.05
					hit na Computer (2016) 51-789	- /	57,575,50	1900	10.00	50100	55.00	52,576.00
03. 17.0.	1.50	- 20	. 35	l	Superior		\$17,316,65	Sestant-20	\$51,3655	\$36,152,10	\$7\$ 1.16 × !	\$1-1-1-11 81
					Tay May per ISN SEC 45	•	\$17,579.76	51 W.	51%	5216	77.00	5.05787-
					Total Marwook and Anapters		5, 30	50.00	5000	53.00	2011.	300° 200,
					A modes for Gods A god offert Street		2.0	53.00	53.8	3316	2010. 2016	20 0. 70 0°
					4 COSC amples Stories		27.20	50.00	30.00	52.00	32.07	30.07
					Contraward occurrence un Nomen, du Com-	15	5, 30	1930	1000	5516	5000	50.0
					A copposition of a party stretch dates from		501	\$3.00	SUUC	\$3.00	200.	(0.0)
					3. Mile of a result for example	'	97.30	51%	51%	5216	72.00	30.00
					L. Other (PCR) Sociologic and Sequencings	10"	37,7107.70	501,000.00	50 0 000 00	507 (000 (0)	500,000,00	380 570 07
					Interest Const.		\$10 cm 12	\$91,095.70	Sec. 299.54	54) TO 10	\$18144.81	\$771.7089.
					4. CHRECT & MODIFIED CHRECT COSTS Talent Class		\$ 1002931	5 225,561.59	\$ 205,021.31	5 Int 22 in	5 197 776.55	\$10*10-385
					Midra di pressonato			£ 270,402.09				5 931 625 85
					4 (\$440000000000							
				IDC Rate in	le's		\$ 19,980.00	5 12,5 (+ 56)	\$ 1956.16	\$ 0.125.15	\$ 5,1000	\$ \$91-75\$
					Ford recent Costs		5 (5,001)6	5 10/10° 26	5 75 1,17 10	5 10 17745	5 30,272 67	5 35 11755
					"DIAL CIPELI AND INDIREL" CDS15							
					People Using		\$ 712,905 (0)	\$ 748,001.25	\$ 77970515	\$ 20,000.00	\$ 2.09-580	\$11,029.40
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					A TOTAL COSTS AND THIS Constitute		L 21 102 10	5.215,7113		L		
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				1 July 2020	1 July 2021	1 July 2022	1 July 2023	
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MONTHS			YEAR 1	YEAR 2	YEAR 3		OPTION YEAR 2	0%
Y1 Y2 Y3 OY1 OY2 BASE	A. SENIOR KEY PERSONNEL		TEAN	TEAN 2	TEARS	OF HOIS TERRET	OF HOW TEAK 2	5/11
1 1 1 1 1 5 -	Nambiar, Prejit	Laboratory Lead	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00
1 1 1 1 7 7	TOTAL SENIOR KEY PERSONNEL	Caporatory Ceau	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00
	B. OTHER PERSONNEL		30.00	30.00	30.00	20.00	,50.00	20.00
12 12 12 12 12 \$ 5,000	TBD	Research Assistant 01	55,000.00	\$5,000.00	\$5,000.00	\$5,000.00	\$5,000.00	\$25,000.00
12 12 12 12 3 3,000	TOTAL OTHER PERSONNEL	Research Assistant of	55,000.00	\$5,000.00	\$5,000.00	\$5,000.00	\$5,000.00	\$25,000.00
	TOTAL SALARY		\$5,000.00	\$5,000.00	\$5,000.00	\$5,000.00	\$5,000.00	\$25,000.00
	C. EQUIPMENT		\$3,000.00	\$3,000.00	33,000.00	33,000.00	\$5,000.00	\$23,000.00
	C. EQUIPMENT		\$0.00	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00
	TOTAL EQUIPMENT		50.00	50.00	50.00	\$0.00	\$0.00	\$0.00
	D. TRAVEL	1	30.00	30.00	30,00	\$0.00	20,00	50.00
			50.00	50.00	50.00	\$0.00	£0.00	\$0.00
	1. Domestic	-	\$0.00	S0.00 S0.00	SQ.00	\$0.00	\$0.00 \$0.00	\$0.00
	2. International		\$0.00		\$0.00	\$0.00	\$0.00	
	TOTAL TRAVEL		\$0.00	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00
	E. PARTICIPANT/TRAINEE SUPPORT COSTS		40.00	20 00	241.00	de an	An un	de au
	1. Turtion/Fees/Health Insurance		\$0.00	\$0.00	\$0,00	\$0.00	\$0.00	\$0.00
	2. Stipends		50.00	50.00	50.00	50.00	\$0.00	\$0.00
	3. Travel		50.00	50.00	50.00	\$0.00	\$0.00	\$0.00
	4. Subsistence		50.00	50.00	50.00	\$0.00	\$0.00	\$0.00
	5. Other		\$0.00	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00
	Total Participant/Trainee Support Costs		\$0.00	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00
	F. OTHER DIRECT COSTS							
	Materials and Supplies		52,950.02	52,950.02	\$2,950.02	52,950.02	\$2,950.02	\$14,750.10
	2. Publication Costs		50.00	50.00	50.00	\$0.00	\$0.00	\$0.00
	3. Consultant Services		50.00	50.00	50.00	\$0.00	\$0.00	\$0.00
	4. ADP/Computer Services		\$0.00	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00
	5. Subawards/Consortium/Contractual Cos	ts	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00
	6. Equipment or Facility Rental/User Fees		\$0.00	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00
	7. Alterations and Renovations		\$0.00	\$0.00	\$0.00	\$0.00	\$0.00	\$0.00
	8. 2-Day Annual Meeting Costs		52,000.00	52,000.00	\$2,000.00	52,000.00	\$2,000.00	\$10,000.00
	Total Other Direct Costs		\$4,950.02	\$4,950.02	\$4,950.02	54,950.02	\$4,950.02	\$24,750.10
	G. DIRECT & MODIFIED DIRECT COSTS							
	Direct Costs		\$ 9,950.02	\$ 9,950.02	\$ 9,950.02	\$ 9,950.02	\$ 9,950.02	\$ 49,750.10
	Modified Direct Costs		\$ 9,950.02	\$ 9,950.02	\$ 9,950.02	\$ 9,950.02	\$ 9,950.02	\$ 49,750.10
	H. INDIRECT COSTS							
IDC Rate>	10.00%		\$ 995.00	\$ 995.00	\$ 995.00	\$ 995.00	\$ 995.00	5 4,975.01
	Total Indirect Costs		\$ 995.00	\$ 995.00	\$ 995.00	\$ 995.00	\$ 995.00	5 4,975.01
	I. TOTAL DIRECT AND INDIRECT COSTS							
	Direct + Indirect		\$ 10,945.02	\$ 10,945.02	\$ 10,945.02	\$ 10,945.02	\$ 10,945.02	\$ 54,725.11
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# HDTRA1-14-24-FRCWMD-BAA

## CBEP = Thrust Area 6 - Cooperative Counter WMD Research with Global Partners

HDTRA1-14-24-FRCWMD-BAA

CTRP = Thrust Area 6 – Cooperative Counter WMD Research with Global Partners Biosurveillance for Spillover of Zoonotic Viruses in Rural Communities in India

Abstract: Zoonotic viruses including certain henipaviruses, filoviruses, and coronaviruses constitute a significant threat to global health. Nipah virus (NiV), Ebola virus (EBOV), and SARS coronavirus are highly pathogenic viruses and select agents capable of causing public health emergencies of international concern. In 2018, the WHO listed these pathogens among the top ten threats to global health and security. In 2014, a single spillover event of Ebola Zaire from an animal host (presumed to be a bat) caused an epidemic in West Africa that infected more than 28,000 people, killing approximately 40%. In Bangladesh and India, Nipah virus causes outbreaks of acute encephalitis in people and has an associated mean mortality rate of 70%. Saudi Arabia and the Gulf states continue to have new cases of MERS CoV, a zoonotic coronavirus related to SARS CoV. Bats are reservoirs for these and related viruses, and zoonotic transmission from bats to humans either directly or via domestic animals has previously occurred in Southeast Asia. Despite their importance as threats to public health and security, the full diversity of these viruses in bats and their potential to infect livestock and people remains poorly understood. This project will test the hypothesis that henipaviruses, filoviruses, and coronaviruses are circulating in bats in India and have spilled over into animal and human populations undetected, which presents a threat to human health. In line with Thrust Area 6 - Cooperative Weapons of Mass Destruction Research with Global Partners, this project will reduce threat from these select agents and viruses of high concern by enhancing early detection and surveillance capacity in India by: 1) conducting biological surveillance in bats, macaques, livestock and people in rural communities in Uttar Pradesh; 2) transferring PCR protocols and a multiplexed serology platform that can detect RNA from and IgG antibodies against all known henipaviruses, filoviruses, and SARS & MERS coronaviruses, to Indian partner labs; and 3) training laboratory personnel to develop and safely utilize these assays to identify human and animal infection and exposure to these viruses. This project will contribute to the Global Health Security Agenda, India and we will share results with key Indian government partners. The proposed project is closely aligned with the aims of the Cooperative Threat Reduction Program in that it will support threat reduction through biosurveillance and capability building, engagement with partnercountry scientists, and use of a One-Health approach.

Background: India has been identified as a hotspot for emerging infectious diseases (EIDs). The majority of EIDs are zoonotic and caused by pathogens that circulate in wild animal populations. People across South Asia, in both rural and urban settings, live in close association with domestic and wild animals, including bats, which have been identified as important reservoirs for zoonotic viruses. Zoonoses such as SARS CoV, Influenza and HIV, have already caused significant global pandemics, and there is potential for other zoonotic viruses (known and unknown) to cause widespread epidemics if given opportunities to jump into human populations (e.g. Ebola virus in West & Central Africa). The henipavirus, filovirus, and coronavirus viral groups include important zoonotic agents such as Nipah virus (NiV), Ebola virus (EBOV), and SARS CoV. Bats are natural reservoirs for all three viral groups. Nipah virus and Ebola virus are listed as select agents by CDC and USDA, and the WHO has identified them and SARS coronavirus as three of the ten most important global infectious disease threats. Enhancing surveillance for these significant zoonotic viruses in animal and human populations to improve detection and reporting in areas highly vulnerable to disease emergence is critical for reducing the risk of epidemics.

There is a need for enhanced surveillance for zoonotic viruses in India. Acute encephalitis syndrome (AES) accounts for more than 10,000 deaths each year in India. Uttar Pradesh (UP), in northern India, is a hot spot for AES, and more than 50% of encephalitides India-wide remain undiagnosed. Nipah virus is a

## HDTRA1-14-24-FRCWMD-BAA

## CBEP = Thrust Area 6 - Cooperative Counter WMD Research with Global Partners

zoonotic virus, carried by bats, that causes acute encephalitis in domestic animals and people. Since 2001, more than 20 NiV outbreaks have been reported in Bangladesh and India, with a mean mortality rate of 70% (100% in some outbreaks). Repeated spillover from bats; high mortality; and human-to-human transmission signifies NiV's importance as an emerging zoonotic pathogen with pandemic potential. Kerala, India reported its first outbreak of Nipah virus encephalitis in May 2018, which infected 23 people (case fatality rate = 91%). *Pteropus medius* bats are a reservoir for henipaviruses in Bangladesh and India. Our group has shown that Nipah virus incidence in pteropid bats in Bangladesh is low (~1%-4%). Viral infections are transient and periodic and spillover depends on coincident active infection within a local bat population and a route of transmission. In Bangladesh, spillover happens seasonally when people consume date palm sap that has been contaminated by bat excreta. The mechanism of spillover in Kerala remains unknown, and despite evidence of NiV in bats in northern and southern India, there is a lack of surveillance in people and livestock. As a result, the extent to which NiV contributes to AES in India is unknown.

Bats also carry filoviruses, which cause viral hemorrhagic fever. Several species of bats have been associated with Ebola and Marburg viruses in Africa. The 2014 Ebola outbreak in West Africa was hypothesized to have begun with a single spillover event from a bat, though which bat species remains unknown. In Asia, outbreaks of viral hemorrhagic fever have not been associated with filoviruses; however, antibodies against Reston ebolavirus, a non-pathogenic filovirus, were detected in pigs and people in the Philippines in 2008. Our group identified Reston ebolavirus RNA and antibodies in local *Mineopterus schreibersii*, a common insectivorous bat, suggesting that pig and human cases were due to spillover from bats. There is now more evidence that various bats in Asia carry filoviruses, though the diversity of those viruses and their threat to human health is unknown. Mengla virus is a new species of filovirus discovered in *Rousettus* bats in China. It is different from Ebola and Marburg, but shares structural and functional traits with them, suggesting that it can infect people. Prior to this discovery, we found antibodies against unknown Ebola-like viruses in *Rousettus leschenaultii* and *Pteropus medius* (the Nipah reservoir) in Bangladesh – both of which are also common in India, suggesting that related filoviruses also circulate in South Asia. Despite this, there is currently no surveillance for filoviruses in India.

SARS CoV emerged in live animal markets in southern China in 2002 and caused a severe respiratory and gastrointestinal disease in people. It spread to 26 countries and infected more than 8,000 people, with 9% mortality. Middle East respiratory syndrome (MERS) CoV, related to SARS CoV, was first identified in a person who died from severe respiratory disease in Saudi Arabia in 2012. Our group identified Chinese horseshoe bats as the natural reservoir for SARS and SARS-like CoVs. Horseshoe bats are also common in India and it is unknown whether SARS-like CoVs have contributed to human respiratory disease in India.

Scientific & threat reduction impact: The possible presence of bat reservoirs for zoonotic viruses and lack of surveillance presents a situation where high consequence viral pathogens may emerge in human populations undetected. This project will establish a surveillance platform that will help mitigate the threat of zoonotic viral outbreaks in animals and people in India. The proposed project closely aligns with the goals of the Cooperative Biological Threat Reduction Program. Improving coordination among public health and animal health agencies; increasing capacity to detect viral zoonotic agents in animals and people; and identifying high risk behaviors for spillover in communities where there is contact among these groups, is necessary to adequately respond to outbreaks and reduce the threat to people's health and well-being. This project will provide important information about the risk of spillover of high consequence zoonotic viruses to livestock and people where people and animals live in very close association. This project will implement biosurveillance using a One Health framework, and capability building through engagement with local and federal institutions. It will also support and complement

#### HDTRA1-14-24-FRCWMD-BAA

## CBEP = Thrust Area 6 – Cooperative Counter WMD Research with Global Partners

India's ability to meet IHR and OIE obligations and achieve Global Health Security Agenda (GHSA) objectives by enhancing local capacity to identify natural spillover or deliberate release of known and novel zoonotic agents. Early detection and prevention of spillover of high-risk viral pathogens in India, one of the most populous and globally connected regions of the world, will be critical to preventing local outbreaks and global pandemics.

Scientific approach & methods. We propose to test the following overarching hypothesis: Henipaviruses, filoviruses and coronaviruses are circulating in bats in rural India where people, bats and livestock live in close proximity, and spillover has occurred undetected. This study will answer the following questions: 1) Which bat species carry henipaviruses, filoviruses and high consequence coronaviruses in India; 2) are there seasonal patterns to viral infections in bat reservoirs in rural communities; 3) have other wildlife species (e.g. macaques) and livestock been exposed to these viruses; and 4) does having livestock increase risk of exposure? Building on our work from PREDICT in Uttar Pradesh, we will expand our human and animal surveillance where bat, human, and livestock populations live in close association. We will work with local university, state, and federal institutions using established and Indian Medical Research Council (ICMR)-approved sampling protocols developed under PREDICT. We have already established relationships with communities in Gorakhpur, a region with high rates of undiagnosed encephalitis and other illnesses. We will add additional communities in other parts of UP to total six field sites: three communities that rear livestock (e.g. pigs, goats, cattle) and three that do not. We will conduct quarterly sampling of wildlife (bats and macaques), livestock, and people at each location over 30 months (or 48 months including option years), and use questionnaires to collect data about how people interact with animals. We will collect ecological data such as bat species diversity and abundance, terrain and vegetation, food cultivation and dietary practices, and potential sources of environmental exposure to animals. Samples will be screened for henipavirus, filovirus, and SARS & MERS coronavirus viral RNA (PCR) or antibodies (Luminex) at King George's Medical University. PCR protocols for viral families developed under PREDICT will be established at KGMU. A validated, multiplexed serology assay (Luminex) that uses soluble glycoproteins from all known henipaviruses and filoviruses, as well as spike proteins from SARS and MERS CoV has been developed by the Broder lab at Uniformed Services University and we have used them in several previous and current research projects, including one funded by DTRA in Malaysia. We will also transfer these assays to other Indian lab partners (e.g. NIV, Pune) to enhance their ability detect novel and known members of these viral groups. We will compare viral detection rates and seroprevalence in humans and animals among study sites and measure changes over time. Serosurveillance will be especially informative to help identify bat reservoirs as these viruses cause acute, transient infections in bats making viral detection more difficult than antibody detection. Using a One Health approach to biosurveillance that includes people and livestock will also allow us to characterize virus and host diversity within these viral groups and identify past spillover events and high-risk behaviors that may have contributed to exposure.

We will report results from this project to relevant government stakeholders in human and animal health sectors, which will help them fulfill their reporting requirements for IHR and OIE. This project also supports the aims of the GHSA and we will present results at India GHSA meetings. This will also facilitate coordination with the US CDC, as they also conduct human disease surveillance in UP. We will publish findings from this study in high-impact, peer-reviewed journals and present at scientific meetings. We have included an optional 4<sup>th</sup> and 5<sup>th</sup> year in which we will continue targeted surveillance in UP, and also expand this project to Kerala state where a recent outbreak of Nipah virus occurred and where our group has already established partnerships with state wildlife, livestock and human health agencies through the state One Health platform.

# HDTRA1-14-24-FRCWMD-BAA CBEP = Thrust Area 6 – Cooperative Counter WMD Research with Global Partners

Capacity Building. This project will build capacity by: 1) developing and training local field teams; 2) transferring diagnostic technologies to partner (including GoI) laboratories and training staff to develop reagents & analyze data; 3) facilitating exchange of research personnel between US partners (EHA, USU) and partner labs in India; 4) providing postdoctoral research opportunities for Indian students interested in One Health, epidemiology, and laboratory diagnostic development; 5) holding One Health meetings to bring together stakeholders from human, domestic animal, and environmental (wildlife) sectors to strengthen cooperation and communication around viral disease surveillance; and 6) supporting the development of the new ICMR National One Health Institute (an MOU between EHA and ICMR is in process). These activities align with CTRP's mission to enhance India's ability to detect and respond to biological threats posed by zoonotic viruses. The increased capacity at both the field and laboratory level in human and animal health sectors, including wildlife, will directly improve livelihoods, reduce the threat from deliberately released agents, and limit the opportunity for zoonotic viruses to emerge undetected.

Partner Institutions and roles: EcoHealth Alliance (EHA): Design and coordinate the study with project partners; supervise local post-doctoral candidate; ensure coordination between field and lab studies and communication to Indian government agencies; and collaboratively analyze and publish data. EHA will be responsible for all contractual obligations with DTRA. King George's Medical University (KGMU), Lucknow: Local implementing partner. KGMU will coordinate & conduct field surveillance activities, sample testing, data analysis and reporting. National Institute of Virology, Pune: The national reference laboratory for new and emerging viruses, as well as BSL 4 agents. NIV Pune will assist with further genetic characterization of viruses. Uniformed Services University, Maryland. Study development. Development and transfer of Luminex reagents to partner labs. Training, data analysis, and reporting.

5-year staged plan: Year One: Formalize local partnerships & update protocols; apply for research approvals through ICMR/HMSC; transfer PCR protocols, Luminex and reagents to partner labs and conduct trainings; hire & train local animal field team; develop database for project; community outreach and begin human & animal sampling. Years 2-3: Continue sampling; coordinate stakeholder meetings to share results and develop surveillance and intervention strategies; publish/present findings (conference & journal). Coordinate with Govt. of India, regional One Health networks (e.g. Bat One Health Research Network) and US CDC India office to disseminate information from project and help develop regional strategies for risk mitigation of bat borne zoonoses. Optional Years 4 & 5: Continue longitudinal surveillance in animals and people in UP; initiate activities in Kerala with established state One Health partners (e.g. Kerala Forest Dept., Kerala Veterinary and Animal Sciences University).

Major Goals and Milestones: 1) Identify bat reservoirs for henipaviruses, coronaviruses and filoviruses: Establish study protocols; train field teams; implement field studies; transfer diagnostic technologies to lab partners; 2) Test the hypothesis that spillover has occurred previously in Uttar Pradesh: Establish community surveillance; integrate questionnaire about animal exposure; screen human and animal samples. 3) Strengthen One Health systems by improving knowledge and capacity: Support the development of a One Health platform in India via ICMR's new One Health Institute. Establish inter-departmental collaborations; conduct integrated training of local technical personnel & graduate students. Report findings to local & federal authorities publish and share project findings via GHSA India meetings & scientific conferences.

Estimated Budget: Y1: \$850,000 Y2: \$950,000 Y3: \$950,000 Y1-3 Total: \$2,750,000

OY 4 \$850,000 OY 5 \$750,000. **OY4-5 total: \$1,600,000** 

Cooperative Biological Engagement Program (CBEP) = Thrust Area 6 – Cooperative Counter WMD Research with Global Partners

## Biosurveillance for Spillover of Henipaviruses and Filoviruses in Rural India

Abstract: The henipaviruses and filoviruses include Hendra virus (HeV) and Nipah virus (NiV), and several species of Ebola (EBOV) and Marburg virus (MARV), respectively, which are highly pathogenic viruses and select agents capable of causing public health emergencies of international concern. In 2018 the WHO listed Nipah virus as one of the top ten threats to global health and security. In Africa, Ebola viruses have caused significant morbidity and mortality in people and wildlife. In 2014, Ebola Zaire caused an epidemic in West Africa that infected more than 28,000 people, killing approximately 40%. The outbreak began with a single spillover event, presumably from contact with an infected bat. Pteropus medius is a reservoir for Nipah virus and occurs throughout India and Bangladesh. In Bangladesh and India, including a recent outbreak in May 2018 in Kerala, India, Nipah virus has caused recurrent, seasonal outbreaks of acute encephalitis in people and has an associated mean mortality rate of 70%. Hemorrhagic fever has also recently been reported in India, but surveillance is limited and to date, no Ebola-related cases have been identified. Bats are reservoirs for both henipa- and filoviruses, and zoonotic transmission of these viruses from bats to humans either directly or via domestic animals has previously occurred in Southeast Asia. Despite their importance as threats to public health and security, the full range of henipa- and filoviruses in bats and their potential to infect livestock and people remains poorly understood. The recent outbreak in Kerala illustrates that Nipah virus circulates in bats throughout the country and spillover is possible wherever bats and people interact. In India, reducing spillover risk is particularly important due to the high population density in the region and high global connectivity via international airports, creating opportunity for local outbreaks to become pandemics. In line with Thrust Area 6 - Cooperative Weapons of Mass Destruction Research with Global Partners, this project will reduce threat from these select agents through enhancing early detection and surveillance capacity in India by: 1) transferring Luminex technology with validated reagents to detect IgG antibodies against all henipaviruses and filoviruses to key partner labs in animal and human health sectors; 2) training laboratory personnel to develop and utilize Luminex assays to identify exposure to henipa-and filoviruses; 3) conduct biological surveillance in bats, nonhuman primates, livestock and people in rural communities in Kerala where there are high levels of contact among people and animals, and with optional expansion into high risk zones in Uttar Pradesh and West Bengal. This project will test the hypothesis that both henipaviruses and filoviruses are circulating in fruit bats in India and have spilled over into animal and human populations, which presents a threat to human health. Activities will be coordinated with the Global Health Security Agenda, and surveillance data will be shared with the government partners at local, provincial and national levels. The proposed project is closely aligned with the aims of the Cooperative Biological Engagement Program in that it will support threat reduction through biosurveillance and capability building, engagement with partner-country scientists, will engage with the CBEP Bat One Health threat reduction network (BOHRN) and promotes a One-Health approach.

Point of Contact & Principal Investigator: Dr. Jonathan Epstein. Epstein@ecohealthalliance.org

Cc: Ms. Emma Lane. lane@ecohealthalliance.org

Cooperative Biological Engagement Program (CBEP) = Thrust Area 6 – Cooperative Counter WMD Research with Global Partners

### Biosurveillance for Spillover of Henipaviruses, Filoviruses, and Coronaviruses in Rural India

Abstract: The henipaviruses and filoviruses include Hendra virus (HeV) and Nipah virus (NiV), and several species of Ebola (EBOV) and Marburg virus (MARV), respectively, which are highly pathogenic viruses and select agents capable of causing public health emergencies of international concern. Severe acute respiratory syndrome (SARS) and Middle East respiratory syndrome (MERS) are zoonotic coronaviruses associated with global spread. In 2018, the WHO listed these pathogens among the top ten threats to global health and security. In Africa, Ebola viruses have caused significant morbidity and mortality in people and wildlife, and Saudi Arabia and the Gulf states continue to have new cases of MERS CoV, some of which are associated with dromedary camels. In 2014, a single spillover event of Ebola Zaire from an animal host (presumed to be a bat) caused an epidemic in West Africa that infected more than 28,000 people, killing approximately 40%. Currently, there is an Ebola outbreak in an insecure region of DRC which threatens to spread to Uganda. In Bangladesh and India, Nipah virus has caused recurrent, seasonal outbreaks of acute encephalitis in people and has an associated mean mortality rate of 70%, including the recent outbreak in May 2018 in Kerala, India. Hemorrhagic fever and acute encephalitis both occur in India, but surveillance is limited and most cases remain undiagnosed. Bats are reservoirs for henipa- corona-, and filoviruses and zoonotic transmission of these viruses from bats to humans either directly or via domestic animals has previously occurred in Southeast Asia. Despite their importance as threats to public health and security, the full diversity of these viruses in bats and their potential to infect livestock and people remains poorly understood. The recent Nipah virus outbreak in southern India and previous findings of Nipah in bats in northern India illustrates that Nipah virus circulates in bats throughout the country and spillover is possible where bats and people interact. In line with Thrust Area 6 - Cooperative Weapons of Mass Destruction Research with Global Partners, this project will reduce threat from these select agents and viruses of high concern by enhancing early detection and surveillance capacity in India by: 1) transferring Luminex technology with validated reagents to detect IgG antibodies against all henipaviruses, filoviruses, and MERS and SARS CoVs to key Kerala state and government of India partner labs in animal and human health sectors; 2) training laboratory personnel to develop and utilize Luminex assays to identify human and animal exposure to henipa-filo- and CoVs; 3) conduct biological surveillance in bats, nonhuman primates, livestock and people in rural communities in Kerala and UP, where there are high levels of contact among people and animals. This project will test the hypothesis that henipaviruses filoviruses, and coronaviruses are circulating in bats in southern India and have spilled over into animal and human populations undetected, which presents a threat to human health. Activities will be coordinated with the Global Health Security Agenda and surveillance data will be shared with the government partners in Kerala and the Government of India (GoI). The proposed project is closely aligned with the aims of the Cooperative Biological Engagement Program in that it will support threat reduction through biosurveillance and capacity building, engagement with and implementation by partner-country scientists, and use of a One-Health approach.

Point of Contact & Principal Investigator: Dr. Jonathan Epstein. Epstein@ecohealthalliance.org

Cc: Ms. Emma Lane. lane@ecohealthalliance.org

						DO	MAINS					
	Domai	n 1	Domain 2	2		Domain	3			Domain	4	
							ETENCIES					
Training Task	1.1.1	1.1.2	2.1.1	2.1.2	2.1.3	3.1.1	3.1.2	3.1.3	3.1.4	4.1.1	4.1.2	4.1.3
Ex. Task 1.3.2: Conduct laboratory- based diagnostic trainings 1.2.2. hire and	x	x	x	X	x	×	X	×			×	x
train field and lab teams												
1.2.5-OY5.2.5. Train students & team through field & lab activities.	х	x	х	X	x	x	x	x			x	х
1.3.3-2.3.3. Train technicians from KGMU to develop reagents & run pseudovirus neutralization assay.	х	x	X	X	x	x	x	x				
1.6.1-OY4.6.1 Hospital staff identified and trained on study protocols	х	x	х			x						

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	Domair	า 5				Domain 6	<b>)</b>		Domain i	7		Domain	8	
				•		COMPETENCIES								
Training Task	5.1.1	5.1.2	5.1.3	5.1.4	5.1.5	6.1.1	6.1.2	6.1.3	7.1.1	7.1.2	7.1.3	8.1.1	8.1.2	8.1.3
Ex. Task 1.3.2:					Х									
Conduct														
laboratory-														
based														
diagnostic														
trainings														
1.2.2. hire and	х	x	х	х	х	x	х	x	x	x	x	х	х	х
train field and														
lab teams														
1.2.5-OY5.2.5.	х	x	х	х	х	x	х	x	x	x	x	х		
Train students														
& team														
through field &														
lab activities.														
1.3.3-2.3.3.	х	x	X	Х	х				x	×	X	X	X	х
Train														
technicians														
from KGMU to														
develop														
reagents & run														
pseudovirus														
neutralization														
assay.														

1.6.1-OY4.6.1					х	x	x	х	х	х
Hospital staff identified and										
identified and										
trained on										
study protocols										

Instructions: this matrix will help track and capture the alignment of BAA research training and professional development activities to BTRP's Domains, Competencies, and Proficiencies Framework. Please note: alignment is only required to the competency level definitions shown below.

- 1. in the left-hand column of the above chart, list every training activity to be performed on a separate line, using their task or sub-task number (ex. Task 2; subtask 2.1; subtask 2.1.1)
- 2. place an x or tally against the competencies that align to the training activity. Competency decisions can be made by determining using the definitions listed below and determining if a) the training subject matter corresponds to the defined competency or b) if the training outcome corresponds to the defined competency. Please note: training activities can align to multiple competencies across the 8 domains. (Note: if completing an annual report, stop at step 2; if drafting a Full Proposal, continue to step 3)
- 3. once completed, list the aligned competencies under its corresponding training activities in the grant proposal.

Domain	Domain Definition	Competency	Competency Definition
1. Disease recognition and prevention of spread	1.1 Understanding of emerging and high-consequence pathogens (EHCP) in the following realms: morbidity, mortality, transmission,	1.1.1 Disease characteristics and impact	Disease characteristics and epidemiology of especially high consequence pathogens (EHCPs) and the diseases they cause, including etiology, signs and symptoms, morbidity, mortality, risk factors, incubation period, period of communicability, mode of

	control, disease presentation and distribution, and impact.	1.1.2 Clinical Infection control and animal health biosecurity	transmission, clinical presentation, mechanisms of spillover, reservoir species, spatiotemporal and geographic distributions of affected populations, and demographic characteristics of affected populations (species/age/sex/occupational status/ etc.), including outcomes on population health.  Understanding of basic infection prevention and control procedures related to especially high consequence pathogens (EHCPs) for human and animal clinical settings in line with GHSA and IHR milestones (when applicable), including standard precautions (PPE use; disinfection; sterilization); environmental infection control; laboratory testing; nosocomial infections, including AMR; and animal health biosecurity principles, procedures.)
2. Disease identification, detection, confirmation, and reporting	2.1 Positive identification of emerging and high consequence pathogens (EHCP) and reporting to appropriate authorities.	2.1.2 Test procedures (presumptive/confirmatory)	Knowledge, understanding and proficiency with procedures for safely and securely collecting appropriate specimens/samples for testing of pathogens (human; animal; environmental), especially EHCPs, and handling and packaging of samples for transport to diagnostic testing laboratories, based on national and international guidelines and best practices, while ensuring specimens will are maintained in condition required for testing (cold chain and or fixation, etc.)  Knowledge, understanding and proficiency with procedures for conducting screening, diagnostic, and confirmatory testing and interpreting test results for ECHPs from

			different types of samples using different types of tests (ELISA, PCR, etc.) for high consequence pathogens, including test accuracy, reliability, specificity, and sensitivity and quality control/quality assurance (QA/QC) measures needed to validate testing results.
		2.1.3 Reporting	Knowledge, understanding and proficiency in policies and procedures for disease notification and/or bi-directional reporting of surveillance and laboratory data through national information-sharing systems, or to notify regional/international authorities such as WHO/OIE/FAO of a disease, other significant epidemiological event, or potential public health emergency of international concern (as appropriate to roles and responsibilities at each operational level).
<b>3.</b> Analytics, assessment, and research	<b>3.1</b> Appropriate study design, data analysis, and implementation of processes, as applies to each discipline.	<b>3.1.1</b> Study design and ethics	Knowledge and understanding of scientific methodologies and study design principles (hypothesis development, research plan development, sampling design, strategy for data collection and analysis) and application of research ethics to responsibly address safety, security, and ethical issues related to the research aims and hypotheses, including understanding of issues around dual-use research of concern.
		3.1.2 Methodologies and analysis	Knowledge, understanding and proficiency in the use of statistical and other relevant methodologies for scientific research analyses including practices and approaches for data collection, use and interpretation of statistical data analyses, data visualization, and use of

		statistical data software tools.
	<b>3.1.3</b> Dissemination of findings and outreach to stakeholders	Knowledge, understanding and proficiency in the dissemination of research findings through peer-reviewed scientific journal publications, conference presentations and public outreach activities, including engagement with appropriate stakeholders and partners at national, regional, and international levels.
	3.1.4 Resource mobilization and management	Knowledge, understanding and proficiency in the planning, acquisition, mobilization, allocation, management and tracking of financial and non-financial resources for research projects, including proposal development and grant writing.
4.1 Critical skills, knowledge and behaviors necessary to effectively manage projects and programs, lead teams, champion solutions, inspire action, and sustain programs.	<b>4.1.1</b> Plan, implements, and evaluates a project or program	Knowledge, understanding and proficiency in program planning, implementation, monitoring and evaluation, including development and measurement of appropriate indicators, and advocacy, or management/oversight of the process.
	<b>4.1.2</b> Apply program management techniques throughout the project or program	Knowledge, understanding and proficiency in program management, including timeline and deliverables management, team building, and generating high-level stakeholder support.
	and behaviors necessary to effectively manage projects and programs, lead teams, champion solutions, inspire action, and sustain	3.1.4 Resource mobilization and management  4.1 Critical skills, knowledge and behaviors necessary to effectively manage projects and programs, lead teams, champion solutions, inspire action, and sustain programs.  4.1.2 Apply program management techniques throughout the project or

		<b>4.1.3</b> Lead culturally diverse teams to complete a project or program	Knowledge, understanding and proficiency in developing and leading culturally diverse teams, ensuring collaboration among team members, and mentorship of future team leaders.
5. Quality management systems	5.1 Knowledge, skills, and abilities required for developing a culture of quality. Operations, services, and infrastructure integrated in a system that meets applicable standards, professional guidelines, and customer requirements for ensuring and maintaining quality and continually improving services.	5.1.1 Physical environment and equipment  5.1.2 Qualified, well-trained, and competent workforce	Knowledge, understanding and proficiency with facility and equipment operations and maintenance, and facility management policies and procedures, in support of establishing a culture regarding physical environment and equipment necessary for EHCP detection, management, and control, or management/oversight of these operations, in a system that meets applicable standards, professional guidelines, and customer requirements for ensuring and maintaining quality and continually improving services.  Knowledge, understanding and proficiency with the principles, processes, and methodologies for workforce development and sustainment in relation to EHCP detection, management, and control, including workforce planning, training, refresher training, continuing education, and/or professional development, in a system that meets applicable standards, professional guidelines, and national licensing requirements where applicable.
		5.1.3 Materiel resource management	Knowledge, understanding and proficiency with procedures and policies for procurement

			and inventory management of materiel resources, including supply chain systems, operations, and infrastructure, or management/oversight of these processes.
		5.1.4 Quality review practices and procedures	Knowledge, understanding and proficiency with quality assurance, quality control, and/or quality management review practices and procedures, based on national or international best practices.
		5.1.5 Document management	Knowledge, understanding and proficiency with document and information management policies and procedures, and program documentation and records/archival management processes.
<b>6.</b> Outbreak preparedness and surveillance	<b>6.1</b> Use of operational guidelines, identification and analysis of outbreak situations, and effective communication with stakeholders.	6.1.1 Operational guidelines	Knowledge, understanding and proficiency with development and implementation of outbreak preparedness, surveillance, investigation, laboratory practices, and response planning and operations guidelines, including case definitions, and demonstrated ability to use related operational tools such as emergency preparedness and response plans, contingency plans, SOPs, and job aids when available. Special note and care should be

			given to documents designed for health care workers directly combatting an outbreak.
		6.1.2 Identification and analysis	Knowledge and proficiency with principles and processes for developing and maintaining disease outbreak situational awareness capabilities, including coordination and communication or reporting across surveillance, laboratory, and other data sources, epidemic intelligence, data analysis, and field epidemiological and outbreak investigation techniques, such as contact tracing, and decision-making in support of these activities.
		<b>6.1.3</b> Communication systems	Knowledge and proficiency with development, implementation, and sustainment of communications systems for outbreak situational awareness and response, including multisectoral and intrasectoral coordination (including via emergency operations centers or equivalent assets), operational risk communication, and public risk communication.
7. Biosafety	7.1 The application of knowledge, techniques, and equipment to prevent accidental exposure to and release of emerging and high consequence pathogens (EHCP).	<b>7.1.1</b> Biohazard Identification and classification	Knowledge, understanding and proficiency with identification and classification of biohazards associated with EHCPs, for laboratories, hospitals, other biomedical and clinical facilities, and the community, including biohazard risk assessment and characterization, and biohazard identification policies and compliance procedures.
		7.1.2 Biohazard Control Measures	Knowledge, understanding and proficiency

		(e.g., personal protective equipment, safety practices and equipment, facility design, and administrative controls).	with development and implementation of international best practices and procedures for EHCP biosafety and biohazard control measures (substitution, elimination, mitigation) for personnel, equipment, facilities (laboratories, hospitals, and other biomedical and clinical facilities) and the community.
		7.1.3 Biohazard Incident Management	Knowledge, understanding and proficiency with development and implementation of policies, procedures, and operational guidelines, including emergency operation communications systems, for Biohazard Incident Management for EHCP outbreaks, releases or exposures.
8. Laboratory biosecurity	8.1 Prevent theft, misuse, loss, or deliberate release of emerging and high consequence pathogens (EHCP).	8.1.1 Biosecurity Risk Identification	Knowledge, understanding and proficiency with principles and procedures for Biosecurity Risk identification, risk assessments, characterization and compliance with recommended international standards and best practices to prevent the loss, theft, or misuse of biological organisms and laboratory/research technologies.
		8.1.2 Biosecurity Risk Mitigation	Knowledge, understanding and proficiency with international best practices for Biosecurity Risk Mitigation, and related risk reduction approaches to enhance physical security, personnel reliability, material control and accountability, transport security, and information security/cybersecurity, and the development and implementation of Biosecurity Risk Mitigation strategies for laboratories, facilities, and personnel to

8.1.3 Biosecurity Incident Management	prevent the loss, theft, or misuse of biological organisms, especially EHCPs, and laboratory/research technologies.  Knowledge, understanding and proficiency with international best practices for Biosecurity Incident Management, and development and implementation of Biosecurity incident management policies and procedures for suspected or confirmed illicit
	access to pathogens, intruder alerts, and other biosecurity incidents.

Please fill out the requested information on all tabs (click on each bottom colored tab or sheet) labeled Cover Page, Personnel, Publications, Presentations, Impacts, Intellectual Property, and Transitions. For annual reports, please only describe <u>DTRA funded work</u> for the reporting period (July 1, 2019 - June 30, 2020).

Tab	Description
Cover Page	
	Providing information about the project participants and collaborating organizations allows an assessment of performance in promoting partnerships and collaborations.
<u>Publications</u>	The products from an effort demonstrate the excellence of the research and efficacy with which the results are being communicated to colleagues, potential users, and the public. Publications are the characteristic product of research. If there is nothing to report during this reporting period, the awardee shall state "Nothing to Report".
Presentations	The products from an effort demonstrate the excellence of the research and efficacy with which the results are being communicated to colleagues, potential users, and the public. Many projects (thought not all) may develop products other than publications. In all cases, if there is nothing to report during this reporting period, the awardee shall state "Nothing to Report".
Impacts	knowledge, enlarges the pool of people trained to develop that knowledge and techniques or put it to use, and improves the physical, institutional, and information resources that enable those people to get their training and perform their functions. In all cases, if there is nothing significant to report during this reporting period, the awardee shall state "Nothing to Report".
	Information on any intellectual property demonstrates the value of protectable innovative ideas resulting from sponsored research and includes patents, inventions, and licenses. Submission of this information is not a substitute for other invention reporting requirements under the awards terms and conditions. In all cases, if there is nothing
Intellectual Property	significant to report during this reporting period, the awardee shall state "Nothing to Report".  Refers to cases where knowledge resulting from your DTRA sponsored effort will be further developed for, or will be
<u>Transitions</u>	used in, a technology application. Transition sponsors can be entities in DoD, other federal agencies, or industry.

SHADED FIELDS ARE REQUIRED

Report Submitted to Defense Threat Reduction Agency

Grant/Award # HDTRA1-20-1-0026
Organization/Institution EcoHealth Alliance

Project Title Biosurveillance for Spillover of Henipaviruses and Filoviruses in Rural India

Name of Submitting Official Dr. Jonathan Epstein
Title of Submitting Official Principal Investigator

Email Address epstein@ecohealthalliance.org

**Phone Number** 1.212.380.4471

Submission Date 7/1/2021

Reporting Period End Date June 30th, 2021

Reporting Period Annual

Foreign Spending Country Foreign Spending (\$)

55891.3

Country 1 India

Country 2
Country 3

# **Proceed to Personnel**

Enter the principle investigator (PI) and all other personnel that have worked at least one person month ("160 hours) on this DTRA project during the reporting period, regardless of the source of compensation.

regardless of the s	regardless of the source of compensation.								
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HDTRA1-20-1-0026	Co-PI	Epsteir	Jonathan			EcoHealth Alliance	44		
HDTRA1:20-1-0026	Co-PI	Jain	Amita		2	King George Medical Unviersity	43		
HDTRA1:20-1-0026	Research Associate/Scientist	Nambiar	Prent		1	Kerala Vetermanan and Aericultural Sciences University			
HDTRA1-20-1-0026	Research Associate/Scientist		Ava		7	EcoHealth Alliance			
						Department of Forests and			
HDTRA1-20-1-0026	Research Associate/Scientist	Zacariah	Arun		1	Wildlife, Government of Kerala			
HDTRA1-20-1-0026	Research Associate/Scientist	Sterling	Spencer		7	Uniformed Services University			
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**Proceed to Publications** 

Enter all publications (Journal Articles, Books, Book Chapters, Conference Papers, and Theses/Dissertations) resulting from this DTRA project during this reporting period. DO NOT include publications that have not yet been submitted.

	p																		
Instructions	Select the apprecriate publication type from the drop down list. So not "nothing to report". I there were no cubications.	Enterthe IV I name of		Enter the EUL in name of the article title (mapter title if book)				Letter tea first page number of the article or chapter	Selectific audication status from the drop down list (w.g. Published Submitted)	ident Fer type	Anner the publication dentifier our bor for type selected in the anexious column.	an acknowledgment	Shirici whether or not a journal acticle or contention due may gone through the pecuniary acticles.						
Award #	Publication Type	Author	Journal [Book Title)	Article Title (Chapter Title)	Volume	Issue	Publication Year	Page	Status	Publication Identifier Type	Publication Identifier	Acknowledgement of DTRA Support		Editors [Book]	Collection [Back]	Publisher (Book)	Publication Location (book)	Media Type (Book)	Edition (Book)
Cearryle	reamai Article	John Leith and iand Doc	Journal of Applied Prysics	Trace chemical analysis using Caherent Anti-Stakes Roman Epectrascopy	1	1	2019	112	Somished	00)	10.1621/jn+23456x	Yes	Yes						
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**Proceed to Presentations** 

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Select the Enter the location of the

(conference) as City, State for domestic

drap down list the acknowledgement of DTRA personnel type — support was included in the for the presenter. presentation. conference, professional meetings or Lity, wereindiges. society, or university). Country for international meetings.

Select from the - Select whether or not an

	presentations.				international meetings.			
Award #	Presentation Type	Presentation (Paper) Title	Conference Date	Conference Name	Conference Location City, State	Author(s)	Personnel Type	Acknowledgement of DTRA Support
		Infrarea Spectroscopy of		Materials Research				
Ехатріғ	Orai Keynate	Metal Carponyls	8/1/2019	Society Fall Meeting	Boston, MA	John Smith	Groauate Student - y	es
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# **Proceed to Impacts**

Enter awards, honors, promotions, or graduations that occurred during the reporting period for the personnel contributing to this DTRA project.

Award #	Impact Type	Impact Description	Personnel Type	Awardee Name	Impact Date	Impact Details/Website
		Best Graduate Student Poster at the				
		American Physical Society winter				
Example	Best Poster	conference	Graduate Student	John Smith	9/20/2019	
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**Proceed to Intellectual Property** 

Enter all patents, invention disclosures, and licenses resulting from this DTRA award that were awarded or applied for during the reporting period. Reporting here does NOT replace the reporting requirements for patents and invention disclosures in the terms and conditions of the award.

Instructions	Select the intellectual property type from the drop down list life intent. Identify, an invention). Select Nathing to Report? If there will no intellectual property.	the intellectual property.	Enter the relevant number for the patient, application, etc.	the nation:	obbicanon oace in	name(s) of the	Select the status of the patention application from the drop down list.
	was no intellectual property.						

	Was that List Service	an britishin indi								
Award #	Intellectual Type	Property	Title	Number	Country	Application Date	Date Issued	Abstract or Description	Inventors	Status
			Accedential Device to					Inscalan regard salamited to State University for an accelerator to delive theority shielded	folia Sando and Cane	
Example	Javesférn		Behat WMDs	723456	1/5A		R/1/2019	MMOs	Anybody	Submitted
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# **Proceed to Transitions**

# Enter transitions of the research or technology from this DTRA project.

# Instructions

Enter a text description of the transitioned Enter the year research or technology. Enter "nothing to" that the transition government agency to the which this report" if there were no transitions.

started.

Enter the organization, company, or work has (is) transitioning.

	report in there were no transitions.	Juiteur	work has (is) cransitioning.
Award #	Transition Description	Year	Transitioned to
	The accelerator technology for detection of WMDs transitioned to funding from other US government agency for further		
Example	development	2020	USGOV Contract # 12-34-5678
HDTRA1-20-1-0026	Nothing to report		
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Biosurveillance for Spillover of *Henipaviruses* and *Filoviruses* in Rural India, Dr. Jonathan Epstein EcoHealth Alliance HDTRA1-20-1-0026

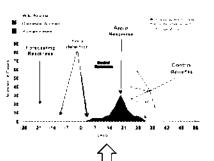
Objective: To enhance capacity for surveillance for high priority zoonoses in animals and people in Uttar Pradesh and Kerala. To identify wildlife hosts for henipaviruses and filoviruses and characterize their distribution near at-risk human populations, and to conduct targeted biosurveillance in wildlife, livestock and humans to detect exposure to or infection with high consequence viruses. Threat Reduction Impact: 1) enhancing field and laboratory capacity to safely conduct biosurveillance in animals and people for henipaviruses and filoviruses; and 2) conducting targeted biological surveillance in wildlife (esp. bats), livestock and people in rural communities in Uttar Pradesh and Kerala to determine reservoir hosts for and detect spillover of these viruses; and 3) assess the role of livestock and geography in exposure rates of henipa- and filoviruses in rural communities. Capacity/Capability Building: Through the development of lab capacity, joint multi-sectoral field surveillance activities, and graduate student training, India's capacity to detect and respond to emerging zoonoses and identify additional causes of AES and VHF will be significantly strengthened

Status of effort: Current collaboration with public health and vet partners in UP & Kerala; laboratory team hired, Luminex and lab supplies procurement underway, local permissions submitted, archived human samples from study sites currently available; 25 reagents for filo and henipavirus antibody detection produced and validated.Personnel Supported: 4 Researchers, 4 University Faculty, 2 Field Veterinarians, 2 Graduate Students, 1 Post-Doctoral Fellow, 2 Scientific Advisors; 6 Lab Research Assistants/Techs; 6 Field Scientists, 1 Program Assistant. Publications & Meetings:

Stakeholders Meeting planned July 2021

PANDEMIC EMERGENCE

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Surveillance in animals
and people at high risk to
detect & respond to
spillover earlier

Enhance capacity within India to conduct surveillance for henipa- and filoviruses Years 1-5Produce & validate reagents to detect henipa- and filovirus antibodies Years 5.Conduct wildlife, livestock, and human biosurveillance rural communities Years 1-5.Carry out hospital-based a encephalitis and viral hemorrhagic fever studies in clinic settings Years 1-5Funding Profile: Y1 \$970,566.84 Y \$995,322.63 Y3 \$999,666.56 OY4 \$998,965.88 OY5 \$998,985.95 TOTAL Y1-Y5 \$4,963,517.86 Contact information: PI: Jonathan H. Epstein 21238-4467 Epstein@ecohealthalliance.orgCo-PI: Amita Jain

#### DTRA Basic Research Annual Report

Please answer all sections of the document. You are welcome to use figures and tables to complement or enhance the text. For annual reports, please only describe work for the period of performance (July 1, 2020 - June 30, 2021). (Project awarded September 24, 2020)

Grant/Award #: HDTRA1-20-1-0026 PI Name: Dr. Jonathan Epstein

Organization/Institution: EcoHealth Alliance

Project Title: Biosurveillance for Spillover of Henipaviruses and Filoviruses in Rural India

### What are the major goals of the project?

List the major goals of the project as stated in the approved application or as approved by the agency. If the application lists milestones/target dates for important activities or phases of the project, identify these dates and show actual completion dates or the percentage of completion. Generally, the goals will not change from one reporting period to the next. However, if the awarding agency approved changes to the goals during the reporting period, list the revised goals and objectives. Also, explain any significant changes in approach or methods from the agency approved application or plan.

Our overall goal is to characterize the distribution of henipaviruses and filoviruses in bats and determine whether undetected spillover into livestock or people has occurred in rural communities in Uttar Pradesh and Kerala, India. By enhancing capacity at partner institutions to conduct targeted surveillance for filoviruses and henipaviruses in wildlife, domestic animals, and high-risk rural human populations, using a One Health approach, we will reduce threat of future outbreaks. The proposed interdisciplinary research effort has three objectives: 1) enhance capacity for surveillance for henipaviruses and filoviruses in animals and people in UP and Kerala; 2) identify wildlife hosts for henipaviruses and filoviruses and characterize prevalence within these hosts; and 3) conduct targeted biosurveillance in wildlife, livestock, and humans to determine exposure rates and detect spillover of these high consequence viruses.

What was accomplished under these goals? For this reporting period describe: 1) major activities; 2) specific objectives; 3) significant results, including major findings, developments, or conclusions (both positive and negative); and 4) key outcomes or other achievements. Include a discussion of stated goals not met. As the project progresses, the emphasis in reporting in this section should shift from reporting activities to reporting accomplishments.

# Task 1.

- 1) Major activities: Apply for permits and ethical approvals.
- **2) Specific objectives:** To gain approval across One Health research and project areas, including human sampling permissions, animal sampling permissions, livestock sampling permissions, ethical approvals, for compliant work to occur across both geographical states (UP and Kerala) through both local and US-based approving bodies. This includes HMSC research permits/ethical approval via the Indian Medical Research Council, local IACUC permissions, wildlife permits will be obtained via the Forest Department, US IRB and US-based IACUC through Tufts Vet School.
- 1.1.1. Apply for HMSC research approvals & local ethical approvals;
- 1.1.2. Apply for IRB, IACUC, HRPO, ACURO approval in US;
- 1.1.3-2.1.3. Apply for wildlife permits in UP and Kerala.
- 3) Significant results: All local permissions and approvals have been generated and submitted for local clearance. Significant efforts related to this objective include the submission of the human-sampling ethics clearance through King George Medical University (KGMU), Kerala State and UP State Forest Department permissions, livestock sampling permissions through Indian Council of Agricultural

Research (ICAR), and Institutional Biosafety Committee clearance through Kerala Veterinarian and Agricultural Sciences University (KVASU), all of which have been submitted and are pending approval. US-based IACUC and IRB have been generated and submission is imminent.

4) Key outcomes: All major local permissions have been submitted, and permission is expected upon committee hearings. Individual meetings, including those with the State Surveillance Officer in Uttar Pradesh, were held to discuss and prime stakeholders on project activities in order to facilitate permissions and approvals. Once all local and US-based permissions are gained the full project activities will be able to commence. Permission and approvals further represent a key opportunity and outlet for stakeholder sensitization and have provided the project with a platform to introduce the objectives of the research, build local relationships with stakeholders, and build interest and buy-in from community and institutional gate-keepers.

# Task 2.

1) Major activities: Enhance capacity within India to conduct biosurveillance for henipaviruses and filoviruses

### 2) Specific objectives:

- 1.2.1. Transfer BioRad Bio-Plex 200 machine & reagents and PCR protocols to KGMU;
- 1.2.2. Hire and train field and lab teams, including a 7-day Luminex training
- 1.2.3 Select 1 PhD student at KGMU and USU and hire a postdoc at EHA
- 1.2.4-2.2.4 Create sample repository with tracking system at KGMU.

### 3) Significant results

The transfer of a Luminex machine has been initiated with Luminex US for shipment of KGMU in Lucknow, UP, India. The shipment of supplies including reagents is underway between partners from Uniform Services University (USU) to King George Medical University, for use with Luminex machine. A Luminex training has been planned to occur once the machine arrives at KGMU. The lab team at KGMU has been hired, including a doctoral research associate, research assistants, laboratory technicians, and a data entry operator. Laboratory SOPs have been shared, and training sessions are being planned to commence once supplies and equipment are received.

4) Key outcomes: 7 hospital collaborators have been identified and 6 laboratory and hospital staff have been hired for this study. The purchase and shipment of a Luminex machine, as well as a shipment of necessary laboratory supplies, has been initiated. Implementing serological surveillance for groups of viruses known to cause encephalitis and hemorrhagic disease will greatly enhance the capacity of local health systems to detect exposure to or past infection in wildlife, domestic animals and human populations. To establish this capacity, EHA and KGMU will establish a joint human and animal surveillance team and train members to safely sample animals and people, using appropriate personal protective equipment and other biosafety techniques. We will provide a BioRad Bio-Plex 200 machine with computer console to KGMU and coordinate training activities for serological surveillance and panhenipavirus and pan-filovirus consensus PCR assays for molecular analysis using existing PCR capabilities at KGMU's virology lab. The hiring of the research team allows for targeted and novel capacity building efforts in at KGMU, a highly skilled and respected diagnostic laboratory, in disease detection methods and One Health approaches to surveillance.

#### Task 3.

- 1) Major activities: Produce & validate reagents to detect henipa- and filovirus antibodies:
- 2) Specific objectives:
- 1.3.1.-OY5.3.1 Provide reagents to partner labs;
- 1.3.2-OY5.3.2 Produce reagents for newly discovered viruses:
- 1.3,3-2.3.3 Train technicians from KGMU to develop reagents & run pseudovirus neutralization assay.
- 3) Significant results: Protein production and assay validation is part of the ongoing collaboration between EcoHealth Alliance, the Uniformed Services University, and the King George Medical University in Lucknow, India. Throughout Year 1, our team has produced and validated reagents

recombinant henipavirus attachment glycoprotein (G) for antigen-based serology. These G protein antigens will be used for the planned retrospective human sera screening of banked samples stored at KGMU. Reagents are prepared to ship and are planned to be sent as choreographed with the arrival of the Luminex machine.

**4) Key outcomes:** These assays will be used to screen bat samples and hospitalized patient sputum and CSF for viral RNA. Testing bat samples and acutely ill hospital patients by PCR may allow us to detect viral RNA during active infection, providing important information about the diversity of viral strains circulating within bat species that may account for human infection and observed immune responses seen on Luminex.

#### Task 4.

- 1) Major activities: Develop/maintain project database
- 2) Specific objectives: Develop a functional database that allows for the secure storage, sharing, and analysis of field and lab data and metadata, that will also connect to sample tracking software so that data is linked to sample IDs in repository. Database will be adaptable to new types of data that may arise during project.
- 3) Significant results: We have created a functional project database which is bespoke to the current project's data needs which are complex and interdisciplinary in nature. The project database uses an Airtable platform which is relational in structure and links data records across tables. Our project database in Airtable allows for collaborative data entry efforts across sampling and lab efforts, as well as secure remote storage, tiered access, and the capacity to hold multitudinous data points. The online database is able to contain all information related to field work and field sites, bat, livestock and human samples, and results of molecular assays. We have also created a SOP for the database which includes clear guidance on use, data quality/data assurance measure, and a data dictionary.
- **4) Key outcomes**: An online database tailored to the specific project needs has been created. This database allows for the secure, organized and comprehensive storage of data across multiple sampling and lab efforts. The database allows for report generation, data cleaning measures, and analysis tools. The database works across data collection platforms and allows multiple data types to be stored together in a secure, central system.

### Task 5.

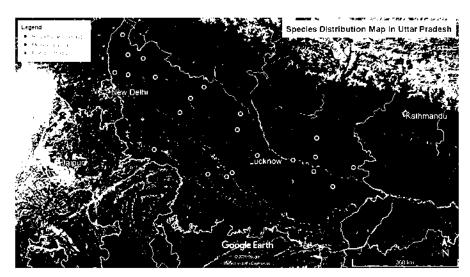
- 1) Major Activities: Screen archived human VHF and AES samples from KGMU
- 2) Specific Objectives:
- 1.5.1-2.5.1. Archived specimens identified/logged/tested
- 1.5.2.-3.5.2 Results entered into database.
- **3)** Significant results: The partnership between EHA and KGMU has been formed through a signed sub awardee agreement, representing the ability for KGMU to use their robust biobank to screen archived samples. While the project awaits the impending shipment of supplies and the Luminex equipment, the KGMU laboratory team has been formed and primed to receive training on the Luminex platform.
- 4) Major Outcomes: The KGMU subaward has been established, and 6 hospital and laboratory staff have been hired, in anticipation of the screening of archived samples. King George's Medical University hospital is a referral hospital that serves patients from all over Uttar Pradesh. Annually, KGMU sees more than 300 BHF cases and 1000 AES cases, many of which are negative to known agents such as Dengue or JEV. Professor Jain has led studies of AES and VHF, and by screening archived serum and CSF from patients who have previously volunteered and consented to use of their samples for this aim, we will be able to assess the baseline prevalence and geographic distribution of henipaviruses and filoviruses in this population of acutely ill patients, which will inform our sampling strategy for the village cross sectional studies as well as deepen our understanding of whether these agents play a significant role in causing severe VHF or AES in UP. We will screen 600 archived sera from VHF patients and from 900 AES patients using the Bioplex assay. Up to 300 CSF samples will be screened using PCR.

### Task 6.

- 1) Major Activities: Hospital-based AES and VHF studies
- 2) Specific Objectives: 1.6.1-OY4.6.1 Hospital staff identified and trained on study protocols
- 3) Significant results: The KGMU hospital staff team has been identified, including collaborators in the KGMU Department of Pediatrics, Medicine, Neurology, Microbiology, as well as research and laboratory associates and assistants. The team awaits the materials and supplies required for a full training on study protocols and methodologies. The human clinical survey has been created as a standardized qualitative and quantitative data collection tool.
- **4) Major Outcomes:** 7 hospital collaborators have been identified and 6 laboratory and hospital staff have been hired for this study. The hiring and training of the hospital-based team will allow for samples to be taken from a clinical setting in an ethical, efficient, and scientifically sound manner. Data from this study will generate new hypotheses about the prevalence of NiV and EBOV related viruses in India, as well as support the human behavioral risk underpinning these diseases of interest.

#### Task 7.

- 1) Major Activities: Wildlife, livestock, and human biosurveillance in rural communities
- 2) Specific Objectives:
- 1.7.1.-OY4.7.1. study sites selected
- 1.7.2-OY5.7.2 human, wildlife, and domestic animal samples collected and tested
- 3) Significant results: While the Indian subcontinent has been experiencing the extremely disruptive effects of the COVID-19 pandemic, the project has thus far been able to address the matter of site selection using a literature review, expert knowledge, and species mapping. Experts in bat biology have been utilized to map bat species distribution in UP and Kerala, and expect use these tools to inform further efforts in site selection when travel is safe. Areas in UP and Kerala where key bat species are found will be considered in conjunction with overlapping areas of livestock farming, as well as human settlements.



**4) Major Outcomes:** Bat distribution mapping efforts have been undertaken by members of the Animal Health team across UP and Kerala. A literature review of relevant bat species has been performed as a foundational source of information relevant to site selection, which will inform the mapping exercise and in-person field visits and reconnaissance once is reasonably safe. A standardized data collection survey tool has been developed to be used in human community sampling efforts.

Task 8.

- 1) Major Activities: Analyze data
- 2) Specific Objectives: Analyze field epi and lab results.
- 3) Significant results: None to report.
- **4) Major Outcomes:** While no major data collection efforts have been undertaken at the point in the project, the project database has been created with this major activity and specific objective in mind. The project database has been designed in order to facilitate rapid analysis and generate, present and publish epidemiological and other project data.

Task 9.

- 1) Major Activities: Disseminate results to relevant stakeholders
- 2) Specific Objectives:
- 1.9.1-5.9.1 Submit annual reports to DTRA.
- 1.9.2-OY5.9.2 Provide results to local stakeholders.
- 3) Significant results: None to report.
- **4) Major Outcomes:** Annual report generated for DTRA, July 2021. A local stakeholders meeting is planned to occur July 2021 with leadership from Co-Pl's Dr. Amita Jain of KGMU, Dr. Prejit Nambiar of KVASU, Dr. Arun Zacariah of Kerala Wildlife Department, and Dr. Rajesh Bhatia, formerly of WHO SEARO. This planned stakeholder's meeting represents a major point of relationship building and stakeholder engagement for the project, and is expected to inform project ability to gain rapid approvals, disseminate project results, and further implement the project successfully.

What samples has the project collected and what methodologies has the project implemented? For this reporting period, use the impact questions below to guide your discussion of: 1) sample collection; 2) sample storage; 3) laboratory techniques; and 4) sustainment of skills. You should provide an annual sample repository list to include all samples collected from this research effort. Discuss how you plan to maintain samples collected during the proposed research effort, along with relevant metadata, for at least 12 months after the project end date. If nearing the end of the project, you should state a plan and schedule for final disposition of EDP samples collected in support of the research effort.

### Samples

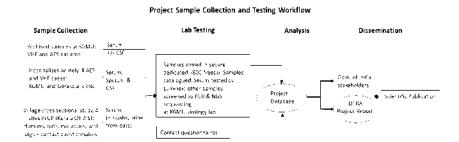
- What are the roles and responsibilities of each partner in sample collection for pathogen identification?
- How will the sample collection, field or clinical, methodologies practiced during this research project have long-term impact on sustained and durable capabilities?

## Laboratory or Clinical Techniques

- What are the laboratory techniques built or reinforced during the duration of the research project?
- How will the techniques and practices taught through the duration of the research project support long-term sustainment and durability of skills in partner country(ies)?

At this point in the project, approvals, permissions, and permits are underway to allow for sample collection to occur. Due to the hardship imposed by the ongoing COVID-19 pandemic in India, this already involved process has an elongated timeline. As such, no samples have been collected for the project thus far. Despite this, progress has been made in terms of the creation of methodologies for future use.

All biological samples will be collected in duplicate and stored in our partner lab at KGMU in secure ultra-cold freezers. Samples will be tracked using specialized lab software and linked to a database with all metadata collected at the time of sampling. Samples will be tracked using specialized lab software and linked to our project database with all metadata collected at the time of sampling.



We will provide protocols and training for sample collection, handling, and storage in the field and lab, using established biosafety and biosecurity protocols, including those developed under PREDICT. We will provide a BioRad Bio-Plex 200 machine with computer console to KGMU and coordinate training activities for serological surveillance and pan-henipavirus and pan-filovirus consensus PCR assays for molecular analysis using existing PCR capabilities at KGMU's virology lab. Co-PI Broder and Co-I Laing (USU) will provide Bioplex / ELISA reagents for all henipa- and filovirus assays to KGMU. Epstein (EHA) will provide the Bioplex machine and PCR protocols. Drs. Broder and Dr. Laing will run a 7-day training course for KGMU lab technicians and invite technicians from UP state medical and veterinary diagnostic labs which currently have ELISA capabilities. To manage and reduce biothreat, a secure, dedicated -80C freezer will be installed at KGMU for project samples. Commercial software will be used to log and curate samples in the lab, creating a sample repository at KGMU.

## What opportunities for training and professional development has the project provided?

If the research is not intended to provide training and professional development opportunities or there is nothing significant to report during this reporting period, state "Nothing to Report."

Use the impact questions below to guide your discussion of opportunities for training and professional development provided to anyone who worked on the project or anyone who was involved in the activities supported by the project. Please indicate for each training: participants, objectives, duration, subject matter, outcomes, and assessment results. For trainings aligned to BTRP Training Domains, Competencies, and Proficiencies Framework (DCPs), please indicate the competencies associated with the training according to the matrix. "Training" activities are those in which individuals with advanced professional skills and experience assist others in attaining greater proficiency. Training activities may include, for example, courses or one-on-one work with a mentor. "Professional development" activities result in increased knowledge or skill in one's area of expertise and may include workshops, conferences, seminars, study groups, and individual study. Include participation in conferences, workshops, and seminars not listed under major activities.

#### Training

- How will the trainings completed through this project contribute to long-term durability of the capabilities this project reinforces?
- What are the capabilities built, reinforced, and/or improved upon in the partner country through the efforts of this research project?
- What are the areas of challenge in terms of transitioning all research capabilities to partner countries and sustaining these efforts?

### Conference Attendance/Knowledge Sharing (Network Building)

- How did you leverage conferences, seminars, and stakeholder meetings to support and engage partner country scientists affiliated with the research?

- How has this research leveraged local and regional threat reduction networks to strategically address current or future biological threats?
- How has this research strengthened the capabilities of partner country scientists to identify and secure competitive research funding through networking and scientific collaboration activities?

Nothing to Report

#### How have the results been disseminated to communities of interest?

If there is nothing significant to report during this reporting period, state "Nothing to Report."

Use the impact questions below to guide your discussion of how the results have been disseminated to communities of interest. Include any outreach activities that have been undertaken to reach members of communities who are not usually aware of these research activities, for the purpose of enhancing public understanding and increasing interest in learning or careers in science, technology, and the humanities.

### Transparency

- How has this research endorsed a culture of transparency within, and between local, regional, and international stakeholders associated with this effort?
- How has this research supported public awareness and prevention strategies in the local and regional communities of partner country(ies)?

### Intellectual Property/Publications

- How have the protocols and information learned during the research project informed disease detection and surveillance knowledge (protocols, SOPs, training) in the long-term?
- How will the information garnered from the research affect standard guidance on disease detection for clinical health professionals? This impact can be in the form of guidance updates, clinical guidelines, or supporting evidence for policy changes at the country-level.

# Evidence Based Data Communication to Stakeholders

- How has this research enhanced or supported mechanisms (SOPs, committees, outreach, etc.) to share resources, expertise, and data between local, regional, and international stakeholders?
- How has this research supported a communication plan to national and international public health stakeholders in the event of a public health emergency of international concern?

Nothing to report

### What do you plan to do during the next reporting period to accomplish the goals?

If there are no changes to the agency-approved application or plan for this effort, state "No Change." Describe briefly, what you plan to do during the next reporting period to accomplish the goals and objectives.

At the time of this report, there are no major changes to the agency-approved plan for this effort, and we expect that all tasks and subtasks will be achieved despite the increased difficulties as a result of the COVID-19 pandemic in India. However, an adjustment has been made in order to best achieve the goals of the project. In the approved proposal, the earliest surveillance efforts for human hospital sampling, human community sampling, wildlife sampling, and livestock sampling were slated to occur in firstly Uttar Pradesh, while our efforts in Kerala were staggered to begin later in Y2, and increase in subsequent years. During the period in which we are seeking permissions, approvals and permits, we have been seeking permissions to work in both states. Based on leadership and identified capacity in Kerala for performing wildlife training, especially through our partners at Kerala Wildlife Department, and KVASU,

we have decided to assemble and train the animal team in Kerala. In doing so, we have identified pragmatic reasons for developing plans to work in both states at the earliest point of approval. This will allow us to commence field work as soon as permissions are granted fully. This shift will give us increased flexibility should there be unavoidable delays in one state or another, based on the pandemic, natural/climate conditions, political conditions, or otherwise. It also allows the project to best utilize the capacity and leadership of the emerging Animal Health team in Kerala, working in concert with team in Uttar Pradesh.

From:	Aleksei Chmura
To:	(b)(6)
Cc;	(b)(6) Whitney Bagge; Joe Riccardi; Billy Karesh Re: [Non-DoD Source] FRBA14-6-2-0471 Clarifications
Subject: Date:	Thursday, September 17, 2020 11:38:05 AM
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Aleksei Chmura,	PhD
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	IvalCovs	-	Call Rev 184		111,251,00	1	450 100 27 - 5	481291-3 5	490,009,01	ŝ	2,299,291,50
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	CAFFA.1 Totals		951 354, 54	Ľ		Ś		99 (97) (32 - 5		5	4859485

tHS Supplies	Jms/4ma. I	Y1	Y2	.,	27.1	0.0	
Laptina.	\$ 7,535.0	\$ 3,575.0					\$ 5,000.0
SF Ca	\$	s	S	\$	S	5	5
trupum	\$ 70, .0	\$ 100	\$ 4,00.	\$ 40,00	S AUTOU	\$ 40,00	\$ 7,000.00
A Life	5 397.50	\$ \$97.50	5 197.97	5 243.50	\$ 197.10	5 247.50	5 1,767.50
Le leuce	9 749.99	5 749.99					5 749.95
MS Clinic	5 0000	\$ 200,00	\$ 5000	3 205 30	\$ 20063	5 205 30	\$ 1,0000
Communications	3 347.70	5 447.70	S. Minn's	3 197.00	5 90000	3 140 20	5,7,5000
Zoten.	3 357.70	5 150,000	5 (50.07)	3 851.00	5 2000	3 150.20	4 2000
Francisco	) 25.70	5 25.70	5 2507	3 25.20	5 2500	3 25.20	3 125.5
IOTALS		\$ 8,240,45	\$ 1,533.50	\$ 2,512,50	\$ 2,512.50	\$ 2,112,50	\$17,499,45

Constitution Services	*1	"		"		Ç4		,	• >
PALEK	\$ 2,50,00	s	\$0.00	s	$\lambda_i  m  n_i$	\$	90, 30	s	Sames
C.L. Winner/S	9.4/06/00								
TOTAL	9 9,500 CO	3	500 CO	- 5	500.00	9	500.00	- 5	500.00

pana necestr	"	- 17	- "			
PAGN Cine			1675	3145		
Secretary Reports			1970	2870		
PRAC			IOIP	15-44		
let-1			52.5	5769	e	,

		Internation	al 1	ravel								TOTAL
Traveler 1 William Karesh (Ammac)					П				Г		Г	
Hitte Mai Per Diem	5	1.958 00	15	1,968,00	5	L958 PO	١:	1,968,00	5	1.958.00		
Meals and the dental Expenses Per Diem	5	1.242 00	S	.,242.00	5	1.242 00	S	242.00	5	1,242 00		
Taxi Estimate	5	570.00	S	570.00	5	570 00	S	570.00	5	570 00	Г	
Flight Estimate	5	2,566	Ş	2, 566	S	2,506	\$	2,360	5		Г	
No Days		5	•	4	l	5	•	4		5	1	
No Tops (⊣r Year	Т	2	_	2	Г	2	_	2	Г	2	Т	
7otos	5	6.346 00	S	6,346.30	5	6.346 00	S	6.346.30	5	0,780 00	S	29,164,00
Fravolor & William Karesh (Field)	П				П				П		Т	
Hose, May Per Diem	5	584.00	٠,	584.00	١,	584.00	٠,	584 30	5	584.00	1	
Meals and the Jental Expenses Per Diern	5	472 50	15	472.50	5	472 90	1	472.50	5	472 53	Т	
Taxi Estimate	5	285 00	S	285.00	5	285 00	S	285,00	5	185 00	Т	
Flight Estimate					Г						Т	
No Days	Т	5	_	4	Г	5		4	Т	5	Т	
No Tops (⇔r Year			•	ı	l		•	ı		1	1	
7otor	5	1.341.50	Ś	.,3450	5	1.341.50	S	3450	5	1,341.50	s	6,707,50
Intivelor si Cathorine Mache daa (Amman)	Т				Т				Т		Т	
Hotel Max Per Diem	5	1,968.00	'5	1,966,00	5	1,968.00	.5	1,968.00	5	1,968.00	$\vdash$	
Meals and Incidental Expenses Per Diern	ls.	1,242.00	4	1,242,00	ls.	L342 M	4	1,242.30	ls.	1,242.03	1	
Taxi Estimate	5	570.00	Š	570.00	5	570 00	S	570,00	5	570 00	т	
Flight Listimate	5	2,566	5	باباد را	s	2,566	5	2,560	5	4,506	т	
No Days	Т	5	_	4	Г	5	-	4	Т	5	Т	
No Topaj ⊕i Year		2	•	2	l	2	•	2		2	1	
7otor	5	6.346 00	Ś	6,346,30	5	6.346 00	Ś	6.346.30	5	6,340 00	s	31,730,00
Intive or 4 - Cathorine Mache coal Pield)					Т						Т	
Hotel Max Per Diem	5	1,414.00	'5	2,528.00	5	7,628.00	'5	2,528.30	5	7,628.00	Т	
Meals and Incidental Expenses Per Diern	ls.	227.50	٠,	1,999,00	ls.	L205 m	٠,	2,395.00	ls.	1,225.00	l	
Taxi Estimate	5	455.00	15	870 GC	5	870 M		870 30	5	870.00	т	
Flgnt Edmate											т	
No Days	Т	10		10	Т	10		10	Т	10	Т	
No Tues ser Year		1	•	7	l		•	7		1	1	
7oto	5	2,745.50	15	5 493 30	5	5,493.00	1	5,493,00	5	5,493.03	5	24,718.90
Traveler 5 - Choly Hagan (Amman)	Т				т				Т		т	
Hote MiskPer Diem	5	1,958,00	S	1,968,00	s	1,968,00	5	1,968,30	5	1,968,03	т	
Meals and Incidental Expenses Per Diem	5	1,247.00	'5	1.742.00	5	1,747.00	Ś	1,747.30	5	1,242.03	Т	
Taxi Estimate	5	973.00	÷	570 30	5	970.00	÷	570 30	5	970.00	T	
Flent Edimate	5	6.4.5	s	0,415	5	6.4.5	s	0,415	5	6.4.5	Т	
No Days	Ė	5	_	5	Ė	2	_	5	Ė	5	T	
No Tups per Year	Т	7		7	Т	7		7	T	7	T	
Zotai	k	13 195 00	٠.	10 199 00	l٠	12:05:00	٠.	10,195,00	k	10,105.00	1.	50,975 M

have en 6 - International Conferences (IM60)		Vienna									1	
Hotel Max Per Diem	5	8.9	S	819	5	1.038	S	1,608	5	1.036	Г	
Meals one the dente. Expenses Per Diem	Ś	240	Ş	240	Ś	480	Ş	480	Ś	483	Г	
Taxi Estimate	5	255.00	5	755.00	5	\$10.00	5	510.00	5	510.00	Г	
Flight Estimate	5	1.495	3	1,493	5	2,986	5	2.986	5	2.986	1	
No Dava		đ		2	Г	đ		2	Г	đ		
No. Trips per Year		:		1		1		2		1		
Torai	5	2,807.00	. 5	2,807,00	5	5,614.00	5	5,504.00	5	5,614.00	5	27,455.00
Visa Eces	١,	504.00	3	504.00	5	504.00	'5	504.00	5	504.00	5	7,520.00
Vaccinet on Costa	5	853 M	_		Г		_		Г		?	850.00
Conference Repisitation Fees	5	500 00	S	500.00	5	1,000,00	S	200.20	5	1,000 00	S	4.000.00
YEARLY TO TAILS	S	51,616.00	Ş	32,532,50	S	sb,8s9.50	Ş	36,839,50	S	64,273.50	\$	173,101.00
conference			•		vi	sa .					-	
§ DyiMED	Г	500			70	i Ge II	_	55	1			

Domestic Travel	П		П			Г				Г	TOTAL
Traveler 1 - William Karosh (NY DC)	П		П			Г				П	
Hotel Max Ren Den	5	912	5	255 \$	515	5	512	٠¸	912	l	
Meals and Incidental Expenses Fer Dein	S	178	ś	114 9	228	ś	228	9	128		
Tax Estorate	S	90.00	ś	45.00 9	3000	ś	90.00	9	90.00		
Transportation Estimate	\$	392	5	156 \$	292	5	35 2	.\$	392	П	
No. Days		7		2	?		2	:_	?		
No Trips per Year		2		1	7		2		2		
Teto!	S	1,722.00	ś	0.1.00 9	1,222.00	ś	1.222.00	9	.,722.00	ś	5.499.00
Traveler 2 - Catherine Mitchalaba (NY DC)	П		П			Г				П	
Horel Max Por Dem	5	756	5	756 5	256	5	756	5	756		
Meals and Incidental Expenses Fer Den	1	214	5	114 5	214	5	114	5	214	Г	
Tax Estorate	S	25 ()(	ś	45.00 9	a£ 00	ś	45.00	9	45.00	Г	
Transportation Estimate	Ş	196	5	156 \$	196	\$	155	ş	196		
No. Eays	Г	7	Г	2	?	Г	2		?	Г	
No Trips per Year	ı	l	l	: '	1		1	•	1	ı	
Feto!	S	611.00	ś	0.1.00 9	61.00	ś	0.1.00	9	61.00	ś	0.055.00
Traveler 3. Attending Meeting in VA											
Horel Max Por Dietro	5	1,024	5	1,324 5	1,024	5	1,024	5	1,074	П	
Meals and Inordental Expenses Fer Den-	5	330	5	380 \$	330	5	380	Š	380	ı	
Tax Estorate	S	450 00	ś	450.00 9	45000	ś	450.00	9	450.00	Г	
Transportation Estimate	Ş	1,011	5	1.311 \$	1,011	S	1,311	ç	1,011		
No. Days	П	3	П	9	÷	Г	4	_	ń	П	
No Trips per Year		2		ε.	2		2	•	2		
Fetot	S	2,864 80	ś	2,864,80 9	2,864.80	ś	2,864,80	9	2,804.60	ś	. 4.324.00
Traveler 4 Presenting Domestic Mtg ASTA	ин (	Maryland)	Г			Г				П	
Horel Max Por Detro	5		5	124 5	924	5	104	5	974	Г	
Meals and Inquiental Expenses Fer Den-	5		5	497 S	497	5	49.7	Īs	497	ı	
Tan Estimate	5		5	570 to 8	310.00	5	573.00	5	210.00	П	
Transportation Estimate	S		ś	684 9	684	ś	684	9	684		
No. Days	Γ	3	Γ	4	4	Γ	4		4	Γ	
People Attending		3	ı	2 .	?		2	:	?	ı	
Fo;al	5		5	2,315.00 5	2,515,00	5	2,305.00	5	2,515,00	5	9,263.00

Conference Registration fees	ś		ś	\$10.00	9	93000	ś	\$10.00	9	930.00	
YEARLY TOTALS	\$	4,697,60	\$	7,321.20	ç	7,942.60	\$	7,542.20	ş	7,942 80	\$ 22,158.00
asmh	5	455 GC	Г								

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						JUST	¥1	Υ2	V3	OY1	OY2	Tutal
Α.	Y2	Y3	0¥1	OY2	Base	A. Senior/Key Personnel						
12	12	12	12	2 9	\$ 127,940.00	Dr. Ehab Abu-Basha, CoPI	\$ 127,940 CO	\$ 127,940.00	\$ 127,940.00	\$ 127,940.00	\$ 127,940,00	\$ 639,700.00
						Dr. Ehab Abu-Basha Fringe	S -	3 -	S -	3 -	S -	s -
- 8	В	8	н	8 9	\$ 19,301,80	Hani Talafha, Biologist	\$ 12,867.87	\$ 12,867,87	\$ 12,867.87	\$ 12.867.87	\$ 12 867 87	\$ 64 339.33
	<del>l                                     </del>		Ť	<u> </u>		Hani Talafha Fringe	\$ .	\$ -	\$ .	4	t .	4 .
2	12	12	12	-2 -5	\$ 32.520.00	Zaidoun Hijazeen, Veterinarian	\$ 5,420.00		*	\$ 32.520.00	\$ 32.520.00	\$ 135,500,00
			- '-		4 02.020.00	Zaidoun Hijazeen Fringe	\$ ·	\$ -	\$ .	\$ -	\$ -	<b>s</b> .
3	3	3	3	3 5	\$ 5439153	Mustafa Ahahneh Virologist/Lah Director	\$ 13,597.88		\$ 13,597.88	\$ 13,597.88	\$ 13 597 88	\$ 67.989.41
		~	,		a ., 301-23	Mustafa Ababneh Emige	\$ .	3 3.207 (112	\$ .	g 5,587 (II)	£ 12,391,00	g () ((((((((((((((((((((((((((((((((((
2.5	2.5	2.5	2.5	2.5	\$ 67.903.1B	Zuhair Bani Ismail, Livestock ID Expert	\$ 12,063.16	-	\$ 12,063.16	\$ 12.063.16	\$ 12.063.16	\$ 60.316.81
. 2.3	2.0		2.3	•	¢ 01.800. u	Zuhair Bani Ismail Fringe	\$ -	s -	⊕ 12,50a.1a	¢ 2.000.10	2.000.10	¢ 00.010.01
- 3	3	3	3	3 5	\$ 69 597 00	Borhan Al-zoghoul Molecular Biologist/Lab Director	T	\$ 17,399.25	\$ 17,399.25	\$ 17,399.25	\$ 17 399 25	\$ 86,996,25
	,	-	,		a calaron	Borhan Al-zughoul Fringe	\$ -	8 -	\$ 17,398.23	a 1.25522	\$ -	a 120 0 0 0 0 0 0 0 0 0
- 3	3	3	3	3 '5	\$ 23,112,00	Bilal Al-Omari, Lab Supervisor	\$ 5,778.00	-	\$ 5,778.00	\$ 5.778.00	\$ 5.778.00	\$ 28.890.00
		3		-	<b>⊉</b> 23.112.00	Bilal Al-Oman Fringe	\$ 3,778.00	\$ 5.776.00	\$ -	\$ 0.770.00	\$ -	\$ 20.090.00
	1			1 5	\$ 177.018.63	Wail Hayajneh Human ID Expert	\$ 14,751.55			\$ 4,751.55	\$ 14.751.55	\$ 73.757.76
	1		1	1 .3	\$ 177 U18.63							
						Wail Hayajneh Ennge	•	*	0	3 -	٠	3 -
			F	-ringo Rato>		Total SemonKey Personnel	\$ 209,817.71	\$ 236,917,71	\$ 236,917.71	\$ 236,917,71	\$ 236 917 71	\$ 1,157.488.57
		,				B. Other Personnel	l		l			
1	1		2	2 .5	\$ 58 241.66	Saad Charaibeh, Avian Pathologist	\$ 4,853.47	\$ 4,853.47	\$ 4,853.47	\$ 9,706.94	\$ 9.706.94	\$ 33,974.30
						Saad Gharaibeh Fringe	\$ ·	3 -	S -	\$ -	S -	\$ -
3.5	3.5	3.5	3.5	3.5	\$ 28 219,16	Admin	\$ 8,230.59		\$ 8,230.59	\$ 8,230,59	\$ 8 230 59	\$ 41 152.94
						Admin Fringe	\$ -		\$ -	\$ -	\$ -	\$ -
2	4	4	3		\$ 47,860.32	Nurses (x2)			+ ,			\$ 103.697.36
12	12	12	12		\$ 2,538.84	Undergrad Student Stipends (x2)		\$ 5,077.68		\$ 5,077.68		
12	12	12	12	12	\$ 338508	Graduate Student Stipends (x2)	\$ 6,770.16	\$ 6,770.16	\$ 6,770 16	\$ 6,770.16	\$ 6770 16	\$ 33,850,80
			H	-ringo Rato>		Total Other Personnel	\$ 40,885 34	\$ 56,838,78	\$ 56,838.78	\$ 53,715,53	\$ 29 785 37	\$ 204 089.50
						C. Equipment						
						Vehicle	\$ 37,150.00	\$ 1,830.00	\$ 1,830.00	\$ 1,830.00	\$ 830.00	\$ 44,470.00
						Luminescence plate reader	\$ 28,846.00					
						Total Equipment	\$ 65,996.00	\$ 1.830.00	\$ 1,830.00	\$ 1.830.00	\$ 1830.00	\$ 73,316,00
						D. Travel			·			
						1 Domestic Travel	\$ 40,108.00	\$ 62,626.00	\$ 62,626.00	\$ 51,367.00	\$ 12,360,00	\$ 229 087.00
						2 Foreign Travel	\$ 37,433.50	\$ 26,890.00			\$ 29,919,50	\$ 148,019.00
						Total Travel	\$ 77,541.50	\$ 89,516 (II)				
						L. Participant/Trainee Support Costs						
						1. Tuilior/Fees/Health Insurance						<b>s</b> -
						2 Stipends		1			1	<b>s</b> .
						3 Travel						3 -
						4 Subsistence						s .
						5. Other						\$ -
						Total Participant/Trainee Support Costs	£ .	s -	š -	s -		s
						F Other Direct Costs		* -		Φ -		
						Materials and Supplies	E 36.033.00	\$ 18,784.77	\$ 7,666.97	\$ 10,255.19	с .	\$ 73,640,89
						2 Publication Costs	0 30.523.85	a 0.70477	a 7.500.91	a 0.235 19		a 73 0417 08
						3. Consultant Services						d -
								1			;	\$ -
						4 ADP/Computer Services	-					\$ - \$ -
						5 Subawards/Consortium/Contractual Costs	-	<u> </u>				*
						6 Equipment or Facility Rental/User Fees	-	l				\$ -
						7. Alterations and Renovations	P 60 500 00	g 50.000.00	B 37 500 00	# PC FOC CT		\$ .
						8 Other - Testing	\$ 53,500 00					\$ 250,000.00
						9 Other - Meetings and Conferences		\$ 39,205.05				\$ 215 251.14
						Total Other Direct Gosts	5 118,023 04	3 111.489.82	\$ 102,533.66	3 114,960 24	s 9188527	\$ 538 892 03
						G. Direct Costs and Modified Direct Costs						
						Direct Costs	\$ 512,263.59	\$ 496.592.31	\$ 487,636.15	\$ 485.6B0.49	\$ 402,693,86	\$ 2,384.866.40
						Modified Direct Costs						\$ -
						H Indirect Costs						
				IDC Rate>	8%	1 Indirect Cost Type	\$ 40,981 09	\$ 39,727,39	88 010,665	\$ 38.854.44	\$ 32,215,51	\$ 190,789.31
				IDC Rale->		2. Indirect Cost Type						\$ -
						I. Total Direct and Indirect Costs						
						Direct + Indirec:	\$ 553,244 68	\$ 536,319.70	\$ 526,647.05	\$ 524,534.92	\$ 434 909 36	
						. Fee						
						Fuc						
						K. Total Costs and Fee						
						Total Costs	\$ 553,244 68	\$ 536,319.70	\$ 526,647.05	\$ 524,534.92	\$ 434 909 36	\$ 2,575 655.72
						L						

	Internation	al Travel				TOTA.
International Contenences (MED yier ru)						
Hotel May Fer Dien:	5 3 276	3 3 276	3 3.276	9 3,275	9 3,276	
Moule and the denial Expenses For Diem	5 1687	3 1683	3 1,680	9 1,650	5 1,650	
	S 1,020 to	5 1,7(20,00)	5 1,020,00	5 102000	9 102000	
L Bor Personale	\$ 1,045	5 7,092	5 8,298	5 3.198	5 3.192	
Re Jug		-1	- 1	٠	-	
No Trips on Year	4	4	4	-	-	
total .	5 9,05000	3 9,188 (0)	3 9 tea (i.)	5 900800	5 7 0.8 %	5 45,86,030
Pogora Corten Sos (EMI D Marhill		1	1		1	1
Hatel Max Fer Oxin	5 3,691	9 3,986	9 3,985	5 3.555	5 4,455	
Mark, and Codental Expenses For Over-	5 4,376	9 1.834	9 1.834	\$ 1.834	\$ 2,298	
	5 765 CO	9 1,020,00	5 1 020.00	\$ 1020.00	\$ 1,275.00	
Figni Eximate	5 1 698	3 2.264	3 2.264	9 2.294	5 2,830	
No Cam	4	J	J	4	4	
No. 15 ps in 150a		- 1	- 1		-	<b>-</b>
Facel	5 6,529 50	3 8,706,00	3 8,706,00	9 B.700.00	9 10 882 30	5 48,530.00
0146 Mig (92)		1	I	I	1	1
Hand May For Dem	\$ 1200	5 1 290	5 1,200	5 0,250	5 1,290	I
Missis and the contail promise For Dom.	\$ 416	5 416	2 418	5 414	5 418	
and Stimules	5 315 mil	3 115 100	3 125 00	3 (05.0)	5 05.00	1
L gill Saudi	5 1983	5 11083	5 1,081	5 (,08)	5 (,081	
No. Cars						
No. To pay on their limits.	S Jaw Ca	1 5 0,000 (c)	1 1 5 (3.1%) (3.1)	5 31%6 nn	S Jumpan	5 19,40mm0
No Trips on New Tr	S Juge go	5 Joseph (00	2 5 3 196 00	1 5 379630	1 5 7 (96 m)	S 1246000
No. Topo ser Sear Testa" Laure Sy (Cara) (3 - de Sc.) - Statie Esp Hallet Me Per Deur	5 June (a)	1 5 June 100	2 3 3 1996 00 3	1 5 309600 5	1 5 / (1900 m) 5	5 13,46000
No. Trips ser Sear Fear Francis (Cara) (3 - de 9 ) - Stantiets Hallot Feb Per Deur Meile und Federal Exercise (50 Deur	5 5 000 5 3,822	1 5 Juner (co. 3 -	1 3 3 199 and 3 4	1 5 30%600 5 5	T S Josephine S S	5 1986mc
No. Type on two tests of the properties of the p	5 3,822 5 1,107 00	5 J. 100 CO 3 9 9	5	1 5 3046 m 5 5 5		S instruct
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Annual Conference (*1 CY2)			E Prope	4 Days		
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MS, Expenses Per Diem	5	188.00	13		5	2,760,00
Domestic Transport	5	414,31	13		5	\$ 143.10
Entring/Scoplins	5	149.00	1		5	146.16
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					5	10 222 39
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tats Experies Per Dom	s	188.00	19	2	s	5,520,00
Dimension Transport	s	414.31	19	1	s	4,143.10
Printing/Supplies	5	1 974.50	- 1	1	s	1,974.60
Trunslation and Sound	9	1.974.64	1	3	5	3 923 92
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Most rgs/Conference Costs	\$ 27,59,08	\$ 37,0805	5 D. B. N.	5 3	7.705 JV	4	21,885,27	
Field Travel Costs	Y1	TY2	Y3	CYL		СY	2	
Fox/oxgra-Pr. Lenn	\$ 15,000.00	\$ 30,000,00	\$ 30,000,00	5 2	3,500,00	5		•
Fac	\$ 4,990.00	\$ 2,590,00	\$ 4,999.00	S .	4,590.00	5	1,360,00	•
Vehicle Meinignance Coviu	5 1,000,00	5 1,000,00	5 1 000,00	5	1 000 00	s	1,000,00	-
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IIIIAI	\$ 90,188,00	\$ 64,626,00	\$ 62,636,00	\$ 5	1,367.00	\$	12,360 00	
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A Ku at	267	26.2	5 1.5?	S	40.09	]		
fela'u	436	43.6	\$ 1.53	S	67.01	]		
4cuta	668	66.8	5 1.32	5	102.20	]		
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	30000	3000	5 1.38	5	£ 390 00			

Field Supplies				Y1		Y2		Y3		071		OY	2	
Туре	Units/Price	Price	Qy Needed	Tota		Qy Needed To	otal	Qy Needed To	/la	Qy Needed To	otal	Qy Needed	ัยโล	٦
Nunc 1 8 internal crypy als	450	5 516.42	12	5	6.19 / 04	1/ \$	8,779.14	8.5	4,131 36	10 \$	5,164.20			
Serum tubes 10 mL	100	\$ 31.09	25	\$	777.25	23 \$	715.07	12 \$	373.08	12 \$	373.08			
Serum tubes 3 mL	100	5 27.31	2	5	54 62	U \$		1 5	27.31	\$				
Vaccutatiner holder	250	\$ 20.77	50	\$	1,038.50	50 \$	1,038.50	5		\$	-			
21g 1m vacutainer safety needles	48	\$ 39.92	30	5	1.197.60	28 \$	1,117.76	15 5	598 80	14 \$	558.88			
18g Lin vacuta ner needles	100	\$ 32.19	5	5	160.95	5 \$	160.95	3 \$	96.57	2 \$	64.38			
3culuer lock syringe	100	\$ 21.80	2	5	43.G0	5		1 \$	21.80	5				
22g lin need e	100	\$ 19.80	2	5	39.60	\$		1 \$	19.80	s	-			
Cotton Swabs	500	\$ 400.00	5	5	2,000.00	10 \$	4,000.00	0 \$		5.5	2,000.00			
Gloves	50	\$ 29.65	7	5	207.55	14 \$	415.10	0.5	-	6 \$	177.90	0	, .	
Masks N95	50	\$ 47.75	7	\$	334.25	7 \$	334.25	7 \$	334.25	5 \$	238.75	0	;	
Tyvek Suits (project supply)	25	\$ 350.00	52	5	18,200 00	U \$		0.5	-	U \$		0	, .	
Googgles (project supply)	12	\$ 135.00	2	5	270.00	0 \$	-	0 \$	-	0 \$	-	0		
Booties (project supply)	200	\$ 300.00	/	5	2,100,00	U \$		0.5	-	U \$		0		
Biohazard Bag (project supply)	200	\$ 155.00	2	\$	310.00	0 \$		0 \$	-	0 \$	-	0		
Sharps Container	20	\$ 200.00	1	5	200.00	1 \$	200.00	1 5	200.00	1 \$	200.00			
Alcohol Swabs	1200	\$ 32.00	4	\$	128.00	7 \$	224.00	2 \$	64.00	4 \$	128.00			
Tablets	1	\$ 200.00	5	\$	1,000.00	5		\$		5			;	
Dry Ice	1 week	\$ 150.00	6	\$	900.00	12 \$	1,800.00	12 \$	1,800.00	9 \$	1,350.00		, .	
Eye wash station	1	\$ 275.00	1	5	275.00	5		5		5			;	
Card reader system		\$ 1,500.00	1	5	1,500.00	\$		5		\$	-		, .	┙
TOTALS				\$	36,933.96	\$	18,784.77	s	7,666.97	\$	10,255.19		<u> </u>	╝

Nunc 1.8 internal cryon als PCRs - 2 swabs/animal or person Serum tubes 10 mL 1 per furman, 2 per animal serology Serum tubes 3 mL 1 per chicken serology 100 21g Inn vacutainer holder 100 18g Lin vacutainer needles 1 per chiman serology 18g Lin vacutainer needles 1 per chiman serology 3cc luer lock syringe 1 per chicken serology 22g Iin needle 1 per chicken serology Cotton Swabs PCRs 2 swabs/animal or person

TESTS
MIRRS PCR Humans
MIRRS PCR Sequencing of positives
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A PCR Sequencing of positives
MIRRS Secrobay Humans PP
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YEAR 1 YEAR 2								YEAR 3					PTI	ON YEAR	:	OPTION YEAR 2				$\neg$						
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Subject: Re: [Non-DoD Source] FRBA14-6-2-0471 Clarifications
All active links contained in this email were disabled. Please verify the identity of the sender, and confirm the authenticity of all links contained within the message prior to copying and pasting the address to a Web browser.
Much appreciated (b)(6)
I cannot imagine how you get through all these documents. We only have a few proposals/awards to justify and it seems like an enormous task for us!
If I may help you quickly locate certain documents in our forms and files, please call or text my handphone anytime day or night.
Many thanks again!
-Aleksei
Aleksei Chmura, PhD Chief of Staff
EcoHealth Alliance 520 Eighth Avenue, Suite 1200 New York, NY 10018-4182
(b)(6) < tel:1.646.413.3437 > (mobile)
Caution-www.ecohealthalliance.org < <a href="http://caution-www.ecohealthalliance.org/">http://caution-www.ecohealthalliance.org/</a> <a href="http://www.ecohealthalliance.org">Caution-http://www.ecohealthalliance.org</a> >
Catton https://www.ceonedithanianeco.org
EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation.
On Sep 2, 2020, at 12:17, (b)(6) < mailta (b)(6) > wrote:
Yes , they were received. Thank you
Original Message From: William B. Karesh <a href="mailto:karesh@ecohealthalliance.org"> &lt; href="mailto:karesh@ecohealthalliance.org"&gt; &lt; href="mailto:k</a>
Caution-mailto:karesh@ecohealthalliance.org >>
Sent: Wednesday, September 2, 2020 11:16 AM  To (b)(6)
< mailto(b)(6) >>
Cc: Aleksei Chmura@ecohealthalliance.org < <u>mailto:chmura@ecohealthalliance.org</u> > < Caution- <u>mailto:chmura@ecohealthalliance.org</u> >>
Subject: Fwd: [Non-DoD Source] FRBA14-6-2-0471 Clarifications

All active links contained in this email were disabled. Please verify the identity of the sender, and confirm the

authenticity of all links contained within the message prior to copying and pasting the address to a Web browser
Dear(b)(6)
I just wanted to confirm that you received the email below earlier this week.
Thanks!!
William B. Karesh, D.V.M Executive Vice President for Health and Policy
EcoHealth Alliance 520 Eighth Avenue, Suite 1200 New York, NY 10018 USA
+1.212.380.4463 (direct) +1.212.380.4465 (fax) Caution-Caution-www.ecohealthalliance.org < http://caution-caution-www.ecohealthalliance.org/> < Caution-http://caution-Caution-www.ecohealthalliance.org/> < Caution-mailto:karesh@ecohealthalliance.org < Caution-mailto:karesh@ecohealthalliance.org > >
President, OIE Working Group on Wildlife
Co-chair, IUCN Species Survival Commission - Wildlife Health Specialist Group
EPT Partners Liaison, USAID Emerging Pandemic Threats - PREDICT-2 Program
EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation.

Begin forwarded message:

From:Aleksei Chmura <a href="mailto:chmura@ecohealthalliance.org">chmura@ecohealthalliance.org</a> <a href="mailto:chmura@ecohealthalliance.org">chmur

Subject:Re: [Non-DoD Source] FRBA14-6-2-0471 Clarifications

Date:August 30, 2020 at 8:14:36 PM EDT

To <sup>(b)(6)</sup>		
< <u>mailto</u>	Caution	1-
Caution	> >>	
C(p)(g)		]
< <u>mailte</u>		'aution-
Cautio		▶ >>, Billy Karesh
<pre><karesh@ecohealthalliance.org <<u="">mailto:karesh@ecohealthalliance.org&gt; &lt;</karesh@ecohealthalliance.org></pre>		
$Caution-\underline{mailto:karesh@ecohealthalliance.org} \ge \le Caution-Caution-\underline{mailto:karesh@ecohealthalliance.org}$	<u>ohealthall</u>	iance.org <
Caution-mailto:karesh@ecohealthalliance.org > >>, Whitney Bagge 	thalliance	e.org
< mailto:bagge@ecohealthalliance.org > < Caution-mailto:bagge@ecohealthalliance.org		
$Caution-\underline{mailto:bagge@ecohealthalliance.org} \leq Caution-\underline{mailto:bagge@ecohealthalliance.org} \leq Caution-mailto:bagge@ecohe$	ce.org >	>>, Catherine
$Machalaba \leq machalaba @ecohealthalliance.org \leq \underline{mailto: machalaba} (\underline{w}ecohealthalliance.org) \leq mailto$	<u>org</u> > <	
$Caution-\underline{mailto:machalaba@ecohealthalliance.org} \ge < Caution-Caution-\underline{mailto:machalaba@ecohealthalliance.org}$	<u>aba@eco</u> l	<u>healthalliance.org</u> <
$Caution-\underline{mailto:machalaba@ecohealthalliance.org} \ge > \ge \texttt{, Joe Riccardi@ecohealthalliance.org}$	ealthalliar	nce.org
$\leq\!$	> < <u>ero.</u> >	Caution-
$Caution-\underline{mailto:riccardi@ecohealthalliance.org} \leq Caution-\underline{mailto:riccardi@ecohealthalliance.org} \leq Caut$	liance.org	>>>

Dear (b)(6)

Please find attached seven files:

- 1) FRBAA14-6-2-0471 Clarifications FINAL.xlsx
- 2) EHA Budget Justification FINAL.docx
- 3) EHA Documentation\_FINAL.pdf
- 4) Human Link Budget Justification\_FINAL.docx
- 5) Human Link Documentation\_FINAL.pdf
- 6) JUST Budget Justification FINAL.docx
- 7) JUST Documentation FINAL.pdf

There are three sets of paired files with the requested documentation (PDFs) and track-change budget justifications (MS Word) for EcoHealth Alliance and our two subcontracts under this proposal. We have also included the clarifications (MS Excel) with responses to the specific questions in each tab.

Please let me know, if there are additional questions or any other documentation is required.

Many thanks!

-Aleksei

Aleksei Chmura, PhD Chief of Staff & Authorized Organizational Representative

EcoHealth Alliance 520 Eighth Avenue, Suite 1200 New York, NY 10018-4182

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-1.212.380.4473 \le \text{tel}: 1.212.380.4469 \le \text{tel}: 1.212.380.4469 \ge \text{(office)}
(b)(6) \Rightarrow (mobile)
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Caution-Caution-www.ecohealthalliance.org < <a href="http://caution-caution-www.ecohealthalliance.org/">http://caution-caution-www.ecohealthalliance.org/</a> < Caution-Caution-<a href="http://www.ecohealthalliance.org/">http://www.ecohealthalliance.org/</a> < Caution-Caution-<a href="http://www.ecohealthalliance.org/">http://www.ecohealthalliance.org/</a> >

EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation.

From:	Aleksei Chmura
To:	(b)(6)
Cc:	(b)(6) Whitney Bagge; Joe Riccardi; Billy Karesh
Subject:	Re: [Non-DoD Source] FRBA14-6-2-0471 Clarifications
Date:	Thursday, September 17, 2020 11:38:05 AM
Attachments:	DTRA Jordan Budget 28 Aug 2020.xlsx
	ontained in this email were disabled. Please verify the identity of the sender, and confirm the links contained within the message prior to copying and pasting the address to a Web browser.
Dear (b)(6)	
	scel file with a breakdown of EcoHealth Alliance costs and with separate tabs for our subcontracts.
•	if you have additional questions or need other documents.
Many thanks!	
-Aleksei	
Aleksei Chmura, Chief of Staff &	PhD
	nizational Representative
EcoHealth Allian 520 Eighth Aven New York, NY 1	ue, Suite 1200
+1.212.380.4473 (b)(6)	< tel:1.212.380.4469 > (office)
	/ (mobile)  ohealthalliance.org < Caution- <u>http://www.ecohealthalliance.org/</u> >
EcoHealth Allian	ice develops science-based solutions to prevent pandemics and promote conservation.
< <u>mailt(</u> b)(6)	2020, at 11:23, Lyles, Latrice D CTR DTRA AL (USA) (b)(6) > wrote:
Good morni	ng,
Thank you f EHA and subs?	or documents as they will come in handy. Would you be able to provide an excel breakdown of
From: Aleks	Message sei Chmura <chmura@ecohealthalliance.org <<u="">mailto:chmura@ecohealthalliance.org&gt; &gt; esday, September 2, 2020 12:34 PM</chmura@ecohealthalliance.org>

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Traveler 3. Attending Meeting in VA											
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Υ.					Base	A. Senior/Key Personnel						
12	12	8         8         8         8         8         8         9         8         \$           12         12         12         12         \$         \$         3         3         \$         \$         2.5         2.5         \$			\$ 127,940.00	Dr Ehab Abu-Basha, CoPI		\$ 127,940.00	\$ 127,940.00		\$ 127,940,00	\$ 639 700.00
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8	В	8	В	8	\$ 19,301,80	Hani Talafha, Biologist	\$ 12,867.87	\$ 12.867.87	\$ 12,867.87	\$ 12.867.87	\$ 12 867 87	\$ 64 339.33
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2	12	12	12	- 2	\$ 32,520,00	Zaidoun Hijazeen, Veterinarian	\$ 5,420.00			\$ 32,520,00	\$ 32,520,00	\$ 135,500,00
						Zaidoun Hijazeen Fringe	\$ ·	\$ -	-	\$ -	\$ -	\$ -
3	3	3	3	- 3	\$ 5439153	Mustafa Ahahneh Virologist/Lah Director	\$ 13,597.88			\$ 13,597.88	\$ 13 597 88	\$ 67 989 41
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						Zuhair Bani Ismail Fringe	3 .	\$ -	\$ -	\$ -	\$ -	3 -
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						Borhan Al-zughoul Fringe	\$ -	5		<u>s</u> -	5 -	\$ .
3	3	3	3	3	\$ 23,112,00	Bilal Al-Omari, Lab Supervisor	\$ 5,778.00	\$ 5.778.00	\$ 5,778.00	\$ 5.778.00	\$ 5.778.00	\$ 28.890.00
						Bilal Al-Omari Fringe	\$ -	\$ -	\$ -	\$ -	\$ -	\$ -
1	1	-	1	1	\$ 77.018.63	Wall Hayajneh, Human ID Expert	\$ 14,751.55	\$ 14,751.56	\$ 14,751.55	\$ 14,751.55	\$ 14.751.55	\$ 73.757.76
						Wail Hayajneh Ennge	5 -	3 -		\$ ·	S -	s -
			H	-ringo Rato-		Total SemonKey Personnel	\$ 209,817.71	\$ 236,917,71	\$ 236,917.71	\$ 236,917,71	\$ 236 917 71	\$ 1,157.488.57
						B. Other Personnel		]				
1	1	-	2	2	\$ 58 241.66	Saad Charaibeh, Avian Pathologist	\$ 4,853.47	\$ 4,853.47	\$ 4,853.47	\$ 9,706.94	\$ 9.706.94	\$ 33,974.30
						Saad Gharaibeh Fringe	S -	3 -		\$ -	S -	s ·
3.5	3.5	3.5	3.5	3.5	\$ 28 219,16	Admin	\$ 8,230.59	\$ 8,230,59	\$ 8,230.59	\$ 8,230,59	\$ 8 230 59	\$ 41 152.94
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12	3 3 3 3 \$ 23  1 1 1 1 1 \$ 77    Fringe Rate		\$ 338508	Graduate Student Stipends (x2)	\$ 6,770.16	\$ 6,770.16	\$ 6,770.16			\$ 33,850,80		
	_					Total Other Personnel	\$ 40,885 34				\$ 29 785 37	
						C. Equipment			,			
	2 12 12 12 12 2 \$ 2.53 2 12 12 12 12 12 5 3.38			Vehicle	\$ 37,150.00	\$ 1,830.00	\$830.00	\$ 1.830.00	s - 830 co	\$ 44 470.00		
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	3 3 3 3 3 3 5 5 2.5 2.5 2.5 3 5 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3					Total Equipment	\$ 65,996.00	\$ 1.830.00	\$ 1,830.00	\$ 1.830.00	\$ 1830.00	\$ 73,316,00
						D. Travel	2 00,500 00	4 1.403.00	2 ,500 00		2 000 00	. 15010.30
	3 3 3 3 3 3 5 5 5 5 5 5 5 5 5 5 5 5 5 5					1 Domestic Travel	\$ 40,108.00	\$ 62,626,00	\$ 62,626.00	\$ 51,367,00	\$ 12,360,00	\$ 229 087.00
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	2 12 12 12 12 12 \$ 2 2 12 12 12 12 12 \$ 3			Total Travel	\$ 77,541.50							
						L. Participant/Traince Support Costs	Ø 17,041.20	9 083710307	<b>2</b> 09.3 07.0	J 70.237 W	0 4E 273.70	3 3/11/00/00
						Tuition/Fees/Health Insurance						\$ .
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						Total Participant/Trainee Support Costs	3 -	\$ -	\$ ·	\$ -	\$ -	\$ -
						F. Other Direct Costs	<u> </u>				_	
						1 Materials and Supplies	\$ 36,933.96	\$ 18,784.77	\$ 7,666.97	\$ 10,255.19		\$ 73,640,89
						2 Publication Costs						\$ ·
						3. Consultant Services		1				\$ -
						4 ADP/Computer Services	1					\$ -
						5 Subawards/Consortium/Contractual Costs						3 -
	1			6 Equipment or Facility Rental/User Fees	1					\$ ·		
						7. Alterations and Renovations						\$ -
						8 Other - Testing	\$ 53,500.00					\$ 250,000.00
						Other - Meetings and Conferences	\$ 27,589.08					\$ 215 251.14
						Total Other Direct Costs	\$ 118,023.04	\$ 111,489.82	\$ 102,533.66	\$ 114,960.24	\$ 91 885 27	\$ 538 892 03
						G. Dried Costs and Modified Direct Costs	1					
						Direct Costs	\$ 512,263.59	\$ 496.592.31	\$ 487,636.15	\$ 485,680,49		\$ 2,384.866.40
						Modified Direct Costs						\$ -
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						1 Indirect Cost Type	\$ 40,381 09	\$ 39,727,39	\$ 39,010,89	\$ 38.854.44	\$ 32 215 51	\$ 190,789.31
				IDC Rale->	>	2. Indirect Cost Type						\$ -
						I. Total Direct and Indirect Costs		]				
						Direct + Indirect	\$ 553,244 68	\$ 536,319.70	\$ 526,647.05	\$ 524,534.92	\$ 434 909 36	
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						Tatal Casts	\$ 553,244 RR	\$ 536 319 70	\$ 526,647.05	\$ 524.534.92	\$ 434,909,36	\$ 2,575,656,72
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	5 765 CO	9 1,020,00	5 1 020.00	\$ 1020.00	\$ 1,275.00	
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No. Topo ser Sear Testa" Laure Sy (Cara) (3 - de Sc.) - Statie Esp Hallet Me Per Deur	5 June (a)	1 5 June 100	2 3 3 1996 00 3	1 5 309600 5	1 5 / (1900 m) 5	5 13,46000
No. Trips ser Sear Fear Francis (Cara) (3 - de 9 ) - Stantiets Hallot Feb Per Deur Meile und Federal Exercise (50 Deur	5 5 000 5 3,822	1 5 Juner (co. 3 -	1 3 3 199 and 3 4	1 5 30%600 5 5	T S Josephine S S	5 1986mc
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No. Topis soch Sections (1994) And Sections (1994) And Sections (1994) And Sections (1994) And Section (1994	5 June Co 5 Super Co 5 Super Co 5 Super Co 5 Linor Co 6 Super Co 7 Co 5 19,546 Co	2 Jules CO	2 3 766 371 5 3 766 371 5 4 5 5 6 7 6 7 7 6 3 7 6 7 7 7 8 8 9 9 7 9	5 3796 m 5 5 296,00	5 / 7 / 7 / 7 / 7 / 7 / 7 / 7 / 7 / 7 /	5 12.569.00 5 24.989.00
We Type see New York Transport State (St. 64 %). Maintest States (St. 64 %). Mic Type St. 64 %). Mic Type St. 64 % (St. 64 %). Mic T	5 5,000 00 5 5,822 5 1107 00 5 2,840 7 6 5 13,596 00 5 14,547 00 5 4,547 00 5 4,547 00 5 4,547 00 5 4,547 00 5	2	2 3 756 00 5 5 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	1 5 3746 m 5 5 5 5 5 7 7 7 7 9 9 5296,00	5 / 746 m 5 / 746 m 5   5   5   5   5   5   5   5   5   5	\$ 12,569.00 \$ 26,989.00 6 7,620.00
No. Topis soch Sections (1994) And Sections (1994) And Sections (1994) And Sections (1994) And Section (1994	5 June Co 5 Super Co 5 Super Co 5 Super Co 5 Linor Co 6 Super Co 7 Co 5 19,546 Co	2 Jules CO	2 3 766 371 5 3 766 371 5 4 5 5 6 7 6 7 7 6 3 7 6 7 7 7 8 8 9 9 7 9	5 3796 m 5 5 296,00	5 / 7 / 7 / 7 / 7 / 7 / 7 / 7 / 7 / 7 /	S 12,569.00 S 24,989.00
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Annual Conference (*1 CY2)			E Prope	4 Days		
Hote May Fer Diem	9	246.00	13		5	4 920 00
MS, Expenses Per Diem	5	188.00	13		5	2 760 00
Domestic Transport	5	414,31	13		5	\$ 143.10
Entring/Scoplins	5	149.00	1		5	146.16
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Peem Ricola	5	1,750,00	1	1	5	1.750 %
ture and Coffic	s	46.03	45		s	2,079 90
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RickUff Meeting (#1 Driv)			A Proper	4 Days		
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Fig. 10g/Sulpho	5	65.60	1		s	65.50
Trunslation and Sound	5	1.974.64	1		5	1 974 94
Recar Rynta	5	1.836.00	1	1	5	1 350 00
Lance and Coffine	5	40.35	20	1.	4	520.00
					5	10 222 39
Regional Meeting (V2, OVI), OV2 CNLVI			t People	A Days	Т	
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Papara Transport - Lesanon	5	409.00			5	A10.00
Pepara Transport Fran	5	49 J 30			5	\$96.00
Hair Mo Pri Dein	5	740, 0.0	h		4	1,690,00
M& Lapresa's Per Denn	5	1.08.00	h		4	2.777.00
Durish of and Sound	5	1574.63	1		5	~523.00
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<mailtd(b)(6)< th=""></mailtd(b)(6)<>
Subject: Re: [Non-DoD Source] FRBA14-6-2-0471 Clarifications
All active links contained in this email were disabled. Please verify the identity of the sender, and confirm the authenticity of all links contained within the message prior to copying and pasting the address to a Web browser.
Much appreciated (b)(6)
I cannot imagine how you get through all these documents. We only have a few proposals/awards to justify and it seems like an enormous task for us!
If I may help you quickly locate certain documents in our forms and files, please call or text my handphone anytime day or night.
Many thanks again!
-Aleksei
Aleksei Chmura, PhD
Chief of Staff
EcoHealth Alliance 520 Eighth Avenue, Suite 1200 New York, NY 10018-4182
(b)(6) > (mobile) > (aution-www.econeannamance.org < http://caution-www.ecohealthalliance.org/> < Caution-http://www.ecohealthalliance.org >
<del></del>
EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation.
On Sep 2, 2020, at 12:17 (b)(6) <mailto: (b)(6)=""> wrote:</mailto:>
Yes, they were received. Thank you
Original Message From: William B. Karesh < karesh@ecohealthalliance.org < <u>mailto:karesh@ecohealthalliance.org</u> > < Caution- <u>mailto:karesh@ecohealthalliance.org</u> >>
Sent: Wednesday, September 2, 2020 11:16 AM To(b)(6)
< <u>mailto</u>
Cc: Aleksei Chmura <chmura@ecohealthalliance.org <<u="">mailto:chmura@ecohealthalliance.org&gt; &lt; Caution-<u>mailto:chmura@ecohealthalliance.org</u>&gt;&gt; Subject: Fwd: [Non-DoD Source] FRBA14-6-2-0471 Clarifications</chmura@ecohealthalliance.org>

All active links contained in this email were disabled. Please verify the identity of the sender, and confirm the

uthenticity of all links contained within the message prior to copying and j	parting the address to a work now.
Dear (b)(6)	
I just wanted to confirm that you received the email below earlier this	week.
Thanks!!	
William B. Karesh, D.V.M Executive Vice President for Health and Policy	
EcoHealth Alliance 520 Eighth Avenue, Suite 1200 New York, NY 10018 USA	
+1.212.380.4463 (direct) +1.212.380.4465 (fax) Caution-Caution-www.ecohealthalliance.org < <a href="http://caution-caution-vution-http://caution-Caution-www.ecohealthalliance.org/">http://caution-caution-vution-vution-http://caution-Caution-www.ecohealthalliance.org/</a> < Caution-mailto:karesh@ecohealthalliance.org < Caution-mailto:karesh@ecohealthalliance.org	·
President, OIE Working Group on Wildlife	
Co-chair, IUCN Species Survival Commission - Wildlife Health Speci	ialist Group
EPT Partners Liaison, USAID Emerging Pandemic Threats - PREDIC	T-2 Program
EcoHealth Alliance develops science-based solutions to prevent pande	emics and promote conservation.
Begin forwarded message:	
From:Aleksei Chmura <chmura@ecohealthalliance.org <<u="">mailto:chmu ution-<u>mailto:chmura@ecohealthalliance.org</u> &gt; &lt; Caution-Caution-<u>mailto</u></chmura@ecohealthalliance.org>	

Caution-mailto:chmura@ecohealthalliance.org > >>

Subject:Re: [Non-DoD Source] FRBA14-6-2-0471 Clarifications

Date:August 30, 2020 at 8:14:36 PM EDT

To <sup>(b)(6)</sup>	
< <u>mailto</u> auti	ion-
Caution > >	>
Cd(p)(e)	
< <u>mailto</u>	aution-
Caution	>>, Billy Karesh
<pre><karesh@ecohealthalliance.org <<u="">mailto:karesh@ecohealthalliance.org&gt; </karesh@ecohealthalliance.org></pre>	<del></del>
$Caution-\underline{mailto:} karesh \underline{@ecohealthalliance.org} > < Caution-Caution-\underline{mailto:} karesh \underline{@ecohealthalliance.org} > < < Caution-Caution-\underline{mailto:} karesh \underline{@ecohealthalliance.org} > < < Caution-\underline{mailto:} karesh \underline{@ecohealthalliance.org} > < < Caution-Caution-\underline{mailto:} karesh \underline{@ecohealthalliance.org} > < < Caution-Caution-\underline{mailto:} karesh \underline{@ecohealthalliance.org} > < < Caution-Caution-\underline{mailto:} karesh \underline{@ecohealthalliance.org} > < < Caution-$	alliance.org <
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Caution-mailto:machalaba@ecohealthalliance.org > >>, Joe Riccardi <riccardi@ecohealthalliance.org< td=""><td>liance.org</td></riccardi@ecohealthalliance.org<>	liance.org
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Dear(b)(6)

Please find attached seven files:

- 1) FRBAA14-6-2-0471\_Clarifications\_FINAL.xlsx
- EHA Budget Justification\_FINAL.docx
- 3) EHA Documentation\_FINAL.pdf
- 4) Human Link Budget Justification\_FINAL.docx
- 5) Human Link Documentation\_FINAL.pdf
- 6) JUST Budget Justification\_FINAL.docx
- 7) JUST Documentation\_FINAL.pdf

There are three sets of paired files with the requested documentation (PDFs) and track-change budget justifications (MS Word) for EcoHealth Alliance and our two subcontracts under this proposal. We have also included the clarifications (MS Excel) with responses to the specific questions in each tab.

Please let me know, if there are additional questions or any other documentation is required.

Many thanks!

-Aleksei

Aleksei Chmura, PhD Chief of Staff & Authorized Organizational Representative

EcoHealth Alliance 520 Eighth Avenue, Suite 1200 New York, NY 10018-4182

+1.212.380.4473 < tel:1.212.380.4469 < tel:1.212.380.4469 > > (office) (b)(6) > (mobile)

Caution-Caution-www.ecohealthalliance.org < <a href="http://caution-caution-www.ecohealthalliance.org/">http://caution-caution-www.ecohealthalliance.org/</a> < Caution-Caution-<a href="http://www.ecohealthalliance.org/">http://www.ecohealthalliance.org/</a> < Caution-Caution-<a href="http://www.ecohealthalliance.org/">http://www.ecohealthalliance.org/</a> >

EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation.

From:	(b)(6)
To:	"William B. Karesh"; "Aleksei Chmura"
Cc:	(b)(6)
	(USA); DTRA Ft Belvoir SI Mailbox SI-FMKA SPS Contracts; "onr_boston@navy.mil" (b)(6)
	(b)(6)
Subject:	Jordan Award
Date:	Tuesday, September 29, 2020 4:22:00 PM
Attachments:	Terms and Conditions (DTRA-specific) 14Sep2020 FINAL.pdf
	HDTRA1-20-1-0029 Award.pdf
	EHA Jordan SOW.pdf

Attached you will find the award documentation for grant HDTRA1-20-1-0029: Reducing the Threat of Middle East Respiratory Syndrome Coronavirus and Avian Influenza in Jordan & Strengthening Regional Disease Surveillance Capacity. Please confirm once you have received this email, thank you.

V/r
(b)(6)
Contract Specialist • Broadleaf
Defense Threat Reduction Agency (DTRA)
Office Phone (b)(6)
(b)(6)

# DEFENSE THREAT REDUCTION AGENCY (DTRA) GENERAL TERMS AND CONDITIONS FOR GRANT AWARDS

# **Table of Contents**

Tabl	e of Contents1
1.	Terms and Conditions Incorporated by Reference 2
2.	Acceptance of Grant2
3.	Recipient Responsibilities2
4.	Standards for Financial Management Systems2
5.	Modification of the Grant2
6.	Payments3
7.	Funding Increments and/or Options
8.	Patent Rights 4
9.	Technical Reporting Requirements4
10.	Financial Reporting Requirements7
11.	Delegation of Administration Duties7
12.	Security
13.	Representations and Assurances8
14.	Data Collection
15.	Publications and Acknowledgement of Sponsorship13
16.	Authorization to Perform Activities Abroad13
17.	Inconsistency between English Version and Translation of Grant 14
18.	Value Added Tax (VAT) and Other Taxes14
19. Fede	Prohibition on the Use of Cooperative Threat Reduction Funds in the Russian

# 1. Terms and Conditions Incorporated by Reference.

The DoD Research and Development General Terms and Conditions, most current version as of the date of grant award, are hereby incorporated by reference and are available for download at website http://www.onr.navy.mil/Contracts-Grants/submit-proposal/grants-proposal/grants-terms-conditions.aspx.

#### 2. Acceptance of Grant.

The recipient is not required to countersign the Grant document; however, the recipient agrees to the conditions specified in the Research Grant and the Articles contained herein unless notice of disagreement is furnished to the Grants Officer within fifteen (15) calendar days after the date of the Grants Officer's signature. In case of disagreement, the recipient shall not assess the Grant any costs of the research unless and until such disagreement(s) is resolved.

# 3. Recipient Responsibilities.

The recipient will bear primary responsibility for the conduct of the research and will exercise judgment towards attaining the stated research objectives within the limits of the Grant's Terms and Conditions.

The Principal Investigator(s) (PI) specified in the Grant award will be continuously responsible for the conduct of the research project and will be closely involved with the research effort. The PI, operating within the policies of the recipient, is in the best position to determine the means by which the research may be conducted most effectively.

# 4. Standards for Financial Management Systems.

Where the Federal Government guarantees or insures the repayment of money borrowed by the recipient, DTRA, at its discretion, may require adequate bonding and insurance if the bonding and insurance requirements of the recipient are not deemed adequate to protect the interest of the Federal Government.

DTRA may require adequate fidelity bond coverage where the recipient lacks sufficient coverage to protect the Federal Government's interest.

Where bonds are required in the situations described above, the bonds shall be obtained from companies holding certificates of authority as acceptable sureties, as prescribed in 31 CFR Part 223, "Surety Companies Doing Business with the United States."

### 5. Modification of the Grant.

The only method by which this Grant may be modified is by a formal, written modification signed by the Grants Officer. No other communications, whether oral or in writing, are valid.

Prior Approvals are required as follows:

- 1) Expenditures on equipment costing \$5,000 or more not specifically identified in the budget at time of award. (Approval via written notification from the Grants Officer.)
- 2) Expenditures for foreign travel not specifically identified in the budget at time of award. (Approval via written notification from the Grants Officer.)

- 3) Prior approval is not required to transfer amounts budgeted for indirect costs to absorb increases in direct costs, or vice versa.
- 4) Prior approval is not required to carry forward an unobligated balance to a subsequent period of performance under this award.

#### 6. Payments.

The 2 CFR 200 governs responsibilities concerning payments, with the following clarifications:

Recipients shall submit requests for payment using Invoicing, Receipt, Acceptance, and Property Transfer (iRAPT) at <a href="https://wawf.eb.mil/">https://wawf.eb.mil/</a>. Any request for advance payments must be approved by the Administrative Grants Office shown in Block 6 of the award. The request shall be submitted to the Administrative Office identified in Block 6 of the Research Grant by entering the following routing codes:

- 1) Pay Office DoDAAC: See Block 12 (Code) on the first page of the Grant.
- 2) Invoice Type: Non Procurement Instrument (NPI) Voucher.
- 3) Issue By DoDAAC: See Block 5 (Code) on the first page of the Grant.
- 4) Admin DoDAAC: See Block 6 (Code) on the first page of the Grant.
- 5) Grant Approver: Same as Admin DoDAAC (Leave Ext. blank).

Payments will be made by the Defense Finance and Accounting Service (DFAS) office specified in the Research Grant (Block 12).

A foreign awardee must have a U.S. bank account and be signed up for electronic payments (electronic funds transfers (EFT)).

# 7. Funding Increments and/or Options.

The recipient is advised that the Grantor's obligation to provide funding for increments and/or options included in the Grant is contingent upon satisfactory performance in the judgment of the DTRA Scientific Officer/Technical Monitor and the availability of funds. Other factors will be considered before options will be exercised (for example, expenditure rate and current programmatic objectives). Accordingly, no legal liability on the part of the Grantor exists unless or until funds are made available to the Grantor and notice of such availability is confirmed in writing to the recipient. Refer to the Funding Profile in Section G of the Grant for additional incremental funding planned, but not currently obligated for the Grant.

Funding Increments – In no event is the Government obligated to reimburse the recipient for expenditures in excess of the total funds allotted by the Government to this agreement. Recipients should note that low expenditure rates reported on payment requests may be cause for deferral of future increments. The Government anticipates unilateral modifications for funding increments.

Options – If the agreement contains Option(s), the Government reserves the right to exercise the Option(s) unilaterally.

# 8. Patent Rights.

Patent Rights are governed by 37 CFR 401.14 with the following clarifications: All DTRA-related disclosures, confirmatory licenses to the government, patent applications, and other communications should be submitted using iEdison (<a href="https://public.era.nih.gov/iedison">https://public.era.nih.gov/iedison</a>), a single web interface for government grantees to report details of inventions and patents.

The 37 CFR Part 401 invention reporting requirements are summarized at iEdison (<a href="https://era.nih.gov/iedison/invention\_timeline.htm">https://era.nih.gov/iedison/invention\_timeline.htm</a>). If the grantee organization is not already an iEdison registrant, then iEdison registration is required prior to submission of the below invention reports. The grant shall not be closed out until all invention reporting requirements are met.

The recipient shall also submit interim and final Reports of Inventions using the DD882 form (<a href="https://www.arl.army.mil/www/pages/218/d882.pdf">https://www.arl.army.mil/www/pages/218/d882.pdf</a>). Interim invention reports shall be submitted annually, listing subject inventions reported during that period or that there are no such inventions. These reports are due no later than 1 July of each year. Grants effective after 31 January will not require a report until 1 July the following year. A final report shall be submitted within ninety (90) days after end of the project, listing all subject inventions or stating that there were no such inventions. These Reports of Inventions should be submitted to:

- 1) The DTRA Grants Officer via email (the address specified in the grant's clause 252.601-9000 may be used);
- 2) The Administrative Officer listed in the Grant via email (Block 6 of the SF-26 award or Block 7 of the SF-30 modification);
- 3) As directed by DTRA, email or portal; and
- 4) E-mailed to <a href="mailed-to-dtrabasicresearch@mail.mil">dtrabasicresearch@mail.mil</a> (file size must be less than 10MB). File should be named by the Grant number and "Invention Report" (e.g. HDTRA1-12-1-9999 Invention Report).

# 9. Technical Reporting Requirements.

**Research Performance Progress Report (RPPR).** Except under rare cases, RPPRs are required annually. The RPPR is due no later than 1 July of each year. Grants effective after 31 January will not require a RPPR until 1 July of the following year.

The RPPR is *not* a cumulative report. The first RPPR shall <u>only</u> include actions that occurred from the Period of Performance start date up to submission of the first RPPR. Each subsequent report shall only include actions that occurred during the 12-month period following the previous year's RPPR.

A RPPR is not required in the final year of the award if the period of performance ends within 60 days of the RPPR due date. In this instance the Final Report will satisfy the requirement. Broadly the RPPR shall address the following items:

- Accomplishments
- Products
- Participants and Other Collaborating Organizations

- Impact
- Changes/Problems

Templates and specific instructions will be provided each year in advance of the submission deadline. All files must be submitted via email to <a href="mailto:dtrabasicresearch@mail.mil">dtrabasicresearch@mail.mil</a> (individual file size must be less than 10MB). A copy of the RPPR should also be provided to the Administrative Office identified in the Grant. The file names should be as follows:

- RPPR: Year Annual Report Grant Number, e.g. 2017 Annual Report HDTRA1-12-1-9999.
- Metrics: Year Metrics Grant Number, e.g. 2017 Metrics HDTRA1-12-1-9999.

**Quad Chart.** An updated quad chart must be submitted annually. A template will be provided each year in advance of the submission deadline. All files must be submitted via email to <a href="mailto:dtrabasicresearch@mail.mil">dtrabasicresearch@mail.mil</a> (individual file size must be less than 10MB). The file name should be as follows:

 Quad Chart: Year Quad Chart Grant Number, e.g. 2017 Quad Chart HDTRA1-12-1-9999.

**Annual Technical Review.** At least one representative (preferably the PI) for each award is expected to attend and present at an annual technical program review meeting, unless otherwise exempted by DTRA in writing. For planning purposes reviews will typically be for two days in Northern Virginia during the spring or summer months.

**Final Technical Report.** A comprehensive final technical report is required: the draft document is required forty-five (45) days prior to the end of the Period of Performance and the final document is required ninety (90) days after the expiration or termination of the award.

The purpose of the final report is to document and to transition the results of the effort into the DTRA and DoD applied research community. The final report will always be sent to the Defense Technical Information Center (DTIC) and unclassified reports may be made available to the public through the National Technical Information Service (NTIS).

The final report is more than an extension of previous annual reports. The final report shall be a **comprehensive** technical summary of the significant work accomplished. The final report, where it is not readily accessible in published form should, where applicable:

- Clearly describe and illustrate the experimental equipment, setup, and procedures;
- Characterize and tabulate collected/computed data in an appendix;
- Sufficiently describe computational codes so they can be reproduced. Include a listing of the code in an appendix if possible and appropriate; and
- When the research effort culminates in the production of one or more student theses or dissertations, in these cases, the most significant advancements and conclusions (equations, figures, relationships, etc.) should be included in an executive summary. The theses or dissertations should be attached as appendices only if they are not readily available. If they are, clearly reference them and how they can be obtained. Also include in the executive summary, cumulative lists of people involved in, and publications

stemming from, the research effort. Do not include copies of already submitted or published articles in the final report.

Standard Form (SF) 298, Report Documentation Page, must be used. Item 13 of the SF-298 should contain a 100 to 200 word abstract summarizing technical progress during the reporting period. The SF-298 may be found on the Internet at: <a href="http://www.gsa.gov/portal/forms/download/116146">http://www.gsa.gov/portal/forms/download/116146</a>

All of the report pages should be prepared for acquisition and distribution by DTIC. All of the report pages should be of good quality for copying purposes. No pages should be missing.

The format and standard required by your institution for the preparation of theses and dissertations shall be used for the final report. In the absence of any institutional standards, you may wish to refer to the American National Standards Institute (ANSI) document Z39.18-1987, "Scientific and Technical Reports: Organization, Preparation, and Production," for guidance. The report may be obtained from:

American National Standards Institute, Inc. 1430 Broadway New York, NY 10018

It is anticipated that all final technical reports will be unclassified and that distribution will not be limited. However, for final technical reports that require a limited distribution as deemed necessary by DTRA, a Distribution List will be provided with the comments on the draft final technical report. The Distribution List should be formatted to match the rest of the report, placed at the end of the report, and added to the Table of Contents. The number of pages in the Distribution List should be added to the total page count and included in the total number of pages cited in Block 15 of the SF-298.

The draft of the final technical report will be due not later than forty-five (45) days prior to the end of the period of performance. The draft of the final technical report (including a draft SF-298) must be submitted electronically as follows:

- Email the draft of the final technical report to <a href="mailto:dtrabasicresearch@mail.mil">dtrabasicresearch@mail.mil</a> (file size must be less than 10MB). The file name should be 'Draft Final Report' and the Grant number, e.g. Draft Final Report HDTRA1-12-1-9999.
- Provide a copy of the report to the Administrative Office identified in the Grant.

Within thirty (30) days, this draft will be reviewed by DTRA and comments will be provided to the Grantee to ensure the report complies with DTRA final report requirements. Such review and comment does not restrict the conduct or reporting of the project research findings/outcomes and, in accordance with Article 35, does not restrict Grantee's ability to publish. Grantee shall incorporate such requested changes so that the report incorporates and complies with agreement final reporting requirements terms. Final Technical Reports are due ninety (90) days after the expiration or termination of the award. The final submission should be made in accordance with the draft final report submission instructions.

**Final Metrics.** A final metrics table (in MS Excel format) is required. A template and specific instructions will be provided in advance of the submission deadline. The final metrics file should be submitted along with the Final Technical Report. The fields contained

in the final metrics file are analogous to those of the annual submissions. The final metrics file shall contain only data from the last annual reporting period until the end of the award's funded Period of Performance.

• Email the final Metrics File to <a href="mailto:dtrabasicresearch@mail.mil">dtrabasicresearch@mail.mil</a> (file size must be less than 10MB). The file name should be 'Final Metrics' and the Grant number, e.g. Final Metrics HDTRA1-12-1-9999.

# 10. Financial Reporting Requirements.

Federal Financial Reports (SF-425) are due no later than 1 July of each year with data "as of" 30 May of that year. Grants effective after 31 January will not require a Federal Financial Report until 1 July of the following year. All financial reports shall be submitted to the Administration Office identified in Block 6 of the Research Grant. In addition, the Federal Financial Report must be submitted electronically as follows:

• Email the Federal Financial Report to <a href="mailto:dtrabasicresearch@mail.mil">dtrabasicresearch@mail.mil</a> (file size must be less than 10MB). The file name should be the Year, 'Federal Financial Report' and the Grant number, e.g. 2015 Federal Financial Report HDTRA1-12-1-9999.

# 11. Delegation of Administration Duties.

Certain grant administration duties have been delegated to the Administration Office identified in Block 6 of the Research Grant. These duties are as follows:

- 1) Provisionally approve all Grant and Cooperative Agreement Vouchers.
- Perform all property administration services except the approval of recipient's requests to purchase equipment with grant funds. Such approvals must be granted by the DTRA Grants Officer.
- 3) Perform all plant clearance functions.
- 4) Approve requests for Registration for Scientific and Technical Information Services (DD Form 1540).
- 5) Obtain all financial report(s) (see Article 10 of this document).
- 6) Execute administrative closeout procedures, which include the following:
  - a. Obtain the final Report of Inventions and Subcontracts (DD Form 882).
  - b. Obtain final payment request, if any.
  - c. Obtain final property report and dispose of purchased property and government furnished equipment (GFE) in accordance with the DoDGARs Part 22, Subpart G.
  - d. Perform a review of final incurred costs and assist the Grants Officer in resolving exceptions, if any, resulting from questioned costs.
  - e. Assure that all refunds due the Government are received by the Grantor.

NOTE: This term and condition is **not applicable** to instrumentation and equipment grant awards.

#### 12. Security.

As a general rule, PI's will not need access to classified security information in the conduct of research supported under this Grant. Should it appear that access to such information is desirable the recipient shall advise the Grantor and request clearance for the investigator. Should information be developed during the course of work under this Grant that, in the

judgment of the PI or the recipient, should be classified, the Grants Officer shall be notified immediately.

# 13. Representations and Assurances.

By accepting funds under this Grant, the recipient assures that it will comply with applicable provisions of the national policies and statutory/regulatory/executive-based requirements detailed below.

**LIVE ORGANISMS.** By signing this agreement or accepting funds under this agreement, the recipient assures that it will comply with applicable provisions of the following national policies concerning live organisms:

# 1) For human subjects:

- a) Adhere to the requirements for protection of human subjects per the DoD level terms and conditions as well as the following DTRA requirements:
- b) The recipient shall adhere to DTRA local clause 252.223-9002 Protection of Human Subjects (Aug 2010). The full text of this clause is as follows:

All research under this grant involving human subjects must be conducted in accordance with 32 CFR 219, 10 U.S.C 980, and DoDD 3216.02, as well as other applicable federal and state regulations. Grantees must be cognizant of and abide by the additional restrictions and limitations imposed on the DoD regarding research involving human subjects, specifically as regards vulnerable populations (32 CFR 219 modifications to subparts B-D of 45 CFR 46), recruitment of military research subjects (32 CFR 219), and surrogate consent (10 U.S.C. 980).

DTRA Directive 3216.01 of June 9, 2010 establishes the DTRA Human Subjects Protection Program, sets forth the policies, defines the applicable terms, and delineates the procedures necessary to ensure DTRA compliance with federal and DoD regulations and legislation governing human subject research. The regulations mandate that all DoD activities, components, and agencies protect the rights and welfare of human subjects of study in DoD-supported research, development, test and evaluation, and related activities hereafter referred to as "research". The requirement to comply with the regulations applies to new starts and to continuing research.

The DTRA directive requires that research using human subjects may not begin or continue until the Defense Threat Reduction Agency's Research Oversight Board (ROB) has reviewed and approved the proposed protocol. Grantees and subcontractors are required to submit a valid federal assurance for their organization (institution, laboratory, facility) that has been issued by either DoD or the Department of Health and Human Services, and documentation of review of proposed protocols by the local Institutional Review Board (IRB) to include consent forms for any planned research using human subjects to the DTRA ROB for its review through the Grants Officer's representative (if assigned) or the Grants Officer. The ROB review is separate from, and in addition to, local IRB review.

A study is considered to involve human research subjects if: 1) there is interaction with the subject (simply talking to the subject qualifies; no needles are required); and

2) if the study involves collection and/or analysis of personal/private information about an individual, or if material used in the study contains links to such information.

Written approval to begin research or subcontract for the use of human subjects under the proposed protocol will be provided in writing from the DTRA ROB, through the Grants Officer. A copy of this approval shall be maintained by both the Grantee and the government. Any proposed modifications or amendments to the approved protocol or consent forms must be submitted to the local IRB and the DTRA ROB for review and approval. Examples of modifications/ amendments to the protocol include but are not limited to:

- a change of the PI;
- changes in duration or intensity of exposure to some stimulus or agent;
- changes in the information requested of volunteers, or changes to the use of specimens or data collected; or
- changes in perceived or measured risks or benefits to volunteers that require changes to the study.

Research pursuant to such modifications or amendments shall not be initiated without IRB and ROB approval except when necessary to eliminate apparent and immediate hazards to the subject(s).

Research projects lasting more than one year require IRB review at least annually, or more frequently as required by the responsible IRB. ROB review and approval is required annually. The Grantee or subcontractor must provide documentation of continued IRB review of protocols for ROB review and approval in accordance with these Terms and Conditions. Research must not continue without renewed ROB approval unless necessary to eliminate apparent and immediate hazards to the subject(s).

Non-compliance with any provision of this clause may result in withholding of payments under the grant pursuant to the grant's payments clause(s) and/or grant termination pursuant to the grant's termination clause(s). The government shall not be responsible for any costs incurred for research involving human subjects prior to protocol approval by the ROB.

# 2) For animals:

- a. Adhere to the requirements for protection of animal subjects per the DoD level terms and conditions as well as the following DTRA requirements:
- b. DTRA local clause 252.235-9001 Prohibition of Use of Laboratory Animals (Jul 2010). The full text of this clause is as follows:

The grant recipient shall obtain approval from the US Army Medical Research and Material Command (MRMC), Animal Care and Use Review Office (ACURO) prior to conducting research on live nonhuman vertebrates. Studies involving non-human primates, dogs, cats, or marine mammals will require a site visit by an ACURO laboratory animal veterinarian as a condition of approval. DoD may also conduct site visits involving research on other animals when deemed appropriate. The animal

research facility is responsible for notifying the DoD sponsor if Association for the Assessment and Accreditation of Laboratory Animal Care accreditation is lost or the facility is under USDA inspection. DoD also has the right to a site inspection under these circumstances.

The grant recipient (including subcontractors) is expressly forbidden to use laboratory animals in any manner whatsoever without the express written approval of MRMC ACURO.

The grant recipient shall complete the ACURO Animal Use Appendix for Research Involving Animals found at the following web site:

http://mrmc.amedd.army.mil/index.cfm?pageid=research\_protections.acuro\_animalap\_pendix. Submit the completed ACURO appendix, contact information, the DTRA grant number and a copy of the grant for processing to the email address listed at the ACURO website. Once ACURO approves the effort, the grant recipient will receive written approval to begin animal use from the US Army MRMC ACURO by separate email. The grant recipient shall promptly provide a copy of the approval to the Grants Officer and Grants Officer representative. After approval, changes or protocol amendments must be submitted to and approved by ACURO before implementation.

The grant recipient, or subcontractors as appropriate, shall submit the most recent U.S. Department of Agriculture Animal Care Inspection Report annually in accordance with instructions provided.

Non-compliance with any provision of this clause may result in termination of the grant.

DoD Instruction 3216.01, dated September 13, 2010, provides policy and requirements for the use of animals in DoD-funded research based on Army Regulation 40-33. The DoD definition of animal is any live nonhuman vertebrate. All proposals that involve the use of animals must be in compliance with DoD Instruction 3216.01 and AR 40-33. DTRA requires that research using animals not begin or continue until the ACURO has reviewed and approved the proposed animal use. For animals, the provisions include rules on animal acquisition, transport, care, handling, and use in: (i) 9 CFR parts 1-4, Department of Agriculture rules that implement the Laboratory Animal Welfare Action of 1966 (U.S.C. 2131-2156); and (ii) the "Guide for the Care and Use of Laboratory Animals," National Institutes of Health Publication No. 86-23

**RESEARCH INVOLVING RECOMBINANT DNA MOLECULES.** Any recipient performing research involving recombinant DNA molecules and/or organisms and viruses containing recombinant DNA molecules agrees by acceptance of this award to comply with the National Institutes of Health "Guidelines for Research Involving Recombinant DNA Molecules," July 5, 1994 (59 FR34496) amended August 5, 1994 (59 FR40170) amended April 27, 1995 (60 FR 20726), or such later revision of those guidelines as may be published in the Federal Register.

**COMBATING TRAFFICKING IN PERSONS.** The recipient agrees to comply with the trafficking in persons requirement in Section 106(g) of the Trafficking Victims Protection Act of 2000 (TVPA), as amended (22 U.S.C. 7104(g)) as implemented by 2 CFR 175.

- 1) Provisions applicable to a recipient that is a private entity.
  - a. You as the recipient, your employees, sub-recipients under this award, and sub-recipients' employees may not—
    - Engage in severe forms of trafficking in persons during the period of time that the award is in effect;
    - Procure a commercial sex act during the period of time that the award is in effect; or
    - Use forced labor in the performance of the award or subawards under the award.
  - b. We as the Federal awarding agency may unilaterally terminate this award, without penalty, if you or a sub-recipient that is a private entity—
    - Is determined to have violated a prohibition in paragraph 1)a. of this award term; or
    - Has an employee who is determined by the agency official authorized to terminate the award to have violated a prohibition in paragraph 1)a. of this award term through conduct that is either
      - o Associated with performance under this award; or
      - Imputed to you or the sub-recipient using the standards and due process for imputing the conduct of an individual to an organization that are provided in 2 CFR Part 180, "OMB Guidelines to Agencies on Government-wide Debarment and Suspension (Non-procurement)," as implemented by our agency at 2 CFR Part 376.
- 2) Provision applicable to a recipient other than a private entity.
  - a. We as the Federal awarding agency may unilaterally terminate this award, without penalty, if a sub-recipient that is a private entity—
    - Is determined to have violated an applicable prohibition in paragraph 1)a. of this award term: or
    - Has an employee who is determined by the agency official authorized to terminate the award to have violated an applicable prohibition in paragraph 1)a. of this award term through conduct that is either
      - o Associated with performance under this award; or
      - Imputed to the sub-recipient using the standards and due process for imputing the conduct of an individual to an organization that are provided in 2 CFR Part 180, "OMB Guidelines to Agencies on Government-wide Debarment and Suspension (Non-procurement)," as implemented by our agency at 2 CFR Part 376.
- 3) Provisions applicable to any recipient.
  - a. You must inform us immediately of any information you receive from any source alleging a violation of a prohibition in paragraph 1)a. of this award term.
  - b. Our right to terminate unilaterally that is described in paragraph 1)b. or 2)a. of this Article:
    - Implements Section 106(g) of the TVPA, as amended (22 U.S.C. 7104(g)), and

- Is in addition to all other remedies for noncompliance that are available to us under this award.
- c. You must include the requirements of paragraph 1)a. of this award term in any subaward you make to a private entity.
- 4) Definitions. For purposes of this award term:
  - a. "Employee" means either:
    - An individual employed by you or a sub-recipient who is engaged in the performance of the project or program under this award; or
    - Another person engaged in the performance of the project or program under this award and not compensated by you including, but not limited to, a volunteer or individual whose services are contributed by a third party as an in-kind contribution toward cost sharing or matching requirements.
  - b. "Forced labor" means labor obtained by any of the following methods: the recruitment, harboring, transportation, provision, or obtaining of a person for labor or services, through the use of force, fraud, or coercion for the purpose of subjection to involuntary servitude, peonage, debt bondage, or slavery.
  - c. "Private entity":
    - Means any entity other than a State, local government, Indian tribe, or foreign public entity, as those terms are defined in 2 CFR 175.25.
    - Includes:
      - A non-profit organization, including any non-profit institution of higher education, hospital, or tribal organization other than one included in the definition of Indian tribe at 2 CFR 175.25(b).
      - o A for-profit organization.
  - d. "Severe forms of trafficking in persons," "commercial sex act," and "coercion" have the meanings given at Section 103 of the TVPA, as amended (22 U.S.C. 7102).

PROHIBITION ON USING FUNDS UNDER GRANTS AND COOPERATIVE AGREEMENTS WITH ENTITIES THAT REQUIRE CERTAIN INTERNAL CONFIDENTIALITY AGREEMENTS. The recipient agrees to comply with the requirements in section 743 of the Financial Services and General Government Appropriations Act, 2015 (Division E of the Consolidated and Further Continuing Appropriations Act, 2015, Pub. L. 113-235):

- 1) The recipient may not require its employees, contractors, or sub-recipients seeking to report fraud, waste, or abuse to sign or comply with internal confidentiality agreements or statements prohibiting or otherwise restricting them from lawfully reporting that waste, fraud, or abuse to a designated investigative or law enforcement representative of a Federal department or agency authorized to receive such information.
- 2) The recipient must notify its employees, contractors, or sub-recipients that the prohibitions and restrictions of any internal confidentiality agreements inconsistent with paragraph 1) of this award provision are no longer in effect.

- 3) The prohibition in paragraph 1) of this award provision does not contravene requirements applicable to Standard Form 312, Form 4414, or any other form issued by a Federal department or agency governing the nondisclosure of classified information.
- 4) If the Government determines that the recipient is not in compliance with this award provision, it:
  - a. Will prohibit the recipient's use of funds under this award, in accordance with section 743 of Division E of the Consolidated and Further Continuing Resolution Appropriations Act, 2015, (Pub. L. 113-235) or any successor provision of law; and
  - b. May pursue other remedies available for the recipient's material failure to comply with award terms and conditions.

# 14. Data Collection.

Data collection activities, if any, performed under this Grant are the responsibility of the recipient. Awarding agency support of the project does not constitute approval of the survey design, questionnaire content, or data collection procedures. The recipient shall not represent to respondents that such data are being collected for or in association with the awarding agency without the specific written approval of the cognizant awarding agency official. However, this requirement is not intended to preclude mention of the awarding agency support of the project in response to an inquiry or acknowledgment of such support in any publication of this data.

# 15. Publications and Acknowledgement of Sponsorship.

Publication of results of the research project in an appropriate professional journal is encouraged as an important method of recording and reporting scientific information. .

The recipient agrees that in the release of information relating to the grant, such release shall include the following statement, "The project or effort depicted was or is sponsored by the Department of the Defense, Defense Threat Reduction Agency. The content of the information does not necessarily reflect the position or the policy of the federal government, and no official endorsement should be inferred." For purposes of this provision, information includes news releases, articles, manuscripts, brochures, advertisements, still and motion pictures, speeches, trade association proceedings, symposia, etc.

When issuing statements, press releases, requests for proposals, bid solicitations, and other documents describing projects or programs funded in whole or in part with federal money, all recipients receiving federal funds, shall clearly state: (i) the percentage of total costs of the program or project which will be financed with federal money, and (ii) the dollar amount of federal funds for the project or program.

#### 16. Authorization to Perform Activities Abroad.

If the award recipient is a foreign institution, the recipient assures that it has been duly authorized to operate and do business in the country or countries in which the grant is to be performed; that it has obtained all appropriate licenses, permits, and approvals required in connection with the grant's proposed activities; and that it will fully comply with all the laws,

decrees, labor standards and regulations of such country or countries during the performance of the grant. U.S. Government funds may not be used in support of a project which is prohibited by law in the country or countries in which it is undertaken. DTRA does not assume responsibility for the recipient's compliance with the laws and regulations of the country or countries in which the activities are to be conducted.

# 17. Inconsistency between English Version and Translation of Grant.

The foreign recipient shall ensure that all contract correspondence that is addressed to the U.S. Government is submitted in English or with an English translation. In the event of inconsistency between the terms of the grant and any translation thereof into another language, the meaning in the English language shall control.

# 18. Value Added Tax (VAT) and Other Taxes

During implementation of grant activities, the recipient will notify DTRA as soon as they become aware of any VAT or other taxes, exceeding \$500.00 per transaction, not identified in the grant proposal and outside those considered VAT costs associated with travel, including and limited to lodging, meals, and transportation. A "transaction" is defined as a single purchase by the recipient and transactions may not be deliberately split in order to avoid compliance with the \$500.00 limit. DTRA approval in writing with documentation of extraordinary circumstances is required prior to the recipient using any DTRA funds for VAT or other taxes exceeding \$500.00 per transaction, and a grant modification may be required. The recipient understands that in the event that DTRA is unable to secure approval to use DTRA funds for VAT or other taxes exceeding \$500.00 per transaction, the purchase of applicable items may not proceed. For those instances where the recipient has received written approval to use DTRA funds to pay VAT or other taxes on any item(s) exceeding \$500.00 per transaction, the recipient will include this information in its financial reports (e.g., SF 425) to DTRA.

# 19. <u>Prohibition on the Use of Cooperative Threat Reduction Funds in the Russian</u> Federation

Recipients of grants or cooperative agreements in Thrust Area 6 are prohibited from using DTRA funds for activities in the Russian Federation.

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CONTRACTING OFFICER WILL CONTRACTORS NEGOTIATED AGREEMENT document and return—copies to issuing office.) Committens or perform all the services set foult or otherwise identifisheets for the consideration stated herein. The rights and oblig contract shall be subject to and governed by the following dociny they have solicitation, if any, and re) such provisions, represented as are attached or incorporated by reterence herein.  [Attachments are listed herein.]	IS.     SEALED-BID AWARD   (Contractor is not required to sign this document.)  Your bid on Solicitation Number   (Contractor is not required to sign this document.)  Including the additions or changes made by you which additions or changes are set forth in tull above, is hereby accepted to the terms listed above and on any continuation sheets. This award consummates the contract which consists of the following documents: (a) the Government's solicitation and your bid, and (b) this award-contract. No turther contractual document is necessary. (Block 18 should be checked only when awarding a scaled bid contract.)  20A. NAME OF CONTRACTING OFFICER					accepted as				
			(b)(6)		CONTRA	CTING OF	_	1/6)		_
19B. NAME OF CONTRACTOR	19C. DAT	E SIGNED		,0)(0) <u>UNITED ST.</u> b)(6)	ATES OF	AMERI	ekati.: <mark>(b)</mark> CA	)(O)	20C. DATE S	IGNED
BY			BY.		(Signan	ne al Contro	reting (Officer)		29-Sep-202	20

Section B - Supplies or Services and Prices

ITEM NO 0001	SUPPLIES/SERVICES	QUANTITY 1	UNIT Lot	UNIT PRICE \$2,957,164.19	AMOUNT \$2,957,164.19
	Years 1-3: FRBAA14-6-2	Ψ2,>-/,,1()/			
	FFP Reducing the Threat of M: Influenza in Jordan & Stre accordance with the follow and DTRA Terms and Cor attachment 2 FOB: Destination U009	ingthening Regiona ving attachements:	al Disease Sur SOW at attacl	veillance Capacity. In hment 1 dated 3/18/2020	
				NET AMT	\$2,957,164.19
ITEM NO	SUPPLIES/SERVICES	QUANTITY	UNIT	UNIT PRICE	AMOUNT
000101	Funding to support CLIN FFP	0001			\$0.00
				NET AMT	\$0.00
	ACRN AA CIN: HDTRA1036112000	)1			\$2,956,308.78

Page 3 of 10

\$942,857.34

ITEM NO SUPPLIES/SERVICES QUANTITY UNIT UNIT PRICE AMOUNT 0002 \$999,970.32 \$999,970.32 1 Lot OPTION Year 4: FRBAB14-6-2-0471 **FFP** Reducing the Threat of Middle East Respiratory Syndrome Coronavirus and Avian Influenza in Jordan & Strengthening Regional Disease Surveillance Capacity. In accordance with the following attachements: SOW at attachment 1 dated 3/18/2020 and DTRA Terms and Conditions for Grant Awards dated 09/14/2020 at attachment 2 FOB: Destination U009 NET AMT \$999,970.32 SUPPLIES/SERVICES **QUANTITY** UNIT UNIT PRICE AMOUNT ITEM NO 0003 l Lot \$942,857,34 \$942,857.34 OPTION Year 5: FRBAA14-6-2-0471 FFP Reducing the Threat of Middle East Respiratory Syndrome Coronavirus and Avian Influenza in Jordan & Strengthening Regional Disease Surveillance Capacity. In accordance with the following attachements: SOW at attachment 1 dated 3/18/2020 and DTRA Terms and Conditions for Grant Awards dated 09/14/2020 at attachment 2 FOB: Destination U009

NET AMT

Section C - Descriptions and Specifications

### CLAUSES INCORPORATED BY FULL TEXT

### 252.601-9002 GRANT REFERENCE INFORMATION (MAY 2009)

- a. This grant is awarded as a result of Broad Agency Announcement (BAA) **HDTRA1-11-16-BRCWMD-BAA**, Research and Development Enterprise, Basic and Applied Sciences Directorate, Basic Research for Combating Weapons of Mass Destruction (C-WMD).
- b. CFDA #: 12.351
- c. Authority: 10 U.S.C 2358 as amended.

(End of Clause)

Section E - Inspection and Acceptance

### INSPECTION AND ACCEPTANCE TERMS

### Supplies/services will be inspected/accepted at:

CLIN	INSPECT AT	INSPECT BY	ACCEPT AT	ACCEPT BY
0001	Destination	Government	Destination	Government
000101	N/A	N/A	N/A	N/A
0002	Destination	Government	Destination	Government
0003	Destination	Government	Destination	Government

### Section F - Deliveries or Performance

### DELIVERY INFORMATION

CLIN	DELIVERY DATE	QUANTITY	SHIP TO ADDRESS	DODAAC / CAGE
0001	POP 29-SEP-2020 TO 28-SEP-2023	N/A	DEFENSE THREAT REDUCTION AGENCY/CT-I (b)(6) 8725 JOHN J. KINGMAN ROAD MSC 6201 FORT BELVOIR VA 22060-6201 (b)(6) FOB: Destination	HDTRAI
000101	N/A	N/A	N/A	N/A
0002	1 yr. AOE	1	DEFENSE THREAT REDUCTION AGENCY/CT-I (b)(6) 8725 JOHN J. KINGMAN ROAD MSC 6201 FORT BELVOIR VA 22060-6201 (b)(6) FOB: Destination	HDTRA1
0003	1 уг. ЛОЕ	1	(SAME AS PREVIOUS LOCATION) FOB: Destination	HDTRA1

### ACCOUNTING AND APPROPRIATION DATA

AA: 044315 097 0134 000 N 20182020 D 34HQ 0901515BR\_KD\_BP\_OT\_18 1820\_0134\_34HQ\_SCNCT DTRA 410 AMOUNT: \$2.956,308.78

ACRN CLIN/SLIN CIN AMOUNT

000101 HDTRA10361120001 AA\$2,956,308.78

### CLAUSES INCORPORATED BY FULL TEXT

### 252.601-9000 GRANT POINTS OF CONTACT (MAY 2009)

Grant Specialist: a.

Name: (b)(6)

Defense Threat Reduction Agency/AL-ACC 8725 John J. Kingman Road, MS 6201

Fort Belvoir, VA 22060-6201

Telephone(b)(6)

Email address: (b)(6)

Grantee Business Office: b.

Name: Dr. Aleksei Chmura

Title: Executive Organizational Representative

Phone: (212) 380 - 4473

E-mail: chmura@ecohealthalliance.org

Grantee Principal Investigator (PI): c.

Name: Dr. William Karesh

Executive Vice President of Health and Policy Title:

Phone: (212) 380 - 4463

E-mail: karesh@ecohealthalliance.org

(End of Clause)

### GRANTS OFFICER'S REPRESENTATIVE (GOR) (MAY 2009) 252.601-9001

d. Grants Officer's Representative (GOR) for this Grant is:

Name: (b)(6) Defense Threat Reduction Agency/BTRP

8725 John J. Kingman Road

Fort Belvoir, VA 22060-6201 telephone(b)(6)

email address:

## b. SCOPE AND LIMITATIONS OF AUTHORITY FOR DTRA GRANTS OFFICER'S REPRESENTATIVE

- 1. The Grants Officer's Representative (GOR) is responsible for monitoring the technical and programmatic performance of the grant or cooperative agreement (hereinafter referred to as agreement) including compliance with the agreement's reporting requirements. Perceived deviations from the agreement's terms and conditions shall be brought to the attention of appropriate recipient personnel and if not corrected, to the attention of the Grants Officer. Deviations or other occurrences indicative of the recipient's inability or unwillingness to conform to the agreement's requirements shall be brought to the immediate attention of the Grants Officer.
- 2. The GOR has no authority to modify the stated terms of the agreement or specifications in any manner, or to approve any action that would result in additional charges to the Government. The DTRA Grants Officer must make all such changes in writing.
- 3. The GOR is authorized to correspond directly with the recipient on matters within the scope and limitations of his or her authority. All such correspondence shall be signed personally by the GOR and a copy will be furnished the Grants Officer for placement in the official agreement file.
- 4. The GOR will provide technical advice to the recipient to the extent such advice will not alter any of the agreement's terms and conditions or result in an increase in the estimated cost of the agreement's performance as specified in the agreement. The GOR is encouraged to discuss with the recipient new ideas related to the original scope of the agreement that may be of special interest to DTRA. The GOR should also be alert for indications that any part of the research is becoming unfruitful. If, as a result of such discussions, questions arise as to the possible need for a modification, the matter shall be brought to the immediate attention of the Grants Officer. The GOR shall not direct the recipient to undertake any action that the recipient believes to be contrary to the agreement's scope, terms or conditions without the written approval of the Grants Officer.
- 5. Any visits to the recipient's facilities on matters pertaining to the agreement will be documented in a brief memorandum. A copy will be provided the Grants Officer for placement in the official agreement file.
- 6. The GOR will maintain such work files as he or she deems necessary to properly document performance of the duties and responsibilities as a GOR. Upon completion of the agreement and delivery of all required reports, such work files will be forwarded to the Grants Officer for placement in the official agreement file.
- 7. In performance of the GOR's duties, the GOR shall constantly stress protection of the Government's interests. Similarly, the GOR shall avoid any act which may tend to compromise the position of DTRA, any individual member of DTRA or which will impact confidence in the integrity of DTRA with the business community.

(End of Clause)

### FUNDING PROFILE:

The amount of \$2,956,308.78 is obligated for work to be performed during the period beginning with grant award and continuing through September 28, 2023. Additional incremental funding planned, but not obligated, is:

FY 21 \$855.41

The Government's liability is limited to the amount obligated

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Invoice Schedule		
1	9/29/2020	\$79,900.24
2	10/29/2020	\$79,900.24
3	11/29/2020	\$79,900.24
4	12/29/2020	\$79,900.24
5	1/29/2021	\$79,900.24
6	2/28/2021	\$79,900.24
7	3/29/2021	\$79,900.24
8	4/29/2021	\$79,900.24
9	5/29/2021	\$79,900.24
10	6/29/2021	\$79,900.24
11	7/29/2021	\$79,900.24
12	8/29/2021	\$79,900.24
13	9/29/2021	\$79,900.24
14	10/29/2021	\$79,900.24
15	11/29/2021	\$79,900.24
16	12/29/2021	\$79,900.24
17	1/29/2022	\$79,900.24
18	2/28/2022	\$79,900.24
19	3/29/2022	\$79,900.24
20	4/29/2022	\$79,900.24
21	5/29/2022	\$79,900.24
22	6/29/2022	\$79,900.24
23	7/29/2022	\$79,900.24
24	8/29/2022	\$79,900.24
25	9/29/2022	\$79,900.24
26	10/29/2022	\$79,900.24
27	11/29/2022	\$79,900.24
28	12/29/2022	\$79,900.24
29	1/29/2023	\$79,900.24
30	2/28/2023	\$79,900.24
31	3/29/2023	\$79,900.24
32	4/29/2023	\$79,900.24
33	5/29/2023	\$79,900.24
34	6/29/2023	\$79,900.24
35	7/29/2023	\$79,900.24
36	8/29/2023	\$79,900.24
37	9/28/2023	\$79,900.14

(End of Clause)

### Section J - List of Documents, Exhibits and Other Attachments

### Exhibit/Attachment Table of Contents

DOCUMENT TYPE	DESCRIPTION	PAGES	DATE
Attachment 1	Statement of Work	7	18-MAR-2020
Attachment 2	Terms and Conditions	14	14-SEP-2020

**Project Title:** Reducing the Threat of Middle East Respiratory Syndrome Coronavirus and Avian Influenza in Jordan & Strengthening Regional Disease Surveillance Capacity.

Document Date: 18 March 2020.

Objective: The objective of this grant is to reduce the threat of high-consequence zoonotic pathogens in Jordan and improve regional disease surveillance capacity. Jordan faces risk of several zoonoses of concern to human and animal health, including Middle East Respiratory Syndrome (MERS-CoV) and Avian Influenza (AI), with fundamental knowledge and capacity gaps around the distribution and determinants of zoonoses. Recent work by our team detected MERS-CoV in camels and people in Jordan, suggesting ongoing and unmitigated transmission risk of a priority pathogen. The proposed study will generate critical advances in determining the presence of zoonotic pathogens in the country and opportunities for public health intervention. Through a prospective human cohort study coordinated with animal sampling, we will conduct biological and behavioral surveillance in five regions of Jordan with livestock production interfaces to determine the presence and risk factors for MERS-CoV, AI, brucellosis, and leptospirosis to identify modifiable risk factors. By testing three core policy-relevant hypotheses and providing multi-disciplinary training opportunities, the awardee shall enhance scientific capacity in Jordan and support disease detection and reporting in Jordan, Iraq and Lebanon. Taking a coordinated, multi-hazard approach to threat reduction, the proposed study will add critical understanding of presence and risk factors for zoonotic diseases in Jordan and advance scientific capacity and application of the One Health concept to counter biothreats in the region.

**Scope:** The awardee proposes a five-year One Health study of zoonotic diseases in Jordan. The awardee team shall focus on the following major goals and milestones:

- 1) Determine the presence of MERS-CoV, AI, and other zoonoses among livestock and poultry in Jordan: *Implement animal study (YI-OYI)*; *Conduct PCR and serology testing (YI-OY2)*
- 2) Determine whether (and if so, the extent to which) humans at human-livestock/poultry interfaces display evidence of infection with these zoonoses: *Implement human cohort study (Y1-OY1)*; Conduct PCR and serology testing (Y1-OY2)
- 3) Characterize causal factors in animal-to-human transmission of these zoonoses: *Implement behavioral risk survey (Y1-OY1)*; *Conduct epidemiological analyses (Y1-OY2)*
- 4) Produce geospatial maps of zoonoses in livestock and poultry and of human zoonotic transmission risk: *Generate geospatial distribution and risk maps (Y1-OY2)*
- 5) Strengthen local capacity for detection and reporting of MERS-CoV, AI, and other zoonoses in Jordan, Iraq, and Lebanon: *Host workshops (YI-2, OYI-2); Submit reports and publications (YI-OY2); Attend presentations/meetings (YI-OY2)*
- 6) Identify specific modifiable risk factors for human infection with these zoonoses: Conduct causal interference analyses (Y3-OY2); Policy recommendations on modifiable risk factors (Y3-OY2)

This research will generate critical advances in detecting the presence of and exposure risk to high-consequence zoonotic viruses and bacteria across geographic regions and interfaces in Jordan, contributing to enhanced understanding of modifiable risk factors to inform government authorities on prevention activities to reduce disease threats.

Background: Jordan has experienced outbreaks of several high-consequence zoonotic diseases, including MERS-CoV and AI. Since its emergence in 2012, MERS-CoV has resulted in over 2,400 human cases globally and is recognized as a priority disease under the WHO R&D Blueprint. While the first human cases of MERS-CoV were traced back to Jordan, little is known about its underlying and ongoing risk of zoonotic disease in animal populations, particularly around primary transmission risk factors and pathways that precede spread in healthcare settings. MERS-CoV is one of several zoonoses posing threat to animal and human populations in the country. Highly Pathogenic Avian Influenza outbreaks, including subtype H5N1, have been reported, and brucellosis is considered endemic in Jordan, with recurring spillover but inadequate understanding of specific modes of transmission and poor vaccination coverage. While human cases of leptospirosis have not been reported in Jordan to date, high rates of certain Leptospira serovars have been detected in animals. Key questions and capacity gaps hinder understanding of the presence, distribution and risk factors in the country, leaving the country vulnerable to zoonotic biothreats, as well as wider regional gaps in detection and reporting. The threat of zoonotic disease is especially pertinent given the rapidly-growing poultry production industry in

Jordan and camel, livestock, and poultry rearing in the region.

Jordan has recently made recent several notable scientific advances in detection of MERS-CoV, generating preliminary data that the proposed project will build on. While camels are the presumptive source of primary human MERS-CoV infections, the exact mechanisms of transmission and the possible role of other livestock species are unclear. Blood samples collected from camels and humans in the northern region of Jordan were positive for MERS-CoV, leading to the first-ever report of this disease to OIE in camels in Jordan in 2016. To date, only few countries have reported virus-positive MERS-CoV test results to the OIE so this is a significant and important step toward improving both MERS-CoV detection and reporting in the Middle East, Studies of zoonotic pathogens in the region to date have largely focused on single sites or interfaces, cross-sectional sampling events, or select taxonomic groups, limiting understanding of causal factors. Research is needed to monitor presence and transmission of zoonotic pathogens spatially, temporally, and by biohazard exposure (e.g. blood, urine, feces, and/or nasal secretions) and practices and conduct epidemiological analyses to identify modifiable risk factors for public health intervention. As a key area of stability in the region, it is crucial to support Jordan's biosurveillance capacity, enable understanding of baseline disease risk to allow differentiation of natural versus nefarious emergence events, and ensure responsible bio-risk management to ethically monitor and reduce disease threats. The country has also recently established a One Health platform that indicates the country's commitment to countering zoonotic disease threats. Given the volume of migration between Jordan and its neighbors, including in animal trade, regional coordination in disease monitoring and threat reduction is a critical component in effectively characterizing and addressing disease risk. This project seeks to fill current capacity gaps to advance Jordan as a leader in scientific research and zoonotic disease management and creating new capacity and scientific collaboration pathways for hypothesisdriven research and zoonotic disease monitoring and threat reduction in the region.

Key references include (additional references can be found in the Project Narrative): van Doremalen N, Hijazeen ZS, Holloway P, Al Omari B, McDowell C, Adney D, Talafha HA, Guitian J, Steel J, Amarin N, Tibbo M, Abu-Basha E, Al-Majali AM, Munster VJ, Richt JA. High Prevalence of Middle East Respiratory Coronavirus in Young Dromedary Camels in Jordan. *Vector Borne Zoonotic Dis.* 2017 Feb;17(2):155-159.

Alagaili AN, Briese T, Mishra N, Kapoor V, Sameroff SC, Burbelo PD, de Wit E, Munster VJ, Hensley LE, Zalmout IS, Kapoor A, Epstein JH, Karesh WB, Daszak P, Mohammed OB, Lipkin WI. Middle East respiratory syndrome coronavirus infection in dromedary camels in Saudi Arabia. *MBio*. 2014 Feb 25;5(2):e00884-14.

World Health Organization. Joint External Evaluation of IHR Core Capacities of the Hashemite Kingdom of Jordan. Geneva. 2016.

Tasks/Scientific Goals: (Format: Year #(s), Task #. Subtask#).

# TASK 1: Y1.1-O2.1: Determine the presence of MERS-CoV, AI, and other zoonoses among livestock and/or poultry in Jordan.

The awardee shall study presence of MERS-CoV, AI, Brucellosis and Leptospirosis in animals. Sites will be selected in five geographically-representative regions across Jordan: Northern Jordan (Al Ramtha), Middle Jordan (Al Zarga), and Southern Jordan (Al Karak, Ma'an, and Aqaba). These regions were selected based on preliminary findings by our team and the presence of livestock production activities. Sites in each region shall reflect interfaces with poultry, camels, and other livestock animals are represented (e.g. farms or markets). The awardee shall receive permissions and approvals to work with animals prior to initiating sampling. Nasopharyngeal and oropharyngeal swabs will be collected from 300 camels, poultry and other livestock quarterly in Year 1, Q3 through Option Year 1, Q3 (a total of 13 sampling visits). At three points during the project (in Y1, Y3, and OY1 during the 1st, 7th, and 13th sampling visits, respectively), blood samples (serum) will be collected from 300 animals. Nasopharyngeal and oropharyngeal swabs will be tested for MERS-CoV and AI using real time PCR. Serum samples will be tested using serology for MERS-CoV, AI, Brucellosis, and Leptospirosis. Staff training on proper study techniques (e.g. sampling, transport, and laboratory) will occur as detailed in Task 5. This task will contribute to testing the hypothesis that livestock species and poultry in Jordan show evidence of infection with MERS-CoV, AI, and/or other assayed zoonoses, as well as inform type of transmission pathways.

- Y1.2.1 Identify sites for project study.
- Y1.2.2 Obtain local permissions and approvals to work with animals.
- Y1.2.3-O1.2.3 Conduct biological sampling in animals within the study area.
- Y1.2.4-O2.2.4 Conduct PCR and/or serology testing for AI, MERS-CoV, brucellosis and leptospirosis.

# TASK 2: Y1.2- OY2.1: Determine whether (and if so, the extent to which) humans at human-livestock/poultry interfaces display evidence of infection with these zoonoses.

The awardee shall conduct a prospective cohort study of humans to evaluate behavioral and occupational risk factors for zoonotic infectious diseases (MERS-CoV, avian influenza, brucellosis, and leptospirosis) among persons living in Jordan in any of the five study regions. The study will enroll persons regularly working with livestock or poultry or sharing their living areas with these animals as well as persons unexposed to these factors. Sites will be selected as in Task 1 and in surrounding communities to enroll both exposed and unexposed populations. In Year 1, the awardee will receive local permissions and approvals to work with human subjects prior to initiating enrollment and sampling. Nasopharyngeal and oropharyngeal swabs will be collected from 300 enrolled humans quarterly in Year 1, Q3 through Option Year 1, Q3 (a total of 13 sampling visits). At three points during the project (in Year 1, Year 3, and Option Year 1

Reducing the Threat of Middle East Respiratory Syndrome Coronavirus and Avian Influenza in Jordan & Strengthening Regional Disease Surveillance Capacity/PI Karesh

Thrust Area 6 - CCWMD Research with Global Partners FRCWMD

during the 1<sub>st</sub>, 7th, and 13th sampling visits, respectively), blood samples (serum) will be collected from 300 humans. Nasopharyngeal and oropharyngeal swabs will be tested for MERS-CoV and AI using real time PCR. Serum samples will be tested using serology for MERS-CoV, AI, brucellosis, and leptospirosis (Years 1-4). Biological sampling will be paired with behavioral surveillance data as in Task 3.

- Y1.2.1 Identify sites and conduct enrollment for human study component.
- Y1.2.2 Obtain local permissions and approvals to work with human subjects.
- Y1.2.3-O1.2.3 Conduct biological sampling (nasopharyngeal and oropharyngeal swabs, and/or blood) of in people within the study area.
- Y1.2.4-O2.2.4 Conduct lab testing for AI, MERS-CoV, brucellosis and leptospirosis.

# TASK 3: Y1.3-OY2.3: Characterize causal factors in animal-to-human transmission of these zoonoses.

The awardee shall monitor behaviors of persons enrolled in the prospective cohort study (as in Task 2) to identify and characterize causal factors for animal-to-human transmission of MERS-CoV, AI, brucellosis and leptospirosis. The PREDICT-2 Human Behavioral Risk Questionnaire will be augmented with animal-specific exposure frequency questions to collect demographic data, symptom and medical history data, social history data, and specific animal-related behaviors and practices that are possible risk factors for infection with MERS-CoV, avian influenza, brucellosis, or leptospirosis. The project team will administer the survey to enrolled participants at the first sampling visit (Year 1), During all future visits (Year 1- Option Year 1), enrolled participants will be administered a brief follow-up questionnaire designed primarily to capture time-varying exposure and covariate data. This longitudinal information will provide critical information about the context of exposures and pathways to link biological sampling and provide the wider context of practices and risk factors. Epidemiologic analysis of the questionnaire data (including time-varying data) in this Task will identify current practices and exposure pathways and initial modifiable risk factors. Specific animal-related behaviors and practices that involve exposure to specific animal biohazards (e.g., blood, urine, feces, and/or nasal secretions) will be assessed under Task 6 for whether they pose a greater risk of infection with MERS-CoV, avian influenza, brucellosis, or leptospirosis. Results will be shared via annual reports and stakeholder meetings as in Task 5.

- Y1.3.1 Obtain local permissions and approvals to work with human subjects.
- Y1.3.2-O1.3.2 Conduct behavioral risk factor surveys in people.
- Y1.3.3-O2.3.3 Analyze epidemiologic causal inference to identify modifiable risk factors.

# TASK 4: Y1.4-OY2.4: Produce geospatial maps of zoonoses in livestock and poultry and of human zoonotic transmission risk.

The awardee shall map findings from Tasks 1-2 to show the distribution of detected pathogens across the regions and sampling sites. Geospatial mapping is a highly relevant visual tool to assist authorities in targeting surveillance and risk management activities. Using QGIS, laboratory results of human and animal testing for MERS-CoV, avian influenza, brucellosis, and leptospirosis will be populated and mapped to show the distribution of detected pathogens. Maps will be generated beginning Y1, Q3 when laboratory findings are available. Geospatial risk maps will be generated using statistical models to link human behavioral data to laboratory findings beginning in Y1, Q4. Geospatial maps will be updated quarterly and shared with USG/DTRA and Jordanian project partners. Training on geospatial analysis will be provided to project team

members and will be covered under a regional workshop to promote broader uptake of geospatial mapping as a low-cost tool to enhance disease monitoring programs in the region (Task 5).

Y1.4.1-O2.4.1 Generate maps of laboratory results

Y2.4.2-O2.4.2 Generate risk maps of human behavioral risk data

# TASK 5: Y1.5-OY2.5: Strengthen local capacity for detection and reporting of MERS-CoV, AI, and other zoonoses in Jordan, Iraq, and Lebanon through meetings and workshops.

Strengthening disease detection and reporting capacity is a key contributor to health security and the ability to target risk factors for threat reduction. The project team is committed to improving this capacity in Jordan as well as more widely in the region through involvement of scientists and government officials in Iraq and Lebanon. We will train local staff in proper techniques for all project activities (Year 1 and Option Year 1). Each year we will support four students from medical and veterinary disciplines in Jordan to participate in training, field activities, project analyses and workshops to provide scientific mentorship and promote the One Health concept. Jordanian scientists and students will also be trained in laboratory techniques to be deployed in Jordan by the Human Link laboratory consultant that has led capacity development for MERS-CoV diagnostics in other countries. The awardee shall organize workshops in Jordan open to scientists from the Ministries of Health and Agriculture from Jordan, Iraq, and Lebanon (Years 1-2, Option Years 1-2). A secure project database will be developed for use in the epidemiological analyses. The awardee shall provide submission of annual sample repository information using a DTRA-specified format and shall grant access to all samples collected and data generated during the course of the project, up to and including at least 12 months after the project end date. The awardee shall conduct presentations/meetings at times and places specified in the grant schedule (Year 1 Task 5), including the DTRA Annual Technical Review. In Year 1 the awardee shall hold a kick-off meeting to introduce the project objectives and promote buy-in in study findings and capacity sustainment. Annual reports and stakeholder meetings and scientific presentations and publications will be used to disseminate results. The project will reinforce Jordan's national One Health platform, promoting the One Health concept for multiple diseases of concern for public and animal health and advancing capacity for hypothesis-driven research with direct application for policy decisions for threat reduction.

Y1.5.1 Conduct project kick-off meeting in Amman with local stakeholders.

Y1.5.2 & O1.5.2 Train local project staff in proper techniques.

Y1.5.3-O2.5.3 Host annual partners and stakeholders meeting.

Y1.5.4-O2.5.4 Complete annual report and share with project partners and local stakeholders.

Y1.5.5-Y2.5.5 & O1.5.5-O2.5.5 Conduct local/regional training workshops.

Y1.5.6- O2.5.6 Data management.

Y1.5.7- O2.5.7 Conduct presentations/meetings at times and places specified in the grant schedule, including DTRA Annual Technical Review.

# TASK 6: Y3.6-OY2.6: Identify specific modifiable risk factors for human infection with these zoonoses.

The awardee shall integrate the findings of the biological testing (animal and human) and behavioral surveys in Tasks 1-3, assessing specific animal-related behaviors and practices that involve exposure to specific animal biohazards (e.g., blood, urine, feces, and/or nasal secretions)

for whether they pose a greater risk of infection with MERS-CoV, avian influenza, brucellosis, or leptospirosis. These potential modifiable risk factors will be determined through epidemiologic analysis of the questionnaire data (including time-varying data) paired with individual laboratory results (e.g., detection of viral RNA, bacterial DNA, or antibodies). Using quarterly, longitudinal data collected in Tasks 1-3, we will examine any major temporal dynamics that may be associated with elevated transmission risk (e.g. seasonality, animal birthing periods). Causal inference techniques within the potential outcomes framework will be employed to solidify statistical associations as concretized biological/clinical/environmental pathways by operationalizing exposures as well-defined interventions and ensuring exchangeability is maintained through confounder control informed by directed acyclic graphs of the conceptual exposure-disease pathways. The awardee will examine multiple zoonotic pathogens (as above) to inform general threat reduction guidelines. Identification of modifiable risk factors will inform intervention development that may interrupt future zoonotic transmission of the four pathogens in this study. Findings will be shared with USG/DTRA and the Jordanian Ministry partners at the annual stakeholders meetings, the threat reduction workshop, scientific presentations and publications as specified in Task 5. This information will provide a strong basis for potential policy guidance and the generation of solutions at the threat reduction workshop in OY2.

Y3.6.1-O2.6.1 Conduct causal inference analyses bases on Tasks 1, 2, 3. Y3.6.2-O2.6.2 Share results of analyses at annual partners and stakeholders meeting. Y3.6.3 & O2.6.3 Submit written report to DTRA and local partners and stakeholders.

### Performance Schedule:

Task	Year 1	Year 2	Year 3	OY 1	OY 2
Task 1: Determine the presence of MERS-CoV, AI, and other zoonoses a	mong livestoc	k and/or poul	try in Jordan		
1.1. Identify sites for project study					
1.2 Obtain local permissions and approvals to work with animals					
1.3 Conduct biological sampling in animals within the study area					
1.4 Conduct lab testing for A1. MERS-CoV, brucellosis and leptospirosis					
Task 2: Determine whether (and if so, the extent to which) humans at hu	man-livestock	/poultry inter	faces display e	vidence of inf	ection
2.1 Identify sites and conduct enrollment for human study component					
2.2 Obtain local permissions and approvals to work with human subjects					
2.3 Conduct biological sampling					
2.4 Conduct testing for AI, MERS-CoV, brucellosis and leptospirosis					
Task 3: Characterize causal factors in animal-to-human transmission of t	hese zoonoses	:			
3.1 Obtain local permissions and approvals to work with human subjects					
3.2 Conduct behavioral risk factor surveys in people					
3.3 Analyze epi causal inference to identify modifiable risk factors					
Task 4: Produce geospatial maps of zoonoses in livestock and poultry and	of human zo	onotic transm	ission risk		
4.1 Generate maps of laboratory results					
4.2 Generate risk maps with human beh, risk data					
Task 5: Strengthen local capacity for detection and reporting of MERS-C	oV, AI, and o	ther zoonoses	in Jordan, Ir	aq, and Lebar	101
5.1 Conduct project kick-off meeting in Amman with local stakeholders					
5.2 Train local project staff in proper techniques					
5.3 Host annual partners and stakeholders meeting					
5.4 Complete annual report and share with partners and stakeholders					
5.5 Conduct local/regional training workshops					
5.6 Data management					
5.7 Conduct presentations/meetings at times and places specified					
Task 6: Identify specific modifiable risk factors for human infection with	these zoonose	S			
6.1 Conduct causal inference analyses based on Task 1, 2, 3					
6.2 Share results of analyses at annual partners and stakeholders meeting					
6.3 Submit written report to DTRA and local partners and stakeholders					

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Traveler 3 - Catherine Machalaba (Animan)		l.		ļ.		ļ.		ļ.		⊢	
Hotel Max Por Diem   \$	1,998.00	15	1,968.03	5	1,968.03	3	1,968.03	5	1,968.00	⊢	
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Gloves	50	\$ 29.65	7	5	207.55	14	5	415.10	0 :	š -	6	\$ 177.90	0	\$	
Masks N95	50	\$ 47.75	7	\$	334.25	7 :	\$	334.25	7 :	334.25	5	\$ 238.75	0	\$	
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HDTRA1-14-24-FRCWMD-BAA
CBEP = Thrust Area 6 - Cooperative Counter WMD Research
with Global Partners FRCWMD

"Reducing the Threat of Middle East Respiratory Syndrome Coronavirus and Avian Influenza in Jordan & Strengthening Regional Disease Surveillance Capacity"

Abstract: In line with Thrust Area 6 - Cooperative Counter WMD Research with Global Partners FRCWMD, we propose a multi-disciplinary One Health research project to identify causal factors in the transmission of Middle East Respiratory Syndrome Coronavirus (MERS), avian influenza (AI), and other zoonoses from animals to humans in Jordan, while simultaneously strengthening local capacity for prevention, detection, and reporting of these zoonoses in Jordan, Iraq, and Lebanon. Little is known about the transmission routes of MERS, AI, and other key zoonoses at human-livestock/poultry interfaces in Jordan. To fill this gap and contribute to biological threat reduction, we propose a hypothesis-driven One Health research project to characterize exposure risks for MERS and AI in Jordan using epidemiological causal inference techniques and improving serological technology. This prospective cohort study involves interviews and longitudinal sampling of humans and opportunistic non-lethal sampling of livestock and poultry at animal markets, farms, and abattoirs for MERS, Brucella spp., and Leptospira spp. (humans/livestock) and for AI subtypes (humans/poultry). Partner laboratories in Jordan will screen and characterize these zoonotic pathogens, and capacity-building crosstrainings will be conducted with visiting laboratory scientists from Iraq and Lebanon. This multifaceted One Health approach will advance scientific knowledge on the risk of MERS, AI, and other zoonoses in Jordan, strengthen local zoonotic disease surveillance capabilities, and enhance scientific partnerships in Jordan, Iraq and Lebanon to prevent, detect, and report MERS, AI, and other zoonotic threats.

Background: There have been over 2,200 cases and 790 deaths across 27 countries from MERS since 2012. Although most cases occurred in the Kingdom of Saudi Arabia (KSA), the first known human infections occurred in Jordan. Most KSA cases have been healthcare-associated, stemming from secondary generations of transmission. Identification of factors involved in *primary* transmission may inform interventions to prevent future primary cases and therefore prevent them from seeding healthcare or community outbreaks. Camels are the presumptive source of primary human MERS infections; however, the exact mechanisms of transmission and the possible role of other livestock species are unclear. These uncertainties highlight the need to better understand the causal factors of primary human infection with MERS at human-livestock interfaces. Understanding modifiable risk factors may lead to development of public health interventions that can prevent animal-to-human transmission of zoonoses.

Avian influenza viruses circulate worldwide among migratory bird populations and occasionally spill into poultry at mass production sites, farms, or backyard coops. Although not every AI subtype is capable of infecting humans or causing severe illness, there have been worrisome outbreaks. AI A/H5N1 has been detected in humans in 16 countries since 2003, resulting in 860 cases and 454 deaths by July 2018. Of these, nearly 42% were in neighboring Egypt. Due to antigenic shifts typical of the influenza viral genome, there is always a possibility of new AI subtypes emerging capable of infecting humans. In Jordan, the rapidly expanding poultry industry driven by ever-increasing demand poses a distinct global health security risk.

Identification and characterization of AI subtypes circulating among poultry across Jordan will provide critical information needed to understand baseline activity and risks. If human AI infections are detected, the project could identify causal factors in poultry-to-human influenza transmission, and therefore inform interventions to prevent future primary cases. Given the human AI infections in neighboring countries, this is a critical gap to be filled in Jordan.

Other zoonoses, such as *Brucella* spp. and *Leptospira* spp., also cause morbidity in Jordan, and surveillance for them can be easily integrated into the research study as secondary outcomes. All samples may be made available to other DTRA- or USG-funded researchers.

Scientific Impact for C-WMD Science: Given that transmission of high-consequence zoonotic pathogens such as MERS and AI, whether by nature or ill intent, poses a significant threat to global security, existing gaps in biosurveillance/detection capabilities, prevention/response activities, and reporting weakens our ability to counter biological WMD threats. Over 60% of emerging infectious diseases originate in animals, so it is imperative to strengthen local capacity to conduct biosurveillance for zoonotic disease threats to understand existing baselines of transmission, characterize risk factors to inform prevention activities that can interrupt transmission, and preempt and/or rapidly respond to emerging disease threats.

Our primary objectives are to characterize causal factors in animal-to-human transmission of MERS, AI, and other zoonoses in Jordan and to strengthen regional capacity for detection of these pathogens in order to reduce the threat of infectious diseases. Our proposed project will rigorously test the following initial hypotheses:

**H<sub>1</sub>.** Livestock species and poultry in Jordan show evidence of infection with MERS, AI, and/or other assayed zoonoses. (Output: Geospatial maps of naturally-occurring zoonotic viruses and bacteria capable of infecting humans developed with and shared with government authorities)

**H<sub>2</sub>.** Humans occupationally exposed to livestock and/or poultry have a greater risk of infection with MERS, AI, and/or other assayed zoonoses compared to other community members in Jordan. (Output: Geospatial risk maps developed with and shared with government authorities)

H<sub>3</sub>. Some specific occupational practices that promote exposure to specific animal biohazards (e.g., blood, urine, feces, and/or nasal secretions) pose a greater risk of infection with MERS, AI, and other assayed zoonoses compared to others. (Output: Enhanced understanding of modifiable risk factors to inform government authorities on prevention activities)

Knowledge generated from this project will confirm the presence of MERS, AI, and other zoonoses at human-livestock/poultry interfaces in Jordan, characterize specific occupational practices as causal factors in human infection with these zoonoses, and identify modifiable risk factors that may be used to inform prevention activities. The project will build and boost local capacity for prevention, detection, and reporting of zoonotic disease threats in Jordan, Iraq and Lebanon through regional cross-training workshops held in Jordan, all of which have substantial camel and poultry production (Fig. 1). This will fill capacity gaps, strengthen collaborative relationships, and foster goodwill in a politically fragile region, enhancing our ability to combat biological threats.

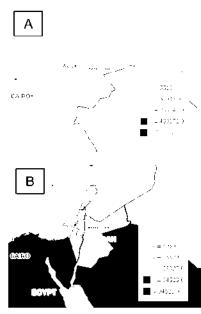


Figure 1: Regional map of live camel (A) and poultry (B) production, 2015.

**Methodology:** We propose to conduct a prospective cohort study with repeated surveys at various human-livestock/poultry interfaces across Northern, Middle, and Southern Jordan. Each interface will be visited thrice a year for 3 years (9 total visits per site) or 4 and 12 respectively if option years are exercised.

During each human-livestock interface visit, livestock (e.g., camels, goats, sheep, cattle) will be opportunistically tested for evidence of active infection with MERS using nasal swabs. During the first, fifth, and final visits, livestock will be opportunistically tested for past infection with MERS, *Brucella* spp., and *Leptospira* spp. using sera. Swab samples will be pooled by species + site + visit date and tested via real-time PCR. Serum samples will be tested via Luminex. During each human-poultry interface visit, poultry will be sampled for AI using oropharyngeal swabs. During the first, fifth, and final visits, poultry will be opportunistically tested for past infection with AI using sera.

Individuals at each site and its surrounding community will be approached for enrollment at the first visit. Those providing informed consent will complete a standardized questionnaire (the previously Jordan-approved and tested PREDICT-2 questionnaire augmented with study-specific items) during the first visit and complete a shorter questionnaire at each subsequent visit to account for time-varying factors. Participants will provide oropharyngeal and nasopharyngeal swabs at each visit and sera at the first, fifth, and final visits. These samples will be tested using PCR for the swab samples (MERS and AI) and Luminex serology for the serum samples (all 4 zoonoses).

Data will be analyzed to generate geospatial maps of livestock and poultry zoonoses, geospatial maps of human zoonotic disease risk (including static and seasonal versions), exposure risk models using causal inference techniques, and prediction models for transmission that can account for seasonal variations and wide-scale livestock/poultry practice trends.

Implementation of Luminex Scrological Technology: Luminex technology for serology and associated equipment will be acquired for participating laboratories in Jordan. Training workshops will be held in Jordan for regional scientists, with refresher trainings held annually.

Regional Capacity Building: Iraqi and Lebanese scientists from their Ministries of Health and Agriculture will be invited to participate in capacity-building workshops in Jordan. Workshops will cover topics including implementing/using Luminex technology, animal and human sampling fieldwork, ethical human subjects research, biosafety and biosecurity best practices, data management and storage, and reporting procedures pursuant to OIE, WHO, IHR, and GHSA obligations. Advance technical training for key individuals is proposed for option years.

Our research project will serve multiple goals and objectives of the CBEP mission, by: 1) engaging partner country scientists in hypothesis-driven research; 2) supporting biosurveillance capacity building by enhancing partner capability to detect and report select agents; 3) enhancing

understanding of zoonotic diseases to allow differentiation of natural versus nefarious emergence events; 4) employing responsible bio-risk management best practices; 5) training partner country researchers to think critically about ethical research and be competitive in soliciting international funding; and 6) promoting a One Health concept.

### **Core Team and Project Organization:**

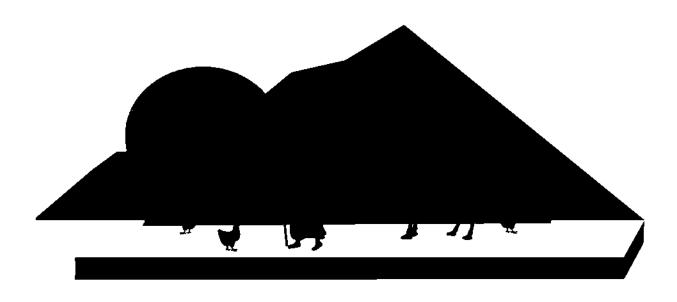
- EcoHealth Alliance (EHA, prime), led by Dr. William Karesh (PI) and Dr. Patrick Dawson (co-PI), will design, implement, and oversee the research project; lead all contractual obligations with DTRA and project sub-awardees; coordinate travel and organize workshops; manage the information database; ensure coordination between partners; and collaborate with our partners in analyzing the data and publishing it.
- Jordan University of Science and Technology (JUST), led by Dr. Ehab Abu-Basha (co-PI), will conduct fieldwork and laboratory work on the research study. EHA has worked extensively with JUST and Dr. Abu-Basha on the USAID PREDICT-2 project, conducting viral surveillance among humans and animals and risk characterization since 2016.
- Partner Laboratories: JUST (Irbid, Jordan) and Jordan Ministry of Agriculture (Amman, Jordan). These institutions have demonstrated capacities for all required laboratory work, training workshops, and specimen storage. Additional laboratory partners may be added following consultation with national partners and DTRA.

Potential Challenges, Solutions, and Statistical Considerations: The main potential challenge will be a low detection rate of MERS, AI, and other zoonoses among human participants. However, power calculations to determine an appropriate sample size will be informed using recent, credible findings from the PREDICT-2 project. Another potential challenge is addressing cultural sensitivities while conducting work, such as avoiding stigmatization of livestock workers or working with various Bedouin communities. EHA and JUST have extensive experience in this domain through PREDICT-2, and have fostered positive relationships with community leaders and liaisons throughout Jordan. To reduce regional instability risk, we will be work in Jordan, a stable bedrock in the region, and will invite regional scientists to workshops inside Jordan rather than traveling to them.

Major Goals and Milestones: 1) Determine the presence of MERS, AI, and other zoonoses among livestock and/or poultry in Jordan. 2) Determine whether (and if so, the extent to which) humans at human-livestock/poultry interfaces display evidence of infection with these zoonoses. 3) Characterize causal factors in animal-to-human transmission of these zoonoses. 4) Produce geospatial maps of zoonoses in livestock and poultry and of human zoonotic transmission risk. 5) Identify specific modifiable risk factors for human infection with these zoonoses. 6) Implement Luminex serological technology in Jordan for enhanced diagnostic and detection capabilities. 7) Strengthen local capacity for detection and reporting of MERS, AI, and other zoonoses in Jordan, Iraq, and Lebanon through workshops.

### **Proposed Budget (in US\$)**

Organization	Year 1	Year 2	Year 3	OY1	OY2
EcoHealth Alliance (Prime)	\$492,000	\$495,000	\$498,000	\$490,000	\$445,000
JUST	\$487,000	\$493,000	\$498,000	\$500,000	\$430,000
TOTALS	\$979,000	\$988,000	\$996,000	\$990,000	\$875,000



# Reducing the Threat of Middle East Respiratory Syndrome Coronavirus and Avian Influenza in Jordan & Strengthening Regional Disease Surveillance Capacity

### **Annual Report Year 1**

**Principal Investigator**: Dr. William Karesh, DVM

EcoHealth Alliance



Grant/Award #: HDTRA12010029 PI Name: Dr. William B. Karesh

Organization/Institution: EcoHealth Alliance

Project Title: Reducing the Threat of Middle East Respiratory Syndrome Coronavirus and

Avian Influenza in Jordan & Strengthening Disease Surveillance Capacity

Goal number, Milestone	Proposed completion
1, Implement animal study	Y1-OY1
1, Conduct PCR and serology testing	Y1-OY2
2, Implement human cohort study	Y1-OY1
2, Conduct PCR and serology testing	Y1-OY2
3, Implement behavioral risk survey	Y1-OY1
3, Conduct epidemiological analyses	Y1-OY2
4, Generate geospatial distribution and risk maps	Y1-OY2
5, Host workshops	Y1-2, OY1-2
5, Submit reports and publications	Y1-OY2
5, Attend presentations/meetings	Y1-OY2
6, Conduct causal interference analyses	Y3-OY2
6, Policy recommendations on modifiable risk	Y3-OY2
factors	

### **Project Goals**

- 1. Determine the presence of MERS-CoV, Al. and other zoonoses among livestock and poultry in Jordan
  - a. Description: Implement animal study (Y1-OY1); Conduct PCR and serology testing (Y1-OY2)
  - b. Status of Effort: Approval for animal research has been obtained from the Jordan University of Science and Technology (JUST) and Tufts University, and surveillance determination has been obtained from ACURO. Obtaining these approvals included achieving many milestones, including drafting and finalizing procedures, and the animal owner consent to sample. The animal field and laboratory teams are finalizing SOPs for data collection and testing.
- 2. Determine whether (and if so, the extent to which) humans at human-livestock/poultry interfaces display evidence of infection with these zoonoses
  - a. Description: Implement human cohort study (Y1-OY1); Conduct PCR and serology testing (Y1-OY2)
  - b. Status of Effort: Approval for human subjects research has been obtained from JUST (in-country approval) and Health Media Lab Research and Ethics (HML) IRB (U.S.-based approval), and we are awaiting the determination for human subjects research approval from HRPO. The data collection tools have been finalized including the surveys, code of surveys into Open Data Kit (ODK) for use on tablets, and the data management system in Airtable for laboratory data.

- Characterize causal factors in animal-to-human transmission of these zoonoses
  - a. Description: Implement behavioral risk survey (Y1-OY1); Conduct epidemiological analyses (Y1-OY2)
  - b. Status of Effort: Final approval for data collection is pending from HRPO, but data collection tools have been finalized. To develop the final behavioral risk survey, several milestones were met, including collaboratively determining the data analysis plan and the creation of the survey guide. Our causal inference approach necessitated a literature review of confounders and pathways, construction of causal diagrams, and a collaborative discussion amongst team members to determine which human behavioral risk questions would be important variables in our analysis (the data analysis plan). Once consensus was achieved on variables to include in our models, the team collaboratively determined the appropriate formulation of questions, the order of questions, and team responses to typical potential participant questions (addressed in the survey guide).
- 4. Produce geospatial maps of zoonoses in livestock and poultry and of human zoonotic transmission risk
  - a. Description: Generate geospatial distribution and risk maps (Y1-OY2)
  - b. Status of Effort: The data management system was put into place (Airtable for managing laboratory data) and a draft report written with mock data in order to ensure the data curation system is in place for the start of data collection. The final interface sites were selected for the project for the five regions.
- 5. Strengthen local capacity for detection and reporting of MERS-CoV. Al. and other zoonoses in Jordan, Iraq, and Lebanon
  - a. Description: Host workshops (Y1-2, OY1-2); Submit reports and publications (Y1-OY2); Attend presentations/meetings (Y1-OY2
  - b. Status of Effort: The Project Launch Workshop and Jordan Training Workshop 1 were successfully completed virtually due to the COVID-19 situation, except for the third day of the Jordan Training Workshop 1 which was safely held in-person on May 27<sup>th</sup>. Training at the Human Link lab at the National Research Centre in Egypt is slated to be held in late summer due to the COVID-19 situation in the region. The team will update and evaluate timing of this training considering team safety and local travel and government restrictions to be completed at the earliest possible date.
- 6. Identify specific modifiable risk factors for human infection with these zoonoses
  - a. Description: Conduct causal interference analyses (Y3-OY2); Policy recommendations on modifiable risk factors (Y3-OY2)
  - b. Status of Effort: The associated tasks are slated for Y3-OY2.

### Y1 Accomplishments

# <u>Task 1: Determine the presence of MERS-CoV. Al, and other zoonoses among livestock and poultry in Jordan.</u>

Subtask 1.1 Identify sites for project study

Activities and results this year:

- Project staff reviewed site selection criteria and collaborated with the team to produce a solution for spatial sampling using a spatial sampling application. This application was developed by an EHA scientist for potential applications in studies that sample from mixed human and animal subjects, and from the environment.
- Project staff reviewed site selection criteria and with input from local subject matter experts (SMEs). With their input, the list of sites was narrowed down to include villages with the three interface sites represent

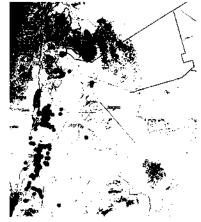


Figure 1. Map of the study area and villages possible to sample from.

include villages with the three interface sites represented and sufficient human population. Using the Shiny App, the team then randomly selected five sites in Jordan from this narrowed down list.

Subtask 1.2 Obtain local permissions and approvals to work with animals

### Activities and results this year:

- The animal owner consent forms were finalized with input
  - from local subject matter experts (SMEs) familiar with sampling communities.
- The JUST IACUC for research with animal subjects was submitted and full approval was obtained.
- JUST approval for use of the animal owner consent forms
   was obtained.
- ACURO approved the animal IACUC and determined the projects protocol was a surveillance project. No further action is required.
- The first draft of the animal sampling protocol has been completed and is currently being reviewed by partners.

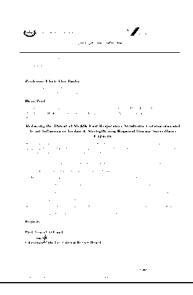


Figure 2. Approval from JUST IRB

### Future Tasks

Subtask 1.3 Conduct biological sampling of animals inthe study areas

Subtask 1.4 Conduct lab testing for AI, MERS-CoV, brucellosis and leptospirosis

# <u>Task 2: Determine whether (and if so, the extent to which) humans at human-livestock/poultry interfaces display evidence of infection.</u>

Subtask 2.1 Identify sites and conduct enrollment for human study component

### Activities and results this year:

- The project staff determined there will likely be a higher rate of loss to follow-up than initially projected. To account for an increased loss to follow-up, the target number for human enrollment for the first year will be increased to 100 people at each site for a total of 500 people for the first year. Protocol and approval documents have been updated as appropriate.
- The project staff have selected the villages below for sampling:
  - 1. WC34+58 Dabet Hanut, Quaira, Aqaba
  - 2. HHF4+55 Al-Mansoura, Shoabak, Ma`an
  - 3. 8G3R83WF+5G Al Damkhi, Karak
  - RRM8+8F Azrag, Zarga
  - 5. F4PG+QH, Hosha Sub-District, Ramtha, Irbid

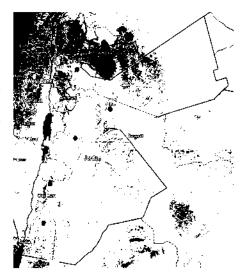


Figure 3. Map of the study area and villages selected within the 5 regions.

 The team developed a draft Human and Animal sampling protocol in preparation for enrollment of human and animal participants.

Subtask 2.2 Obtain local permissions and approvals to work with human subjects

### Activities and results this year:

- Obtained human subjects research approval from the local ethics review board at JUST (2/23/21). The human subjects research and human sampling protocol was submitted to HML IRB for US-based approval.
- Prior to purchasing equipment, the team needs to acquire a tax exemption letter from the Jordanian government.
- Supported JUST in their application for a Federal Wide Assurance number.
- Approval of the human subjects research and human sampling protocol from HML IRB was obtained (4/23/21).
- Submitted an amendment to the local ethics review board at JUST and full approval was given (4/26/21).
- The human subjects research and human sampling protocol was submitted to the Human Research Protection Office (HRPO) (5/4/21).

### Future Tasks

Subtask 2.3 Conduct biological sampling

- Equipment purchased
  - Field Vehicle: 2021 MGRX8

Subtask 2.4 Conduct testing for AI, MERS-CoV, brucellosis and leptospirosis

- Equipment purchased
  - Synergy HTX Multi-Mode microplate reader and PC

### Task 3: Characterize causal factors in animal-to-human transmission of these zoonoses.

Subtask 3.1 Obtain local permissions and approvals to work with human subjects

### Activities and results this year:

- The Jordanian team and EcoHealth Alliance (EHA) behavioral risk scientists collaboratively built and finalized the Animal History Form, Field Data Sheet, Human Questionnaire, and Site Characterization Form.
- Local JUST IRB approval obtained.
- The Human questionnaire, Site Characterization Form, and Animal History Form have been drafted and coded for Open Data Kit (ODK). Tablets were purchased for the Jordan team.



Figure 4. Virtual meeting regarding project questionnaire development and analysis plan

- The JUST IRB approved an amendment for the human subjects research.
- HML IRB approved the protocol for U.S.-based IRB approval.
- The human subjects research and human sampling protocol were submitted to HRPO.

Subtask 3.3 Analyze epi causal inference to identify modifiable risk factors

### Activities and results this year:

- Worked with the project staff and SMEs to compile a list of epidemiological risk factors, behaviors, and general exposure risk factors to be used in data analysis.
- During the human subjects research workshop, the team worked to collaboratively compose the survey guide to provide clarity of question purpose and application for field staff to support data fidelity across sampling events. In scenarios where subjects might need support in understanding questions, ensuring that field staff understand the risk factors being analyzed in each

question will support survey strength.

### Future Subtask(s)

Subtask 3.2 Conduct behavioral risk factor surveys in people

# Task 4: Produce geospatial maps of zoonoses in livestock and poultry and of human zoonotictransmission risk.

Subtask 4.1 Generate maps of laboratory results

### Activities and results this year:

The project staff created a geospatial risk summary report using the package R Markdown in RStudio for future stakeholder meetings that includes maps to demonstrate the percentage of positive tests over the total amount of humans and animals tested for the four zoonotic pathogens in Jordan for this project.

### Future Subtask(s):

Subtask 4.2 Generate risk maps with human behavioral risk factors

# Task 5. Strengthen local capacity for detection and reporting of MERS-CoV. Al. and other zoonoses in Jordan, Iraq, and Lebanon

Subtask 5.1 Conduct project kick-off meeting in Amman with local stakeholders

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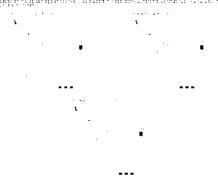


Figure 5. Geospatial map R Markdown file with mock laboratory data

### Activities and results this year:

The Jordan MERS/AI project launch meeting was held on January 27<sup>th</sup>, 2021. In order to adhere to COVID-19 safety guidelines, the meeting was held virtually using a password-protected system. The launch meeting was a success and highlighted the centrality and investment in collaboration for this project. The launch was also an opportunity to engage additional key stakeholders to promote synergies with existing biodefense/health security activities, and provided a forum to share One Health approaches that could potentially be sustained by the Ministry of Health.

### Subtask 5.2 Train local project staff in proper techniques

All research team members are equipped with CITI training in human subjects research, and the global team will continue to provide capacity-building through trainings and workshops on subjects integral to this project and to future public health work, including best practices regarding research and data security, survey tool development, and risk communication.



Figure 6. First and second days of virtual first annual project training workshop



- The first annual project training workshop took place over the course of three days, and was attended by team members, including university students. The first and second day of the workshop was held virtually, and the third day of the workshop was held in person.
- o The sessions held during the two-day virtual workshop included training on:
  - Data management use and storage facilitated by Jamie Cooksey and Dr. Whitney Bagge
  - Ethical human subject research facilitated by Emily Hagan and Stephanie Martinez
  - Ethical animal subject research facilitated by Hani Talafha
  - Questionnaire administration facilitated by Emily Hagan and Stephanie Martinez
  - Safe human and animal sampling facilitated by Dr. Basil Amarneh and Dr. Zaidoun Hijazeen

- The third day of the workshop (May 27<sup>th</sup>, 2021) included training on basic laboratory safety: biosafety and biosecurity, facilitated by Dr. Mustafa Ababneh, and cold chain implementation, facilitated by Dr. Borhan Al-Zghoul.
  - Due to the COVID-19 group limitation, 7 people attended the third in-person workshop;
    - Dr. Mustafa Ababneh (JUST)
    - Dr. Borhan Al-Zghoul (JUST)
    - Yasmin Issa
       Daradkeh (graduate student, JUST)



Figure 7. In-person workshop on basic laboratory safety

- Farah Al. Nabulsi (graduate student, JUST)
- Abed Elrahman Talfha (undergraduate student, JUST)
- Leen Al-Bayari (undergraduate student, JUST)
- Dr. Omar Al-Omari, MD
- March 29, 2021: Dr. Ehab Abu-Basha reviewed the diagnostic laboratories in Al-Karak and Ma'an to determine the limitations and status of veterinary services in the south of Jordan (the report was a follow up to previous visits in 2017 and 2019).
  - Recommendations are as follows:
    - Complete laboratory building infrastructure with suitable benches, sinks, and cabinets (Karak Lab).
    - Hire adequately trained veterinarians and laboratory technicians to the laboratories.
    - Provide adequate training on safety, proper use, and maintenance of laboratory equipment and chemicals.
    - Provide adequate amount of laboratory diagnostic chemicals, supplies and kits.
    - MOA should recruit staff to Ma'an and provide incentives for relocation.

### Subtask 5.6 Data Management

- The project staff have finalized data management tools and decided to use ODK Connect servers to securely store survey data and Airtable servers for human and animal lab data. Only the specific app users for each of the servers can access the secure data.
- Training for the data management systems was completed in the project training workshop.

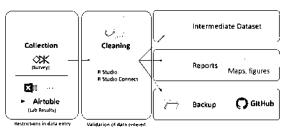


Figure 8. Data management system, from collection to cleaning and reporting results.

- Incorporated the Arabic version of surveys to ODK.
- The project team developed field data sheets and input into the Airtable data management software, and reviewed accessibility and overall design for improvements in conjunction with their review of the field and laboratory protocols.
- The project team has created specific QR codes on ODK Collect for each of the team members to access their own survey to practice using the tablets for when sampling begins.
- For real-time quality control of data collection and entry, the project team has begun coding and developing a series of automated reports



Figure 9. A mock example of the QR codes that each team member received.

using R Studio Connect to be in place for when data collection begins. The intention is that field and laboratory staff serve an active role in data cleaning as they can review data on a biweekly basis and work with EHA data management staff to identify any issues in real time.

The data management team has developed a draft project dashboard for full project progress, to be shared amongst partners to share team and project progress at-a-glance. The dashboard ensures the full team can see the project on all fronts – laboratory, surveys, and human and animal sampling. The dashboard differs from the automated sampling reports, in that the sampling reports are highly detailed accounts of each effort tailored to the sub-team doing that sampling for data cleaning purposes, while the dashboard summarizes the full team effort.

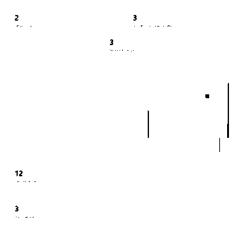


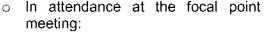
Figure 10. Project Dashboard with mock data.

Subtask 5.7 Conduct presentations/meetings at times and places specified

- Project kick off for HDTRA1-20-1-0029 Jordan was held on January 13<sup>th</sup>, 2021.
   The kick-off meeting was a success and the team had a positive collaboration with DTRA and Jordan partners.
- O Project staff also attended a meeting on March 7<sup>th</sup> at Jordan Armed Forces GHQ (JAF) with LTC. Mohammad Malkawi, Arms Control & International Organizations branch, Directorate of Planning & Organization (DPO), JAF followed-up with letters to the Ministry of Agriculture, Ministry of Health, and Ministry of Environment to form a Focal point committee to the project. Letters were also sent to RSS, Badia Fund, WHO, FAO, USAID, AOAD, and RVC to join the committee. This committee will be updated as part of our projects team ensuring a One Health approach is successfully employed.

#### o Focal Point Meeting

The Focal Point meeting took place in Amman, Jordan on April 6, 2021. Dr. Ehab Abu-Basha presented an overview of the project. This event provided an opportunity to clearly establish the role of the Ministry of Agriculture in the context of the project and to discuss upcoming lab visits in the south of Jordan. Follow-up meetings will be conducted online due to COVID-19 regulations.



- Dr. Ehab Abu-Basha, JUST
- Dr. Zaidoun Hijazeen, JUST
- Dr. Yasmin Daradkeh, JUST
- · Dr. Laura Sawalha, WHO
- Dr. Hussein Zyoud, Head of Animal Health Labs, MoA
- · Dr. Essam Hawwa, CVO, MoA
- · Colonel Oday Mrayat, Jordan Armed Forces

#### Jordan Team Meeting

• The Jordan in-country team held a meeting that discussed training and workshops, the human questionnaire, and purchasing activities in preparation for the beginning of data collection on May 10, 2021. The Jordan team is purchasing lab and field materials and establishing and reviewing standard operating procedure (SOPs) for animal and human sampling, questionnaire administration, and laboratory procedures.

#### Future Subtasks

Subtask 5.3 Host annual partners and stakeholders meeting

Subtask 5.4 Complete annual report and share with partners and stakeholders

Subtask 5.5 Conduct local-regional training workshops

OOVID-19 restrictions have limited in-person workshops, but as the restrictions begin to ease, we are hopeful that we will be able to set a date for the local-regional training workshop, with Human Link Lab at the National Research Centre, Egypt, in August. The team looks forward to engaging with our regional stakeholder to further serve the regional biosecurity and capacity-building objectives of the project.



Figure 11. Focal point meeting in Jordan.

#### Tasks to be initiated after Year 1:

#### Task 6. Identify specific modifiable risk factors for human infection with these zoonoses

#### Samples and Laboratory Techniques

#### Samples:

Sample collection will be performed by the JUST field team, under supervision of the Country PI, Dr. Ehab Abu-Basha. This project is conducted by a large group of scientists, researchers and trained field personnel from Jordan University of Science and Technology and Princess Haya Biotechnology Center. Several graduate students are included in the research team where they will receive intensive training on laboratory techniques including serology such as ELISA, complement fixation test, antibody neutralization test as well as molecular diagnostic techniques including DNA isolation, cDNA, PCR, and sequencing. This group of young researchers will continue to provide research and diagnostic services in various locations in the country. Therefore, the project will not only help us be part of a global initiative for the surveillance of major emerging and remerging viral infections of both human and animal origins, but also will provide a nidus for continued national development of capacity in the surveillance and diagnosis of these diseases in Jordan. Furthermore, the project will help provide animal care takers, farmers, and the public at large with valuable data and unique educational opportunities about the occurrence, recognition, and prevention of important zoonotic diseases.

#### Laboratory or Clinical Techniques:

The pseudo-particle neutralization (ppNT) assay is technique new to JUST and Jordan. Human serum will be tested for MERS-CoV antibodies at the Princess Haya Biotechnology Center and Molecular & Virology Laboratory at JUST. Co-I Kayali from Human Link will be training the JUST team to implement this method.

Many molecular and diagnostic assays by a group of well trained, national scientists and young graduate students and technicians will be performed according to the latest scientific standards. Highly advanced molecular diagnostic assays such as cloning, preparing different PCR products for sequencing, sequence analysis, and utilizing data entry will be performed and routinely used in this laboratory. This will reduce the running time to obtain valuable diagnostic results at a significantly lower cost since scientists and researchers from Jordan used to send such samples to international specialized laboratories around the world for cloning and sequencing.

In Jordan, the molecular and virology laboratory is expected to be an important hub for training and preparing future scientists as many graduate students are being trained in the laboratory to perform various techniques from sample handling to DNA extraction, cDNA synthesis, performing different PCR protocols, cloning, plasmid purification, and sequencing analysis. Several Graduate and undergraduate students will be trained during project time.

## Opportunities for training and professional development:

#### Training

The Jordan Training Workshop 1 provided an opportunity for the team to have in-depth conversations about ethics in human subjects research with particular attention paid to informed consent, recruitment, and safe data handling. Our project's strong focus on tailoring tools and approaches to be reflective of our communities of interest and their protections ensures that team members are fully prepared to work with communities in a manner that is respectful, robust, and able to generate nuanced and important insights. This is an asset to this project, and to the team's future research projects.

In addition to the Jordan Training Workshop 1, all personnel who will be working with human subjects data have completed CITI training in Social-Behavioral-Educational Research and Biomedical Research.

Two graduate and two undergraduate students are being trained as part of this project, including:

- Yasmin Issa Daradkeh (graduate student, JUST)
- Farah Al Nabulsi (graduate student, JUST)
- Abed Elrahman Talfha (undergraduate student, JUST)
- Leen Al-Bayari (undergraduate student, JUST)

During the summer months (July through September), the students will have hands on training for 10 weeks on molecular biology techniques: RNA and DNA extraction, RNA and DNA quantification, cDNA synthesis, conventional and real-time PCR. They will also be trained on how to use an Eliza for serology. Once field work begins, the students will also have the opportunity to participate with the animal and human sampling.

#### Dissemination of results to communities of interest:

The team developed a 2-page project summary to share with stakeholders on project overview, goals, and intended outputs. This document was shared with the Ministry of Agriculture, Ministry of Health, Ministry of Environment, World Health Organization (WHO)-Jordan, Royal Scientific Society (RSS), Jordan Army Forces (JAF), Food & Agriculture Organization of The United Nation (FAO), and Badia Fund.

A focal point Steering Committee was established to regularly to update all partners about our project and help in implementation of One Health Approach with WHO. Our partners on this committee are the Ministry of Agriculture (MOA), Ministry of Health (MOH), Ministry of Environment (MOE), World Health Organization (WHO), Food and Agriculture Organization (FAO), Royal Scientific Society (RRS), and Badia Fund. Each organization has identified its' representative and are now fully engaged in the project.

# Change in plan for the next reporting period to accomplish the goals:

Task 1: Determine the presence of MERS-CoV, AI, and other zoonoses among livestock and/or poultry in Jordan

- 1.3 Conduct biological sampling in animals within the study area
- 1.4 Conduct lab testing for Al. MERS-CoV, brucellosis and leptospirosis
  - These tasks were scheduled to begin in Q3 of Project Year Y1 but due to both the COVID-19 pandemic and pending HRPO approval, we anticipate to begin sampling in Q4 of Project Year Y1.

Task 2: Determine whether (and if so, the extent to which) humans at human-livestock/poultry interfaces display evidence of infection

- 2.3 Conduct biological sampling
- 2.4 Conduct testing for Al. MERS-CoV, brucellosis and leptospirosis
  - Same as above, we anticipate to begin sampling in Q4 of Project Year Y1.

Task 3: Characterize causal factors in animal-to-human transmission of these zoonoses 3.2 Conduct behavioral risk factor surveys in people

- 3.3 Analyze epi causal inference to identify modifiable risk factors
  - Same as above, we anticipate to begin sampling in Q4 of Project Year Y1 and preliminary analysis will shortly follow.

Task 4: Produce geospatial maps of zoonoses in livestock and poultry and of human zoonotic transmission risk

- 4.1 Generate maps of laboratory results and share with partners
- 4.2 Generate risk maps with human behavior risk data and share with partners
  - We anticipate to begin sampling in Q4 of Project Year Y1 and to generate the geospatial maps for the following quarter, Q1, Y2.

Task 5: Strengthen local capacity for detection and reporting of MERS-CoV, Al, and other zoonoses in Jordan, Iraq, and Lebanon

- 5.3 Host annual partners and stakeholders meeting
  - No Change.
- 5.4 Complete annual report and share with partners and stakeholders
  - No Change.
- 5.5 Conduct local/regional training workshops
  - Regional trainings will be conducted considering team safety and regional restrictions on travel. However, we anticipate still holding the training at Human Link in Q4 of Project Year 1 as planned.
- 5.6 Data management
  - No Change.
- 5.7 Conduct presentations/meetings at times and places specified
  - No Change.

		DOMAINS										
	Domain	າ 1	Domain 2			Domain 3				Domain 4		
						СОМРІ	TENCIES					
Training Task	1.1.1	1.1.2	2.1.1	2.1.2	2.1.3	3.1.1	3.1.2	3.1.3	3.1.4	4.1.1	4.1.2	4.1.3
			,	,								
Task 5.5.1 Conduct local- regional training workshops			х	Х		X	X	X				

		DOMAINS												
	Domain	ı 5				Domain 6	;		Domain 1	7		Domain	8	
							COMPE	TENCIES						
Training Task	5.1.1	5.1.2	5.1.3	5.1.4	5.1.5	6.1.1	6.1.2	6.1.3	7.1.1	7.1.2	7.1.3	8.1.1	8.1.2	8.1.3
Task 5.5.1 Conduct local- regional training workshops														
<u> </u>														

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Instructions: this matrix will help track and capture the alignment of BAA research training and professional development activities to BTRP's Domains, Competencies, and Proficiencies Framework. Please note: alignment is only required to the competency level definitions shown below.

- 1. in the left-hand column of the above chart, list every training activity to be performed on a separate line, using their task or sub-task number (ex. Task 2; subtask 2.1; subtask 2.1.1)
- 2. place an x or tally against the competencies that align to the training activity. Competency decisions can be made by determining using the definitions listed below and determining if a) the training subject matter corresponds to the defined competency or b) if the training outcome corresponds to the defined competency. Please note: training activities can align to multiple competencies across the 8 domains. (Note: if completing an annual report, stop at step 2; if drafting a Full Proposal, continue to step 3)
- 3. once completed, list the aligned competencies under its corresponding training activities in the grant proposal.

Domain	Domain Definition	Competency	Competency Definition
1. Disease recognition and prevention of spread	1.1 Understanding of emerging and high-consequence pathogens (EHCP) in the following realms: morbidity, mortality, transmission, control, disease presentation and distribution, and impact.	1.1.1 Disease characteristics and impact	Disease characteristics and epidemiology of especially high consequence pathogens (EHCPs) and the diseases they cause, including etiology, signs and symptoms, morbidity, mortality, risk factors, incubation period, period of communicability, mode of transmission, clinical presentation, mechanisms of spillover, reservoir species, spatiotemporal and geographic distributions of affected populations, and demographic characteristics of affected populations (species/age/sex/occupational status/ etc.), including outcomes on population health.

		<b>1.1.2</b> Clinical Infection control and animal health biosecurity	Understanding of basic infection prevention and control procedures related to especially high consequence pathogens (EHCPs) for human and animal clinical settings in line with GHSA and IHR milestones (when applicable), including standard precautions (PPE use; disinfection; sterilization); environmental infection control; laboratory testing; nosocomial infections, including AMR; and animal health biosecurity principles, procedures.)
2. Disease identification, detection, confirmation, and reporting	2.1 Positive identification of emerging and high consequence pathogens (EHCP) and reporting to appropriate authorities.	2.1.1 Sample collection	Knowledge, understanding and proficiency with procedures for safely and securely collecting appropriate specimens/samples for testing of pathogens (human; animal; environmental), especially EHCPs, and handling and packaging of samples for transport to diagnostic testing laboratories, based on national and international guidelines and best practices, while ensuring specimens will are maintained in condition required for testing (cold chain and or fixation, etc.)
		2.1.2 Test procedures (presumptive/confirmatory)	Knowledge, understanding and proficiency with procedures for conducting screening, diagnostic, and confirmatory testing and interpreting test results for ECHPs from different types of samples using different types of tests (ELISA, PCR, etc.) for high consequence pathogens, including test accuracy, reliability, specificity, and sensitivity and quality control/quality assurance (QA/QC) measures needed to validate testing results.
		2.1.3 Reporting	Knowledge, understanding and proficiency in

			policies and procedures for disease notification and/or bi-directional reporting of surveillance and laboratory data through national information-sharing systems, or to notify regional/international authorities such as WHO/OIE/FAO of a disease, other significant epidemiological event, or potential public health emergency of international concern (as appropriate to roles and responsibilities at each operational level).
3. Analytics, assessment, and research	<b>3.1</b> Appropriate study design, data analysis, and implementation of processes, as applies to each discipline.	3.1.1 Study design and ethics	Knowledge and understanding of scientific methodologies and study design principles (hypothesis development, research plan development, sampling design, strategy for data collection and analysis) and application of research ethics to responsibly address safety, security, and ethical issues related to the research aims and hypotheses, including understanding of issues around dual-use research of concern.
		3.1.2 Methodologies and analysis	Knowledge, understanding and proficiency in the use of statistical and other relevant methodologies for scientific research analyses including practices and approaches for data collection, use and interpretation of statistical data analyses, data visualization, and use of statistical data software tools.
		<b>3.1.3</b> Dissemination of findings and outreach to stakeholders	Knowledge, understanding and proficiency in the dissemination of research findings through peer-reviewed scientific journal publications, conference presentations and public outreach

			activities, including engagement with appropriate stakeholders and partners at national, regional, and international levels.
		3.1.4 Resource mobilization and management	Knowledge, understanding and proficiency in the planning, acquisition, mobilization, allocation, management and tracking of financial and non-financial resources for research projects, including proposal development and grant writing.
4. Management, leadership, and advocacy	4.1 Critical skills, knowledge and behaviors necessary to effectively manage projects and programs, lead teams, champion solutions, inspire action, and sustain programs.	<b>4.1.1</b> Plan, implements, and evaluates a project or program	Knowledge, understanding and proficiency in program planning, implementation, monitoring and evaluation, including development and measurement of appropriate indicators, and advocacy, or management/oversight of the process.
		<b>4.1.2</b> Apply program management techniques throughout the project or program	Knowledge, understanding and proficiency in program management, including timeline and deliverables management, team building, and generating high-level stakeholder support.
		<b>4.1.3</b> Lead culturally diverse teams to complete a project or program	Knowledge, understanding and proficiency in developing and leading culturally diverse teams, ensuring collaboration among team members, and mentorship of future team leaders.

5. Quality	5.1 Knowledge, skills, and	5.1.1 Physical environment and	Knowledge, understanding and proficiency
management	abilities required for	equipment	with facility and equipment operations and
systems	developing a culture of		maintenance, and facility management policies
	quality. Operations,		and procedures, in support of establishing a
	services, and infrastructure		culture regarding physical environment and
	integrated in a system that		equipment necessary for EHCP detection,
	meets applicable standards,		management, and control, or
	professional guidelines, and		management/oversight of these operations, in
	customer requirements for		a system that meets applicable standards,
	ensuring and maintaining quality and continually		professional guidelines, and customer requirements for ensuring and maintaining
	improving services.		quality and continually improving services.
	improving services.	5.1.2 Qualified, well-trained, and	Knowledge, understanding and proficiency
		competent workforce	with the principles, processes, and
		,	methodologies for workforce development
			and sustainment in relation to EHCP detection,
			management, and control, including workforce
			planning, training, refresher training,
			continuing education, and/or professional
			development, in a system that meets
			applicable standards, professional guidelines,
			and national licensing requirements where
		5.1.3 Materiel resource management	applicable.  Knowledge, understanding and proficiency
		3.1.3 Materier resource management	with procedures and policies for procurement
			and inventory management of materiel
			resources, including supply chain systems,
			operations, and infrastructure, or
			management/oversight of these processes.

		<b>5.1.4</b> Quality review practices and procedures	Knowledge, understanding and proficiency with quality assurance, quality control, and/or quality management review practices and procedures, based on national or international best practices.
		5.1.5 Document management	Knowledge, understanding and proficiency with document and information management policies and procedures, and program documentation and records/archival management processes.
6. Outbreak preparedness and surveillance	<b>6.1</b> Use of operational guidelines, identification and analysis of outbreak situations, and effective communication with stakeholders.	6.1.1 Operational guidelines	Knowledge, understanding and proficiency with development and implementation of outbreak preparedness, surveillance, investigation, laboratory practices, and response planning and operations guidelines, including case definitions, and demonstrated ability to use related operational tools such as emergency preparedness and response plans, contingency plans, SOPs, and job aids when available. Special note and care should be given to documents designed for health care workers directly combatting an outbreak.
		6.1.2 Identification and analysis	Knowledge and proficiency with principles and processes for developing and maintaining disease outbreak situational awareness capabilities, including coordination and

			communication or reporting across surveillance, laboratory, and other data sources, epidemic intelligence, data analysis, and field epidemiological and outbreak investigation techniques, such as contact tracing, and decision-making in support of these activities.
		<b>6.1.3</b> Communication systems	Knowledge and proficiency with development, implementation, and sustainment of communications systems for outbreak situational awareness and response, including multisectoral and intrasectoral coordination (including via emergency operations centers or equivalent assets), operational risk communication, and public risk communication.
7. Biosafety	7.1 The application of knowledge, techniques, and equipment to prevent accidental exposure to and release of emerging and high consequence pathogens (EHCP).	<b>7.1.1</b> Biohazard Identification and classification	Knowledge, understanding and proficiency with identification and classification of biohazards associated with EHCPs, for laboratories, hospitals, other biomedical and clinical facilities, and the community, including biohazard risk assessment and characterization, and biohazard identification policies and compliance procedures.
		<b>7.1.2</b> Biohazard Control Measures (e.g., personal protective equipment, safety practices and equipment, facility design, and administrative controls).	Knowledge, understanding and proficiency with development and implementation of international best practices and procedures for EHCP biosafety and biohazard control measures (substitution, elimination, mitigation) for personnel, equipment, facilities (laboratories, hospitals, and other biomedical and clinical facilities) and the community.

		<b>7.1.3</b> Biohazard Incident Management	Knowledge, understanding and proficiency with development and implementation of policies, procedures, and operational guidelines, including emergency operation communications systems, for Biohazard Incident Management for EHCP outbreaks, releases or exposures.
8. Laboratory biosecurity	8.1 Prevent theft, misuse, loss, or deliberate release of emerging and high consequence pathogens (EHCP).	8.1.1 Biosecurity Risk Identification	Knowledge, understanding and proficiency with principles and procedures for Biosecurity Risk identification, risk assessments, characterization and compliance with recommended international standards and best practices to prevent the loss, theft, or misuse of biological organisms and laboratory/research technologies.
		8.1.2 Biosecurity Risk Mitigation	Knowledge, understanding and proficiency with international best practices for Biosecurity Risk Mitigation, and related risk reduction approaches to enhance physical security, personnel reliability, material control and accountability, transport security, and information security/cybersecurity, and the development and implementation of Biosecurity Risk Mitigation strategies for laboratories, facilities, and personnel to prevent the loss, theft, or misuse of biological organisms, especially EHCPs, and laboratory/research technologies.
		8.1.3 Biosecurity Incident Management	Knowledge, understanding and proficiency with international best practices for Biosecurity Incident Management, and development and implementation of

	Biosecurity incident management policies and
	procedures for suspected or confirmed illicit
	access to pathogens, intruder alerts, and other
	biosecurity incidents.
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Please fill out the requested information on all tabs (click on each bottom colored tab or sheet) labeled Cover Page, Personnel, Publications, Presentations, Impacts, Intellectual Property, and Transitions. For annual reports, please only describe <u>DTRA funded work</u> for the reporting period (July 1, 2019 - June 30, 2020).

Tab	Description
Cover Page	
	Providing information about the project participants and collaborating organizations allows an assessment of performance in promoting partnerships and collaborations.
<u>Publications</u>	The products from an effort demonstrate the excellence of the research and efficacy with which the results are being communicated to colleagues, potential users, and the public. Publications are the characteristic product of research. If there is nothing to report during this reporting period, the awardee shall state "Nothing to Report".
Presentations	The products from an effort demonstrate the excellence of the research and efficacy with which the results are being communicated to colleagues, potential users, and the public. Many projects (thought not all) may develop products other than publications. In all cases, if there is nothing to report during this reporting period, the awardee shall state "Nothing to Report".
Impacts	knowledge, enlarges the pool of people trained to develop that knowledge and techniques or put it to use, and improves the physical, institutional, and information resources that enable those people to get their training and perform their functions. In all cases, if there is nothing significant to report during this reporting period, the awardee shall state "Nothing to Report".
	Information on any intellectual property demonstrates the value of protectable innovative ideas resulting from sponsored research and includes patents, inventions, and licenses. Submission of this information is not a substitute for other invention reporting requirements under the awards terms and conditions. In all cases, if there is nothing
Intellectual Property	significant to report during this reporting period, the awardee shall state "Nothing to Report".  Refers to cases where knowledge resulting from your DTRA sponsored effort will be further developed for, or will be
<u>Transitions</u>	used in, a technology application. Transition sponsors can be entities in DoD, other federal agencies, or industry.

SHADED FIELDS ARE REQUIRED

Report Submitted to **Defense Threat Reduction Agency** 

HDTRA1-20-1-0029 Grant/Award# Organization/Institution EcoHealth Alliance

Project Title

Reducing the Threat of Middle East Respiratory Syndrome Coronavirus and Avian Influenza in Jordan & Strengthening Regional Disease Surveillance Capacity

Name of Submitting Official Dr. William B. Karesh

Title of Submitting Official Executive Vice President for Health and Policy

karesh@ecohealthalliance.org Email Address

212-380-4463 Phone Number

Submission Date Reporting Period End Date June 30th, 2021

Annual

Foreign Spending Country Foreign Spending (\$)

7/1/2021

Country 1 201,514.72 Jordan

Country 2 Country 3

Reporting Period

#### **Proceed to Personnel**

Enter the principle investigator (PI) and all other personnel that have worked at least one person month (\*160 hours) on this DTRA project during the reporting period, regardless of the source of compensation.

regardless of the	regardless of the source of compensation.								
	Select the appropriate				Forter the number of examples - Forter the I.C.1 name of the	Forter the I.L. L. name of the	Enter the Hirsch index for Pland	Select whother a	Enter a graduation or anticipated
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Award #	Регѕоппе) Туре	Last name	First name	Suffix	Person Months Worked	Organization/Institution	Hirsh Index	Graduated	Graduation date [students]
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HDTRA1-20-1-0029	Research Associate/Scientist	Bagge	Whitney		4	EcoHealth Alliance			
HDTRA1:20-1-0029	Research Associate/Scientist	Hagan	Етіў		2	EcoHealth Alliance			
HDTRA1-20-1-0029	Research Associate/Scientist	Martinez	Stephanie		2	Ecohealth Alliance			
HDTRA1-20-1-0029	Research Associate/Scientist	Cooksey	Jamie		2	Ecohealth Alliance			
HDTRA1-20-1-0029	Research Associate/Scientist	Abu-Basha	Ehab		9	JUST			
HDTRA1-20-1-0029	Research Associate/Scientist	Hijazeen	Zaidoun		9	T\$Ut			
HDTRA1-20-1-0029	Research Associate/Scientist	Talafha	Нал		80	ISDI			
HDTRA1 20 1 0029	Research Associate/Scientist	Ababneh	Mustafa		5	JUST			
HDTRA1-20-1-0029	Research Associate/Scientist	Malkawı	Zuhair		9	JUST			
HDTRA1.20-1-0029	Research Assuciate/Scientist	AL-Zoghoul	Borhan		5	TSDI			
HDTRA1-20-1-0029	Research Associate/Scientist	Al-Omari	Bilal		5	JUST			
HDTRA1-20-1-0029									
HDTRA1-20-1-0029	Research Associate/Scientist	Gharaibeh	Saad		1	JUST			
HDTRA1 20 1 0029	Graduate Student	Daradkeh	Yasmin		5	JUST		Ne	Jar 22
HDTRA1-20-1-0029	Graduate Student	Al Nabulsi	Farah		5	JUST		Yes	Sep-20
		Elrahman							
HDTRA1-20-1-0029	Undergraduate Student	Talfha	Abed		2	UST		No	Jul-23
HDTRA1-20-1-0029	Undergraduate Student	Al-Bayari	Leen		5	JUST		No	Jul-24
HDTRA1-20-1-0029	Technician	Aahmawi	Alaa		1	JUST			
HDTRA1-20-1-0029	Technician	Al-Hanandeh Sief	Sicf		5	ISDI			
HDTRA1 20 1 0029	Technician	Hammdeh	Mehammad		5	JUST			
HDTRA1-20-1-0029	Research Associate/Scientist	Amarneh	Basil		2	JUST			
HDTRA1:20-1-0029	Research Associate/Scientist	Al-Omari	Отаг		2	JUST			
HDTRA1-20-1-0029	Research Associate/Scientist	Jaradat	Saied		1	JUST			
HDTRA1-20-1-0029	Technician	Andre	Amanda		2	Ecohealth Alliance			
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**Proceed to Publications** 

Enter all publications (Journal Articles, Books, Book Chapters, Conference Papers, and Theses/Dissertations) resulting from this DTRA project during this reporting period. DO NOT include publications that have not yet been submitted.

Instructions	Selection appropriate published type from the drop described Selection to the drop described Selection of the drop described to the described to the described selection of the describ	Enter the fall name of		Esterate EUL, name of the article (ide (chapte) in the (flood)				Enter the first page number of the article or chapter		ident Fer type	Anter the pile leation dentifier non-burifier the Type selected in the arey observations	an acknowledgment	Shire Ewhether or not a journal article or conference busernus gene talleuga the even receive a rocces.					
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**Proceed to Presentations** 

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Select the Enter the location of the

presentation type
from the drop down
list. Select Nothing
to Report if there

| The control of the presentation in the present

format. wereino presentations

(conference) as City, State for domestic society, or university). Country for international meetings.

Select from the - Select whether or not an drap down list the acknowledgement of DTRA personnel type — support was included in the for the presenter. presentation.

	presentations.				international meetings.			
Award #	Presentation Type	Presentation (Paper) Title	Conference Date	Conference Name	Conference Location City, State	Author(s)	Personnel Type	Acknowledgement DTRA Support
		infrarea Spectroscopy of		Materials Research				
Fxampie	Orai Keynate		8/1/2015	Society Fall Meeting	Boston, MA	John Smith	Groquate Student	yes
HDTRA1-20-1-0029	Oral Invited	One Health in the Pander	6 24 2021	Walter Reed Army Ins	Is Online		PI	Yes
			6/2/20121				Associate/Scientis	
4DTRA1 20 1 0029	Oral Invited	Preventing the Next Pand	0.7/2021	American Association	o Online		t	Yes
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HDTRA1-20-1-0029								
HDTRA1-20-1-0029								
HDTRA1-20-1-0029								
IDTRA1 20 1 0029								
4DTRA1 20 1 0029								
HOTRA1-20-1-0029								
HDTRA1-20-1-0029								
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HOTRA1-20-1-0029								
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#### **Proceed to Impacts**

Enter awards, honors, promotions, or graduations that occurred during the reporting period for the personnel contributing to this DTRA project.

Award #	Impact Type	Impact Description	Personnel Type	Awardee Name	Impact Date	Impact Details/Website
		Best Graduate Student Poster at the				
		American Physical Society winter				
Example	Best Poster	conference	Graduate Student	John Smith	9/20/2019	
HDTRA1-20-1-0029	Nothing to Report					
HDTRA1-20-1-0029						
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HDTRA1-20-1-0029						
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**Proceed to Intellectual Property** 

Enter all patents, invention disclosures, and licenses resulting from this DTRA award that were awarded or applied for during the reporting period. Reporting here does NOT replace the reporting requirements for patents and invention disclosures in the terms and conditions of the award.

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#### **Proceed to Transitions**

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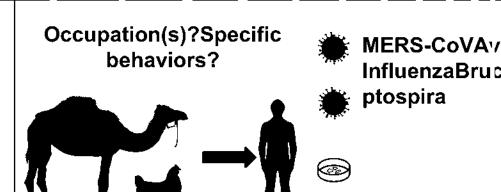
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# Reducing the Threat of Middle East Respiratory Syndrome Coronavirus and Avian Influenza in Jordan, PI: William B. Karesh, EcoHealth Alliance, HDTRA1-20-1-0029

Objective: Work with government and private sector partners to characterize causal factors in animal-to-human transmission of MERS, Al, and other zoonoses in Jordan, implement serological technology for enhanced diagnostic and detection capabilities, and strengthen regional detection and reporting capacity to reduce biological threats. Threat Reduction Impact: Establishing laboratory and field SOPs to assist partners in detecting presence and exposure to highconsequence zoonotic pathogens. Capacity/Capability Building: Pseudoparticle neutralization assay technology transfer from Human Link to JUST, training of graduate and undergraduate students, site selection and sampling design methods, human behavioral risk Status of effort. Animal sampling approval survey development obtained, HRPO approval pending, field and laboratory teams prepping for field work and sample testing. Personnel Supported: This project is supporting 7 faculty, 6 other scientists, 2 graduate students, 2 undergraduate studentsPublications & Meetings: 13 workshop participants



Major Goals and Milestones:Determine evidence of MERS, AI, and other zoonoses in humans, livestock and poultry in Jordan (Y1-OY2)Identify risk factors and modifiable behaviors for threat reduction (Y1-OY2)Improve knowledge and diagnostic capacity (Y1-OY2)Funding Profile Year 1 09/2020-09/2021: \$984,351, Year 2 09/2021-09/2022: \$981,695, Year 3 09/2023-09/2024: \$991,118Contact information PI: Dr. William B. Karesh,

212-380-4463 Karesh@ecohealthalliance.org



### Biological Threats Reduction Program PROJECT QUARTERLY REPORT

	PROJECT INFORMATION	ON
	HDTRA12010029 - Reduc	_
Project and Title:	East Respiratory Syndrome	
Troject and Title.	Influenza in Jordan & Stren	igthening Regional Disease
	Surveillance Capacity	
Quarter and Dates:	Y1, Q1	(9/29/20 - 12/28/20)
Prepared by:	Dr. William B. Karesh	

#### OVERALL PROJECT SUMMARY AND OBJECTIVES

#### **Project Objective**

Utilizing an international consortium of partners, this effort is a multi-disciplinary One Health research project to identify causal factors in the transmission of MERS-CoV, AI, and other zoonoses in Jordan, while simultaneously strengthening local capacity for prevention, detection, and reporting of these zoonoses in Jordan, Iraq, and Lebanon. Our proposed project will rigorously test the following initial hypotheses: 1) That livestock species and poultry in Jordan show evidence of infection with MERS-CoV, AI, respectively and/or other zoonoses. 2) That people regularly exposed to livestock and/or poultry or their living areas have a greater risk of infection with MERS-CoV, AI, and/or other zoonoses compared to other community members in Jordan. 3) That some specific practices may promote exposure to specific animal biohazards (e.g., blood, urine, feces, and/or nasal secretions) and pose a greater risk of infection with MERS-CoV, AI, and other zoonoses compared to others. This study involves interviews and longitudinal sampling of humans, livestock, and poultry at animal markets and farms for MERS-CoV, Brucella spp., and Leptospira spp. (people/livestock) and for AI subtypes (people/poultry). Partner laboratories in Jordan will screen and characterize these zoonotic pathogens, and capacity-building cross-trainings will be conducted with visiting laboratory scientists from Iraq and Lebanon. This multifaceted One Health approach will advance scientific knowledge on the risk of MERS-CoV, AI, and other zoonoses in Jordan, strengthen local zoonotic disease surveillance capabilities, and enhance scientific partnerships in Jordan, Iraq and Lebanon to prevent, detect, and report biological threats.

#### YEAR 1, QUARTER 1 PROGRESS UPDATES FOR EACH TASK

### Task 1: Determine the presence of MERS-CoV, AI, and other zoonoses among livestock and poultry in Jordan.

Subtask 1.1 Identify sites for project study

Activities and results this quarter:

 Project staff reviewed site selection criteria within the 5 project sites; final site selection to occur prior to commencing sampling in Q3.

Subtask 1.2 Obtain local permissions and approvals to work with animals

Activities and results this quarter:

- o JUST IACUC for research with animal subjects was submitted and full approval was obtained.
- IACUC for research with animal subjects was submitted for pre-review, comments were addressed from prereview, and the subsequent amended application was submitted for full review to Tufts University.
- Drafted animal owner consent form.

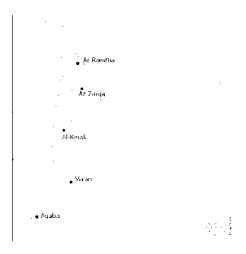


Figure 1. Map of the 5 sites for project study in Jordan.

#### Future Tasks

Subtask 1.3 Conduct biological sampling of animals in the study areas Subtask 1.4 Conduct lab testing for AI, MERS-CoV, brucellosis and leptospirosis.

### Task 2: Determine whether (and if so, the extent to which) humans at human-livestock/poultry interfaces display evidence of infection.

Subtask 2.1 Identify sites and conduct enrollment for human study component

Activities and results this quarter:

 Project staff reviewed site selection criteria within the 5 project sites; final site selection to occur prior to commencing sampling in Q3.

Subtask 2.2 Obtain local permissions and approvals to work with human subjects

Activities and results this quarter:

 Compiled materials for local JUST and HML IRBs, finalized human questionnaire.



Figure 2. Virtual meeting to discuss the project questionnaire

Future Tasks

Subtask 2.3 Conduct biological sampling

Subtask 2.4 Conduct testing for AI, MERS-CoV, brucellosis and leptospirosis

#### Task 3: Characterize causal factors in animal-to-human transmission of these zoonoses.

Subtask 3.1 Obtain local permissions and approvals to work with human subjects Activities and results this quarter:

 Compiled materials for local JUST and HML IRBs. The Jordanian team and EHA behavioral risk scientists collaboratively built and finalized the human questionnaire and site characterization form.

Subtask 3.3 Analyze epi causal inference to identify modifiable risk factors. Activities and results this quarter:

- O Project staff reviewed data analysis plan using causal inference framework to identify all data needed for the models that will be produced to test study hypotheses. All proposed model variables were mapped onto specific questions to be asked in the questionnaire. The questionnaire was then arranged according to logical conversational flow in light of how it will be administered.
- Examined socio-economic factors that could affect vulnerability to both risks and impacts, including household wealth indicators used in prior surveys (e.g. for comparability with World Bank reports). These will be relevant for economic context and policy recommendations for biothreat reduction.
- o Project staff investigated the literature for risk factors and confounders associated with leptospirosis, brucellosis, AI, and MERS-COV.

Future Subtask(s)

Subtask 3.2 Conduct behavioral risk factor surveys in people

### <u>Task 4: Produce geospatial maps of zoonoses in livestock and poultry and of human zoonotic</u> transmission risk.

Subtask 4.1 Generate maps of laboratory results

Activities and results this quarter:

 The project staff identified the workflow and mock-ups for generation of these maps for quarterly reporting.

Subtask 4.2 Generate risk maps with human behavioral risk factors

Activities and results this quarter:

 The project staff identified which exposures would be best to map for inclusion in the questionnaire.

### <u>Task 5. Strengthen local capacity for detection and reporting of MERS-CoV, AI, and other</u> zoonoses in Jordan, Iraq, and Lebanon

Subtask 5.1 Conduct project kick-off meeting in Amman with local stakeholders Activities and results this quarter:

- A date and preliminary schedule for the kick-off meeting was set.
- Coordinated with the Jordan Armed Forces health official, including organizing an upcoming briefing meeting on the project. We anticipate this will be a positive collaboration for the project, expanding key stakeholders to promote synergies with existing biodefense/health security activities and provide a forum to share One Health approaches that could potentially be sustained by Ministry of Health.

#### Subtask 5.6 Data Management

 Software, systems, and workflow was determined for data management (see Figure 3). These will be tailored to outputs for Jordanian partners (e.g., reports that can be shared at the monthly national One Health platform meeting convened by Ministry of Health, as relevant).

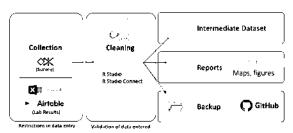


Figure 3. Summary of data management software and data flow for this project.

#### Future Tasks

Subtask 5.2 Train local project staff in proper techniques

• All team members will be equipped with CITI training and the team will continue planning future training to understand research and data security, and survey tool development.

Subtask 5.3 Host annual partners and stakeholders meeting

Subtask 5.4 Complete annual report and share with partners and stakeholders

Subtask 5.5 Conduct local-regional training workshops

Subtask 5.7 Conduct presentations/meetings at times and places specified

#### Tasks to be initiated after Year 1:

#### Task 6. Identify specific modifiable risk factors for human infection with these zoonoses

#### Presentations and Publications

#### 1. Presentations and Other Conference Submissions: All Institutes

 Developed slide set to present project concept for wider awareness of an applied One Health approach for threat reduction.

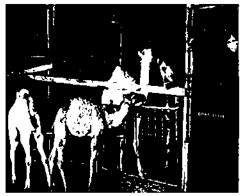
#### 2. Publications: All Institutes

No project scientific publications to report as of yet. However, the project team developed a project summary sheet for outreach to stakeholders in Jordan (as shown below).



### Reducing the Threat of MERS-CoV and AI in Jordan & Strengthening Regional Disease Surveillance Capacity

Jordan has experienced several important zoonotic diseases in the few past years including avian influenza A (AI) strains H1N1 and H5N1 and Middle East Respiratory



Syndrome Coronavirus (MERS-CoV). High-consequence zoonotic pathogens in the Middle East have resulted in disease epidemics of international importance, raising the need for health security enhancements to strengthen the region's ability to prevent, detect and respond to multiple zoonotic disease threats. MERS-CoV is recognized as a threat to global health security, with designation as a priority disease under the World Health Organization (WHO) Research and Development Blueprint. Although most cases have been reported in the Kingdom of Saudi Arabia (KSA), the first known human infections occurred in Jordan in 2012.

Camels are the presumptive source of primary human MERS-CoV infections; however, the exact mechanisms of transmission and the possible role of other livestock species are unclear.

These uncertainties highlight the need to better understand the causal factors of primary human infection with MERS-CoV at human-livestock interfaces given the high livestock-human contact in camel production systems in Jordan. Avian influenza viruses are widely distributed all over the world. In Jordan, the first outbreak of AI strain H5N1 was reported in 2006. The development and wide spread circulation of a new and highly pathogenic strain of AI could present a serious risk to both animal and humans and could result in major socio-economic losses. In fact, there has been a rapid expansion of the poultry industry in Jordan in recent years due to increased demand which increases the risk of AI to public health, locally as



well as regionally and globally. Therefore, identification and characterization of AI subtypes circulating among poultry across Jordan will provide critical information needed to understand baseline activity and risks.



Utilizing an international consortium of partners, we are conducting a multi-disciplinary One Health research project to identify causal factors in the transmission of MERS-CoV, AI, and



community members in Jordan.

- other zoonoses in Jordan, while simultaneously strengthening local capacity for prevention, detection, and reporting of these zoonoses in Jordan, Iraq, and Lebanon. Our proposed project will rigorously test the following initial hypotheses:
- 1) That livestock species and poultry in Jordan show evidence of infection with MERS-CoV, AI, respectively and or other zoonoses.
- 2) That people regularly exposed to livestock and or poultry or their living areas have a greater risk of infection with MERS-CoV, AL and or other zoonoses compared to other
- 3) That some specific practices may promote exposure to specific animal biohazards (e.g., blood, urine, feces, and or nasal secretions) and pose a greater risk of infection with MERS-CoV, AI, and other zoonoses compared to others.

This study involves interviews and longitudinal sampling of humans, livestock, and poultry at animal markets and farms for MERS-CoV. Brucella spp., and Leptospira spp. (people livestock) and for AI subtypes (people poultry). Partner laboratories in Jordan will screen and characterize these zoonotic pathogens, and capacity-building cross-trainings will be

conducted with visiting laboratory scientists from Iraq and Lebanon. This multifaceted One Health approach will advance scientific knowledge on the risk of MERS-CoV. AI, and other zoonoses in Jordan, strengthen local zoonotic disease surveillance capabilities, and enhance scientific partnerships in Jordan, Iraq and Lebanon to prevent, detect, and report biological threats.

The project is supported by the U.S. DoD Defense Threat Reduction Agency's Biological Threats Reduction Program.



### Biological Threats Reduction Program PROJECT QUARTERLY REPORT

	PROJECT INFORMATION	ON
	HDTRA12010029 - Reduci	ing the Threat of Middle
Project and Title:	East Respiratory Syndrome	
Project and Title.	Influenza in Jordan & Strer	gthening Regional Disease
	Surveillance Capacity	
Quarter and Dates:	Y1, Q2	(12/29/20 - 03/28/21)
Prepared by:	Dr. William B. Karesh	

#### OVERALL PROJECT SUMMARY AND OBJECTIVES

#### **Project Objective**

Utilizing an international consortium of partners, this effort is a multi-disciplinary One Health research project to identify causal factors in the transmission of MERS-CoV, AI, and other zoonoses in Jordan, while simultaneously strengthening local capacity for prevention, detection, and reporting of these zoonoses in Jordan, Iraq, and Lebanon. Our proposed project will rigorously test the following initial hypotheses: 1) That livestock species and poultry in Jordan show evidence of infection with MERS-CoV or AI, respectively and/or other zoonoses. 2) That people regularly exposed to livestock and/or poultry or their living areas have a greater risk of infection with MERS-CoV, AI, and/or other zoonoses compared to other community members in Jordan. 3) That some specific practices may promote exposure to specific animal biohazards (e.g., blood, urine, feces, and/or nasal secretions) and pose a greater risk of infection with MERS-CoV, AI, and other zoonoses compared to others. This study involves interviews and longitudinal sampling of humans, livestock, and poultry at animal markets and farms for MERS-CoV, Brucella spp., and Leptospira spp. (people/livestock) and for AI subtypes (people/poultry). Partner laboratories in Jordan will screen and characterize these zoonotic pathogens, and capacity-building cross-trainings will be conducted with visiting laboratory scientists from Iraq and Lebanon. This multifaceted One Health approach will advance scientific knowledge on the risk of MERS-CoV, AI, and other zoonoses in Jordan, strengthen local zoonotic disease surveillance capabilities, and enhance scientific partnerships in Jordan, Iraq and Lebanon to prevent, detect, and report biological threats.

#### YEAR 1, QUARTER 1 PROGRESS UPDATES FOR EACH TASK

### <u>Task 1: Determine the presence of MERS-CoV, AI, and other zoonoses among livestock and poultry in Jordan.</u>

Subtask 1.1 Identify sites for project study Activities and results this quarter:

- Project staff reviewed site selection criteria and collaborated with the team to produce a solution for spatial sampling using a "Shiny App." This novel app will be used to randomly select sampling sites in Jordan.
- The Site Characterization form has been drafted and coded with Open Data Kit (ODK). ODK is software that allows for the creation of survey tools and the collection, management, and usage of data in lowbandwidth environments.

Subtask 1.2 Obtain local permissions and approvals to work with animals

Activities and results this quarter:

- The animal owner consent forms were finalized with input from local subject matter experts (SMEs) familiar with sampling communities.
- Jordan University Science and Technology (JUST) IACUC for research with animal subjects was submitted and full approval was obtained.
- o JUST approval for use of the animal owner consent forms was obtained.
- ACURO approved the animal IACUC and determined the projects protocol was a surveillance project. No further action is required.
- The first draft of the animal sampling protocol has been completed and is currently being reviewed by partners.

#### Future Tasks

Subtask 1.3 Conduct biological sampling of animals in the study areas

Subtask 1.4 Conduct lab testing for AI, MERS-CoV, brucellosis and leptospirosis.



Figure 1. An example of the Shiny Apprandomly selecting sites (the red dots).

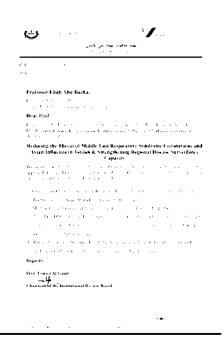


Figure 2. Approval letter from Jordan's IRB committee.

### Task 2: Determine whether (and if so, the extent to which) humans at human-livestock/poultry interfaces display evidence of infection.

Subtask 2.1 Identify sites and conduct enrollment for human study component Activities and results this quarter:

o The project staff determined that it is likely that there will be a higher rate of loss to follow-up than initially projected. To account for an increased loss to follow-up, the target number for human enrollment for the first year will be increased to 100 people at each site for a total of 500 people for the first year. Protocol documents have been updated as appropriate.

Subtask 2.2 Obtain local permissions and approvals to work with human subjects Activities and results this quarter:

- Obtained human subjects research approval from the local ethics review board at JUST (2/23/21). The human subjects research and human sampling protocol will now be submitted to Health Media Lab IRB Research and Ethics (HML) for domestic approval.
- o Prior to purchasing equipment, the team needs to acquire a tax exemption letter from the Jordanian government.
- o Supported JUST in their application for a Federal Wide Assurance number.

#### Future Tasks

Subtask 2.3 Conduct biological sampling

Subtask 2.4 Conduct testing for AI, MERS-CoV, brucellosis and leptospirosis

#### Task 3: Characterize causal factors in animal-to-human transmission of these zoonoses.

Subtask 3.1 Obtain local permissions and approvals to work with human subjects Activities and results this quarter:

- The Jordanian team and EcoHealth Alliance (EHA) behavioral risk scientists collaboratively built and finalized the Animal History Form, Field Data Sheet, Human Questionnaire, and Site Characterization Form.
- Local JUST IRB approval obtained.
- The Human questionnaire, Site Characterization Form, and Animal History Form have been drafted and coded for Open Data Kit (ODK). Tablets were purchased for the Jordan team.

Subtask 3.3 Analyze epi causal inference to identify modifiable risk factors Activities and results this quarter:

- Worked with the project staff and SMEs to compile a list of epidemiological risk factors, behaviors, and general exposure risk factors to be used in data analysis.
- O During the human subjects research workshop, worked to collaboratively fill-out a survey guide to provide clarity of question purpose and application for field staff. In scenarios where subjects might need support in understanding questions, ensuring that field staff understand the risk factors being analyzed in each question will support survey strength.

#### Future Subtask(s)

Subtask 3.2 Conduct behavioral risk factor surveys in people

### <u>Task 4: Produce geospatial maps of zoonoses in livestock and poultry and of human zoonotic</u> transmission risk.

#### Future Subtask(s):

Subtask 4.1 Generate maps of laboratory results Subtask 4.2 Generate risk maps with human behavioral risk factors

### <u>Task 5. Strengthen local capacity for detection and reporting of MERS-CoV, AI, and other</u> zoonoses in Jordan, Iraq, and Lebanon

Subtask 5.1 Conduct project kick-off meeting in Amman with local stakeholders Activities and results this quarter:

The Jordan MERS/AI project launch meeting was held on January 27<sup>th</sup>, 2021. In order to adhere to COVID-19 safety guidelines, the meeting was held virtually using a password-protected system. The launch meeting was a success and highlighted the centrality and investment in collaboration for this project. The launch was also an opportunity to engage additional key stakeholders to promote synergies with existing biodefense/health security activities, and provided a forum to share One Health approaches that could potentially be sustained by the Ministry of Health.

#### Subtask 5.2 Train local project staff in proper techniques

- All research team members are equipped with CITI training in human subjects research, the team will continue to provide capacity-building through trainings and workshops on subjects integral to this project and to future public health research, including best practices regarding research and data security, and survey tool development.
  - The first annual project training workshop took place over the course of three days, and was attended by all team members, project training workshop fig and was attended by all team members, project training workshop fig project training workshop fig and workshop workshop figures and was attended by all team and the string days of the

Figure 3. First day of the virtual first annual project training workshop.

- including senior scientist and university students. The first and second day of the workshop was held virtually and the third day of the workshop will be held in person. The date of the third workshop will be determined soon, depending on local regulations and will include safety measures in place for COVID-19.
- The third day of the workshop will include training on basic laboratory safety: biosafety and biosecurity, facilitated by Dr. Mustafa Ababneh, and cold chain implementation, facilitated by Dr. Borhan Al-Zghoul.
- o The sessions held during the two-day virtual workshop included training on:
  - Data management use and storage facilitated by Jamie Cooksey and Dr. Whitney Bagge
  - Ethical human subject research Emily Hagan and Stephanie Martinez
  - Ethical animal subject research- Hani Talafha
  - Questionnaire administration facilitated by Emily Hagan and Stephanie Martinez

 Safe human and animal sampling - led by Dr. Basil Amarneh and Dr. Zaidoun Jijazeen

#### Subtask 5.6 Data Management

- The project staff have finalized data management tools and decided to use ODK connect servers to securely store survey data and Airtable servers for human and animal lab data. Only the specific app users for each of the servers can access the secure data.
- Training for the data management systems was completed in the project training workshop.



Figure 4. Second virtual day: first annual project training workshop.

#### Subtask 5.7 Conduct presentations/meetings at times and places specified

- Project kick off for HDTRA1-20-1-0029 Jordan was held on January 13<sup>th</sup>, 2021. The kick-off meeting was a success and the team had a positive collaboration with DTRA, and Jordan partners.
- Meeting on March 7<sup>th</sup> at Jordan Armed Forces GHQ (JAF) with LTC. Mohammad Malkawi, Arms Control & International Organizations branch, Directorate of Planning & Organization (DPO), JAF to follow up with letters send to Ministry of Agriculture, Ministry of Health and Ministry of Environment to form a Focal point committee to the project. Letters are also sent to RSS, Badia Fund, WHO, FAO, USAID, AOAD and RVC to join the committee. This committee will follow up on One Health approach as well.

#### Future Tasks

Subtask 5.3 Host annual partners and stakeholders meeting

Subtask 5.4 Complete annual report and share with partners and stakeholders

Subtask 5.5 Conduct local-regional training workshops

 A proposed date has been set for the training at the Human Link Lab at the National Research Centre, Egypt. While details remain to be confirmed due to considerations for COVID-19, the team looks forward to engaging with our regional stakeholder to further serve the regional biosecurity and capacity-building objectives of the project.

#### Tasks to be initiated after Year 1:

#### Task 6. Identify specific modifiable risk factors for human infection with these zoonoses

#### Presentations and Publications

- 1. Presentations and Other Conference Submissions: All Institutes
  - Developed slide set to present project concept for wider awareness of an applied One Health approach for threat reduction.
- 2. Publications: All Institutes

No project scientific publications to report as of yet.

### Biological Threats Reduction Program PROJECT QUARTERLY REPORT

	PROJECT INFORMATI	ON
Project and Title:	HDTRA12010029 - Reduc East Respiratory Syndrome Influenza in Jordan & Stret Surveillance Capacity	•
Quarter and Dates:	Y1, Q3	(03/29/21- 6/29/21)
Prepared by:	Dr. William B. Karesh	

#### OVERALL PROJECT SUMMARY AND OBJECTIVES

#### **Project Objective**

Utilizing an international consortium of partners, this effort is a multi-disciplinary One Health research project to identify causal factors in the transmission of Middle East Respiratory Syndrome Coronavirus (MERS-CoV), Avian Influenza (AI), and other zoonoses in Jordan, while simultaneously strengthening local capacity for prevention, detection, and reporting of these zoonoses in Jordan, Iraq, and Lebanon. Our proposed project will rigorously test the following initial hypotheses: 1) That livestock species and poultry in Jordan show evidence of infection with MERS-CoV or AI, and/or other zoonoses. 2) That people regularly exposed to livestock and/or poultry or their living areas have a greater risk of infection with MERS-CoV, AI, and/or other zoonoses compared to other community members in Jordan. 3) That some specific practices may promote exposure to specific animal biohazards (e.g., blood, urine, feces, and/or nasal secretions) and pose a greater risk of infection with MERS-CoV, AI, and other zoonoses compared to others. This study involves interviews and longitudinal sampling of humans, livestock, and poultry at animal markets and farms for MERS-CoV, Brucella spp., Leptospira spp. (people/livestock), and for AI subtypes (people/poultry). Partner laboratories in Jordan will screen and characterize these zoonotic pathogens, and capacity-building cross-trainings will be conducted with visiting laboratory scientists from Iraq and Lebanon. This multifaceted One Health approach will advance scientific knowledge on the risk of MERS-CoV, AI, and other zoonoses in Jordan, strengthen local zoonotic disease surveillance capabilities, and enhance scientific partnerships in Jordan, Iraq, and Lebanon to prevent, detect, and report biological threats.

#### YEAR 1, QUARTER 1 PROGRESS UPDATES FOR EACH TASK

### <u>Task 1: Determine the presence of MERS-CoV, AI, and other zoonoses among livestock and poultry in Jordan.</u>

Subtask 1.1 Identify sites for project study

Activities and results this quarter:

O Project staff reviewed site selection criteria and with input from local subject matter experts (SMEs), the list of sites was narrowed down to include villages with the three interface sites represented and sufficient human population. From this list of thirteen potential sites, the Shiny App, was used to randomly generate the final five sites in Jordan to be used for the field study components of the project. Jordian project partners participated throughout the process and learned the methodology and rationale for randomizing site selection.

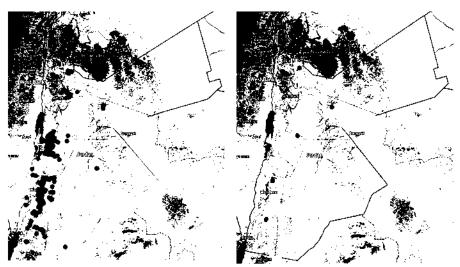


Figure 1. Sites selected using spatial sampling "Shiny App" (red points). Left image contains every point prior to SME local input and the right image contains the randomly selected points.

Subtask 1.2 Obtain local permissions and approvals to work with animals Activities and results this quarter:

 Jordan ACUC and Tufts IACUC approvals and ACURO surveillance project determination were obtained in previous quarters. No further action is required.

#### Future Subtask(s)

Subtask 1.3 Conduct biological sampling of animals in the study areas Subtask 1.4 Conduct lab testing for AI, MERS-CoV, brucellosis and leptospirosis.

### Task 2: Determine whether (and if so, the extent to which) humans at human-livestock/poultry interfaces display evidence of infection.

Subtask 2.1 Identify sites and conduct enrollment for human study component Activities and results this quarter:

- The project staff have selected the villages below for sampling:
  - 1. WC34+58 Dabet Hanut, Quaira, Agaba
  - 2. HHF4+55 Al-Mansoura, Shoabak, Ma'an
  - 3. 8G3R83WF+5G Al Damkhi, Karak
  - 4. RRM8+8F Azraq, Zarqa
  - 5. F4PG+OH, Hosha Sub-District, Ramtha, Irbid
- o The team developed a draft Human and Animal sampling protocol in preparation for enrollment of human and animal participants.

Subtask 2.2 Obtain local permissions and approvals to work with human subjects

Activities and results this quarter:

- o The project team submitted the study protocol to Health Media Lab IRB Research and Ethics (HML) for U.S. IRB approval. Approval was obtained (4/23/21).
- o The project team submitted an amendment to the local ethics review board at JUST and approval was given (4/26/21).
- o The human subjects research and human sampling protocol was submitted to the Human Research Protection Office (HRPO) (5/4/21).

Subtask 2.3 Conduct biological sampling

- Equipment purchased
  - Field Vehicle: 2021 MGRX8, delivery pending completion of tax exemption approval

Subtask 2.4 Conduct testing for AI, MERS-CoV, brucellosis and leptospirosis

- o Equipment purchased
  - Synergy HTX Multi-Mode microplate reader and PC

#### Task 3: Characterize causal factors in animal-to-human transmission of these zoonoses.

Subtask 3.1 Obtain local permissions and approvals to work with human subjects

Activities and results this quarter:

- The JUST IRB approved an amendment for the human subjects research.
- HML IRB approved the protocol for U.S.-based IRB approval.
- The human subjects research and human sampling protocol were submitted to HRPO.

#### Future Subtask(s)

Subtask 3.2 Conduct behavioral risk factor surveys in people

Subtask 3.3 Analyze epi causal inference to identify modifiable risk factors

# Task 4: Produce geospatial maps of zoonoses in livestock and poultry and of human zoonotic transmission risk.

Subtask 4.1 Generate maps of laboratory results

Activities and results this quarter:

o The project staff created an R Markdown document in RStudio for future stakeholder meetings that includes maps to demonstrate the percentage of positive tests over the total amount of humans and animals tested for the four zoonotic pathogens of interest in Jordan for this project.

# Future Subtask(s):

Subtask 4.2 Generate risk maps with human behavioral risk factors

# <u>Task 5. Strengthen local capacity for detection and reporting of MERS-CoV, Al, and other zoonoses</u> in Jordan, Iraq, and Lebanon

Subtask 5.2 Train local project staff in proper techniques

- Due to an increase in COVID-19 cases in Jordan, the third date for the in-person workshop was finalized for May 27<sup>th</sup>, 2021. The workshop included training on basic laboratory safety: biosafety and biosecurity, facilitated by Dr. Mustafa Ababneh, and cold chain implementation, facilitated by Dr. Borhan Al-Zghoul.
  - Due to the COVID-19 group limitation,
     7 people attended the third in-person workshop:
    - Dr. Mustafa Ababneh (JUST)
    - Dr. Borhan Al-Zghoul (JUST)
    - Yasmin Issa Daradkeh (graduate student, JUST)
    - Farah AlNabulsi (graduate student, JUST)
    - Abed Elrahman Talfha (undergraduate student, JUST)
    - Leen Al-Bayari (undergraduate student, JUST)
    - Dr. Omar Al-Omari, MD



Figure 2. In-person workshop on basic laboratory safety.

- March 29, 2021: Dr. Ehab Abu-Basha reviewed the diagnostic laboratories in Al-Karak and Ma'an to determine the limitations and status of veterinary services in the south of Jordan (the report was a follow up to previous visits in 2017 and 2019).
  - o Recommendations are as followed:
    - Complete laboratory building infrastructure with suitable benches, sinks, and cabinets (Karak Lab)
    - Hire adequately trained veterinarians and laboratory technicians to the laboratories
    - Provide adequate training on safety, proper use and maintenance of laboratory equipment and chemicals.
    - Provide adequate amount of laboratory diagnostic chemicals, supplies and kits.
    - MOA should recruit staff to Ma'an and provide incentives for relocation.

# Subtask 5.6 Data Management

Incorporated the Arabic translations of the surveys into ODK.

- The project team developed field data sheets and input into the AirTable data management software, and reviewed accessibility and overall design for improvements in conjunction with their review of the field and laboratory protocols.
- The project team has created specific QR codes on ODK Collect for each of the team members to access their own survey to practice using the tablets for when sampling begins.
- For real-time quality control of data collection and entry, the project team has coded and developed a series of automated reports using R Studio Connect to be in place for when data collection begins. The intention is that field and laboratory
  - staff serve an active role in data cleaning as they can review data on a biweekly basis and work with EHA data management staff to identify any issues in real time
    - The ODK validation report has been coded for the purpose of real-time quality control for the surveys that are delivered by field team members.
- The data management team developed a draft project dashboard for full project progress monitoring, to be shared among partners to share team and project progress at-a-glance. The dashboard ensures the full team can see project activities on all fronts—laboratory, surveys, human and animal sampling. The dashboard differs from the automated sampling reports, in that the sampling reports are highly detailed accounts of each effort tailored to the sub-team doing that sampling for data cleaning purposes, while the dashboard summarizes the full team effort.



Figure 3. A mock example of the QR codes that each individual team member

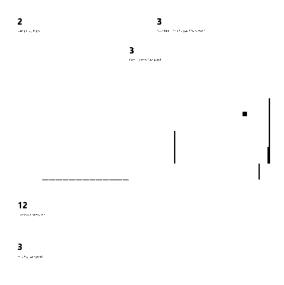
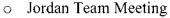


Figure 4. An example of the dashboard demonstrating how data will be summarized.

Subtask 5.7 Conduct presentations/meetings at times and places specified Activities and results this quarter:

o Focal Point Meeting

- o April 6, 2021: The focal point meeting took place in Amman, Jordan, in April. Dr. Ehab Abu Basha gave a brief description of the project. This event provided an opportunity to clearly establish the role of the Ministry of Agriculture in the context of the project and to discuss upcoming lab visits in the south of Jordan. Follow-up meetings will be conducted online due to COVID-19 regulations.
- o In attendance to the focal point meeting:
  - Dr. Ehab Abu Basha, JUST
  - Dr. Zaidoun Hijazeen, JUST
  - Dr. Yasmin Daradkeh, JUST
  - Dr. Laura Sawalha, WHO
  - Dr. Hussein Zyoud, Head of Animal Health Labs, MoA
  - Dr. Essam Hawwa, CVO, MoA
  - Colonel Oday Mrayat, Jordan Armed Forces



May 10, 2021: The Jordan in-country team held a meeting to discuss training and workshops, the human questionnaire, and purchasing activities in preparation for the beginning of data collection. The Jordan team will purchase lab and field materials and establish and review standard operating procedure (SOPs) for animal and human sampling, questionnaire administration, and laboratory procedures.

#### Future Subtask(s)

Subtask 5.1 Conduct project kick-off meeting in Amman with local stakeholders

Subtask 5.3 Host annual partners and stakeholders meeting

Subtask 5.4 Complete annual report and share with partners and stakeholders

Subtask 5.5 Conduct local-regional training workshops

OOVID-19 restrictions have limited in-person workshop, but as the restrictions begin to ease, we are hopeful that we will be able to set a date for the local-regional training workshop, with Human Link Lab at the National Research Centre, Egypt, in August. The team looks forward to engaging with our regional stakeholder to further serve the regional biosecurity and capacity-building objectives of the project.

#### Tasks to be initiated after Year 1:

### Task 6. Identify specific modifiable risk factors for human infection with these zoonoses

Figure 5. Focal point meeting in Jordan.

**Quarterly Report:** HDTRA12010029 - Year 1, Quarter 3 (03/29/21 6/30/21)

#### Presentations and Publications

#### 1. Presentations and Other Conference Submissions: All Institutes

- June 7, 2021: Dr. Whitney Bagge presented to the American Association of University Women (AAUW) on "Preventing the Next Pandemic: Science and Capacity Building.
- June 24, 2021: Dr. William Karesh presented a webinar for the Walter Reed Army Institute of Research (WRAIR) One Health Branch, including AFRIMS and other overseas labs on "One Health in the Pandemic Era" and discussed DTRA BTRP projects including this one in Jordan.

# 2. Publications: All Institutes

No project scientific publications to report as of yet.

# **Biological Threats Reduction Program PROJECT QUARTERLY REPORT**

	PROJECT INFORMATION	ON
	HDTRA12010029 - Redu	ucing the Threat of Middle
Duciest and Title	East Respiratory Syndror	ne Coronavirus and Avian
Project and Title:	Influenza in Jordan & Str	rengthening Regional
	Disease Surveillance Cap	pacity
Quarter and Dates:	Y1, Q4	(06/30/21-9/29/21)
Prepared by:	Dr. William B. Karesh	

#### OVERALL PROJECT SUMMARY AND OBJECTIVES

### **Project Objective**

Utilizing an international consortium of partners, this effort is a multi-disciplinary One Health research project to identify causal factors in the transmission of Middle East Respiratory Syndrome Coronavirus (MERS-CoV), Avian Influenza (AI), and other zoonoses in Jordan, while simultaneously strengthening local capacity for the prevention, detection, and reporting of these zoonoses in Jordan, Iraq, and Lebanon. Our proposed project will rigorously test the following initial hypotheses: 1) That livestock species and poultry in Jordan show evidence of infection with MERS-CoV or AI, and/or other zoonoses. 2) That people regularly exposed to livestock and/or poultry or their living areas have a greater risk of infection with MERS-CoV, AI, and/or other zoonoses compared to other community members in Jordan. 3) That some specific practices may promote exposure to specific animal biohazards (e.g., blood, urine, feces, and/or nasal secretions) and pose a greater risk of infection with MERS-CoV, AI, and other zoonoses compared to others. This study involves interviews and longitudinal sampling of humans, livestock, and poultry at animal markets and farms for MERS-CoV, Brucella spp., Leptospira spp. (people/livestock), and for AI subtypes (people/poultry). Partner laboratories in Jordan will screen and characterize these zoonotic pathogens, and capacity-building cross-trainings will be conducted with visiting laboratory scientists from Iraq and Lebanon. This multifaceted One Health approach will advance scientific knowledge on the risk of MERS-CoV, AI, and other zoonoses in Jordan, strengthen local zoonotic disease surveillance capabilities, and enhance scientific partnerships in Jordan, Iraq, and Lebanon to prevent, detect, and report biological threats.

# YEAR 1, QUARTER 4 PROGRESS UPDATES FOR EACH TASK

# Task 1: Determine the presence of MERS-CoV, AI, and other zoonoses among livestock and poultry in Jordan.

Subtask 1.1 Identify sites for project study

Activities and results this quarter:

• Four project staff from Ecohealth Alliance traveled to Amman, Jordan from August 6<sup>th</sup> - 14<sup>th</sup> to provide a two-day workshop and an investigation of selected study sites. During the trip the project staff visited four different potential human-animal interface sites. Two of the were located in the Tura subdistrict in Al-Ramtha and the other two sites were located in the Hmeimeh and the Al-Dissi subdistricts in Quaira, Aqaba.



Figure 1. A camel interface site in the Tura subdistrict in Al-Ramtha in the north of Jordan.

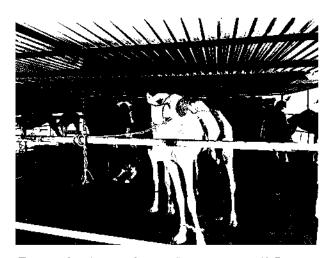


Figure 2. A camel interface site in Al-Dissi, Quaira, Aqaba in south of Jordan.

Subtask 1.2 Obtain local permissions and approvals to work with animals

Activities and results this quarter:

 All required approvals for animal work have been obtained. The team is still awaiting approval from the Office for Human Research Protections (HRPO) so that human and animal work can be conducted simultaneously (One Heath approach).

#### Future Subtask(s)

Subtask 1.3 Conduct biological sampling of animals in the study areas Subtask 1.4 Conduct lab testing for AI, MERS-CoV, brucellosis and leptospirosis.

# <u>Task 2: Determine whether (and if so, the extent to which) humans at human-livestock/poultry interfaces display evidence of infection.</u>

Subtask 2.1 Identify sites and conduct enrollment for human study component Activities and results this quarter:

O During the August 6<sup>th</sup> to 14<sup>th</sup> trip to Jordan, the project staff investigated the sites that were randomly chosen for the project. They communicated and discussed the project's objectives with the people who lived/worked with the camels at four different interface sites. These sites were located in the Hmeimeh and the Al-Dissi subdistricts in Quaira, Aqaba, and the Tura subdistrict in Al-Ramtha, Irbid.



Figure 3. Meeting with the four families from a camel interface site in Hmeimeh subdistrict in Quaira, Agaba.



Figure 4. Discussing the project with the head of the household at a camel interface site in the Tura subdistrict in Al-Ramtha, Irbid.

# Subtask 2.3 Conduct biological sampling

- Equipment purchased
  - o Field Vehicle: 2021 MGRX8 was delivered and is currently with the in-country team.
  - o Lab supplies were purchased for the project. These supplies include PCR testing kits, direct RNA sequencing kits, tubes and caps, pipettes, probes, and etc.
  - The JUST team created a purchasing committee from three members that will review different vendors for product quality and price, and recommend the purchases of items accordingly. In addition, they will also follow up with products once delivered to check specifications and sign the required receipt forms as per JUST regulations.

## Subtask 2.4 Conduct testing for AI, MERS-CoV, brucellosis and leptospirosis.

 During the trip to Jordan, project team members visited Jordan University of Science and Technology (JUST), and visited the laboratories that the project's lab personnel will be using to conduct testing on samples for exposure to AI, MERS-CoV, Brucella Sp. and Leptospira Sp.



Figure 5. Mohammad Hussien Alboom demonstrating how to analyse genetic sequencing results.



Figure 6. (L-R) Zaidoun Hijazeen, Jamie Cooksey, Emily Hagan, Dr. Ehab Abu-Basha, Dr. Mustafa Ababneh, Dr. Whitney Bagge, Dr. William Karesh, Dr. Borhan Al-Zghoul, and Mohammad Hussien Alboom in the molecular biology lab at JUST.

# Future Subtask(s)

Subtask 2.2 Obtain local permissions and approvals to work with human subjects

o The team is still awaiting approval from the Office for Human Research Protections (HRPO).

### Task 3: Characterize causal factors in animal-to-human transmission of these zoonoses.

Subtask 3.1 Obtain local permissions and approvals to work with human subjects Activities and results this quarter:

• The team is still awaiting approval from the Office for Human Research Protections (HRPO).

# Future Subtask(s)

Subtask 3.2 Conduct behavioral risk factor surveys in people

Subtask 3.3 Analyze epi causal inference to identify modifiable risk factors

# <u>Task 4: Produce geospatial maps of zoonoses in livestock and poultry and of human zoonotic transmission risk.</u>

Subtask 4.1 Generate maps of laboratory results Activities and results this quarter:

 Updated the original geospatial map with mock data that is connected to the project's AirTable to create updated automated reports to be sent to our project partners and stakeholders.

# Future Subtask(s):

Subtask 4.2 Generate visk maps with human behavioral visk factors

Figure 7. Updated geospatial map based on laboratory results of humans, livestock, and poultry positive for the four pathogens.

Task 5. Strengthen local capacity for detection and reporting of MERS-CoV,

AI, and other zoonoses in Jordan, Iraq, and Lebanon

Subtask 5.2 Train local project staff in proper techniques

- The JUST graduate and undergraduate students working on the project train for 8 weeks in PCR techniques at the Molecular and Biology Laboratory, JUST. After training, the students may come in on their own time to practice.
- September 14, 2021, the human studies field team, graduate and undergraduate students gathered for training on how to use the tablets and give the questionnaire to the participants.
- The project staff team and the students have created a "Whats Λpp" group chat for communication.

Figure 8. Dr. Basil demonstrates how to use the tablets and questionnaires to the human field team, graduate, and undergraduate students.

Subtask 5.5 Conduct local-regional training workshops. Activities and results this quarter:

On August 10<sup>th</sup> & 11<sup>th</sup>, 2021, EHA project staff travelled to Amman, Jordan, to lead a two-day Survey Design and Risk Communications Workshop at the Grand Hyatt Amman. The workshop was a highly interactive event in which project team leaders facilitated presentations, discussions, and workbook activities designed to build skills around question and survey design and risk communications strategies. The workshop additionally included training on key tenets of ethical research with human populations. Including the project team, 48 participants attended the first day, and 37 participants attended the second day. In keeping with the One Health foundations of the project, an important component of this workshop was having a group of participants who would be representative of a diverse cross-section of government ministries and national One Health stakeholders from the public and private sector. As the participant list below illustrates, this was successfully achieved. Our esteemed participants received certificates at the close of the workshop, and the project team received positive feedback and suggestions for future trainings.



Geospatial Maps of Zoonoses in Livestock, Poultry, and Humans in Jordan.

# Objectives Day One

- o Refresher on ethics in research with human populations
- o Review of key principles of research design and preparation
- o Discussion of strategies of how to design surveys with strong questions

# o Objectives Day Two

- o Application of lessons learned from Day One in drafting real survey questions
- o Introduction to the concept of risk
- o Introduction to the concept of risk communication
- o Practice key message development
- o Practice risk communication skills through examples

# Attendance for Day One

- o Ministry of Health
  - o Dr. Alaa Hamdallah
  - o Dr. Saed Alnserat
  - o Dr. Nizar Masouda
  - o Dr. Akef Sleihat
  - o Dr. Hayel El-Buriqi
  - o Dr. Fatima Zerriouh
- Ministry of Agriculture
  - o Dr. Esam Hawa, Chief Veterinary Officer
  - o Dr. Majed Hawaosheh
  - o Dr. Samer Aldwairi
  - Dr. Khaldoon Malkawi
  - o Dr. Hassan Al-Husainat
  - o Dr. Marwan Awabdeh
  - o Dr. Sharher Abu Dalbouh
  - O Dr. Haitham Λthamneh
- Ministry of Environment
  - Eng. Belal Shaqarin, Director of Climate Change Directorate
  - Eng. Saddam Diab Λl-Khawaldeh, Nature Protection Directorate
  - Eng. Nour M Alghazo, Climate Change Directorate
  - Noura Hamad Λlshra'a, Climate Change Directorate
- Jordan University of Science and Technology
  - Dr. Mohammad Khalifeh, Dean of Faculty of Veterinary Medicine
  - o Dr. Tareq Mukattash, Dean of Research
  - o Dr. Ehab Abu-Basha, Project Lead
  - o Dr. Mustafa Ababneh
  - o Mr. Bilal Al-Omari
  - o Dr. Mohammad Borhan Al-Zghoul
  - o Dr. Basil Hameed Amarneh
  - o Dr. Saied Jaradat
  - o Dr. Omar Al-Omari
  - o Eng. Hani Talafha
  - Alaa Fahmawi, Nurse, King Abdulla Hospital



Figure 9. Invited speakers table at opening of the workshop.



Figure 10. A 10-minute presentation on One Health in Jordan by Dr. Alaa Hamdallah from the Ministry of Health.



Figure 11. Group exercise during day one of the Survey Design and Risk Communication Workshop.

- Ayesha Al-Kofahi, Research assistant
- Nour Alhoda, Research assistant
- o Graduate Students
  - Dr. Yasmin Daradkeh
  - o Dr. Farah Al Nabulsi
- Undergraduate Students
  - Leen Al Bayari
  - Abdelrahman Talafha
- Royal Scientific Society
  - o Dr. Ghadeer Alzghoul
  - o Dr. Mu'men Alrwashdeh
- o WHO
  - o Dr. Lora Al-Sawalha, One Health Professional Officer
- EMPHNET
  - o Dr. Moad Rahamneh
- United Animal Health Inc.
  - Dr. Nadim Amarin, marketing and Technical Manager, METΛ
- Private Sector Veterinarians
  - o Dr. Zaidoun Hijazeen
  - Dr. Brittany Crook, ΛUS
- DTRA
  - LTC. Todd Dahmann

#### Attendance for Day Two

- Ministry of Health
  - Dr. Alaa Hamdallah
  - Dr. Saed Alnserat
  - o Dr. Nizar Masouda
  - Dr. Akef Sleihat
  - o Dr. Hayel El-Burigi
  - o Dr. Fatima Zerriouh
  - Ministry of Agriculture
  - o Dr. Esam Hawa, Chief Veterinary Officer
  - o Dr. Majed Hawaosheh

- o Dr. Samer Aldwairi
- o Dr. Khaldoon Malkawi
- o Dr. Hassan Al-Husainat
- Dr. Marwan Awabdeh
- o Dr. Haitham Athamneh
- Dr. Safa Atoom, Head of Microbiology Unit
- Ministry of Environment
  - Eng. Nour M Alghazo, Climate Change Directorate
  - Noura Hamad Alshra'a, Climate Change Directorate
    - Abdelrahman Talafha
- Royal Scientific Society
  - o Dr. Ghadeer Alzghoul
- Jordan University of Science and Technology
  - o Dr. Ehab Abu-Basha, Project Lead
  - o Mr. Bilal Al-Omari
  - Dr. Mohammad Borhan Al-Zghoul
  - o Dr. Omar Al-Omari
  - o Eng. Hani Talafha
  - Ayesha Al-Kofahi
  - o Nour Alhoda
  - Graduate Students
    - Dr. Yasmin Daradkeh
  - o Undergraduate Students
    - Leen Al Bayari
    - Abdelrahman Talafha
- WHO
  - Dr. Lora Al-Sawalha, One Health Professional Officer
  - o Olle Mjergura
- o EMPHNET
  - o Dr. Moad Rahamneh
- Private Sector Veterinarians
  - o Dr. Zaidoun Hijazeen
  - o Dr. Brittany Crook, AUS



Figure 12. Group photo of the participants from the August training workshop.



Figure 13. Certificates given out at the end of day two after the completion of the workshop.

# Subtask 5.6 Data Management

- On August 8th, the project staff met at the Princess Haya Biotechnology Center at Jordan University of Science and Technology to review the data management system AirTable and discuss any concerns prior to starting sampling.
  - O The team decided to use a general QR code and have team members select their name from a list in the administration section of the survey.
  - The team created validation reports for the field data and lab data. The data validation reports will help the field team check for extreme values or any data entry errors when the project team inputs data into AirTable or ODK surveys.

Figure 14. On August 8, 2021, the team gathered at JUST University, Jordan to discuss AirTable and data management.

Subtask 5.7 Conduct presentations/meetings at times and places specified

- On September 9, 2021, Dr. Ehab Abu-Basha conducted a meeting with both field and lab teams and the following activities were undertaken:
  - Responsibilities and tasks such as sample collection, transportation, sample identification and labeling, receiving, and processing of samples in the lab and questionnaire administration were described and identified for each member of each team.
  - o SOPs previously prepared were reviewed and the human and animal team were instructed to strictly follow the SOPs. A workshop to train team members how to execute each task according to SOP is set to take place in the coming weeks.

- September 14, 2021, the field and lab teams had a meeting to discuss the logistics of cold chain transport of samples from different project sites to JUST. The field team will draw a chart to summarize this process according to the agreed SOPs.
- September 16, 2021, Dr. Whitney Bagge conducted a regional planning meeting to discuss the upcoming training at Human Link Lab at the National Research Centre in Egypt.
  - Those in attendance:
    - EcoHealth: Dr. William Karesh Dr. Whitney Bagge, Jamie Cooksey, Amanda Andre, Darya Ivanova
    - JUST: Dr. Ehab Abu-Basha, Dr. Zaidoun Hijazeen, Dr. Bilal Al Omari, Yasmin Daradkeh, Farah Al. Nabulsi
    - Human Link: Ghazi Kayali
  - The meeting concluded that the training would take place for 5 days in the second half of November. Security clearance for participants to enter the lab will need to be cleared before travel to Egypt since the lab is a governmental facility.

## Future Subtask(s)

Subtask 5.1 Conduct project kickoff meeting in Amman with local
stakeholders
Subtask 5.3 Host annual partners
and stakeholders meeting
Subtask 5.4 Complete annual
report and share with partners and
stakeholders

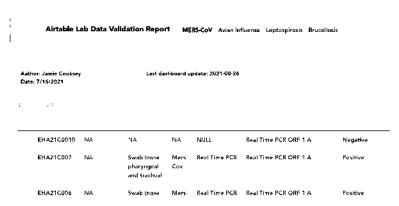


Figure 15. An updated version of the validation reports that will be sent to the team members that will inform of any extreme or missing values.

#### Tasks to be initiated after Year 1:

#### Task 6. Identify specific modifiable risk factors for human infection with these zoonoses

#### Presentations and Publications

# 1. Presentations and Other Conference Submissions: All Institutes

- Or. Ehab Abu-Basha was invited to attend WHO 3rd Quarterly Meeting "Strengthening One Health in Jordan" which was hold on July 15<sup>th</sup>, 2021, in Amman. Dr. Abu-Basha gave a presentation about Strengthening Regional Disease Surveillance MERS CoV and AI in Jordan. He also discussed the methodology and the goals of the current project.
- o Dr. Ehab Abu-Basha was invited as keynote speaker to the World Zoonoses Day 2021 on July 6<sup>th</sup> by the Turkish Veterinary Association and gave a presentation about "One Health in Action". He addressed the importance of One Health in action, strengthening collaboration for detection and surveillance of zoonotic diseases, establishment of a national One Health committee, capacity building and community trust and taising awareness.

# 2. Publications: All Institutes

No project scientific publications to report as of yet.

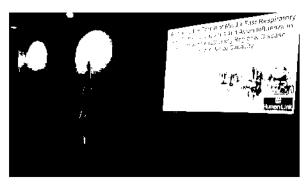


Figure 16. Dr. Ehab Abu-Basha at the WHO's 3<sup>rd</sup> Quarterly Meeting.



Figure 17. World Zoonoses Day 2021 conference was virtual on Zoom.

From: Aleksei Chmura To: (b)(6) Cc: Noam Ross; (b)(6) (b)(6) ; <u>Joe Riccardi</u> [Non-DoD Source] Re: Notification of DTRA Grant Award -- HDTRA12110023 -- "Predicting Biothreat Impacts Subject: from Early-Stage Data via Transfer Learning Date: Monday, July 19, 2021 4:32:15 PM All active links contained in this email were disabled. Please verify the identity of the sender, and confirm the authenticity of all links contained within the message prior to copying and pasting the address to a Web browser. I confirm receipt and we are delighted to revive this Grant award!

Please note our address may need to be updated, since it now is:

**EcoHealth Alliance** 520 Eighth Avenue **Suite 1200** New York, NY 10018

Thank you!

-Aleksei

Aleksei Chmura, PhD Chief of Staff

EcoHealth Alliance 520 Eighth Avenue, Suite 1200 New York, NY 10018-4182

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+1.212.380.4473 \le \text{tel}: 1.212.380.4469 \ge \text{(office)}
(b)(6)
                                                     (mobile)
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Caution-www.ecohealthalliance.org < Caution-http://www.ecohealthalliance.org >

EcoHealth Alliance develops science-based solutions to prevent pandemics and promote conservation.

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On Jul 19, 2021, at 15:5
Caution(b)(6)
                                                      > wrote:
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Dr. Ross and Dr. Chmura,

Congratulations! Attached for your records is the subject Grant award. Please let me know if you have any questions about the attached document. FYSA - also ce'ed to this email is the GOR for this effort, Dr. Batni.

Thank you,

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Contracting Officer	_	
	uction Agency, AL-ACR	
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# **COST PROPOSAL**

Prime 1 (Fill-in) Sub 1 (Fill-in)

#### **GENERAL INFORMATION**

The purpose of the requested information in the attached worksheets is to assist government personnel in the review and evaluation of cost proposals submitted by offerors. Options: Unpriced Options will not be accepted. Any Option that is not fully priced, will not be included in any resulting award.

Enter the proposed cost detail for the Base and each Option period (as needed) on the "Total Amount" tab.

- The formulas in this spreadsheet are based upon common business practices; however, Offerors may edit formulas as necessary (edited formulas must still be visible).
- Ensure all costs from other worksheets are correct.

Below is a summary of the proposed cost. This chart will automatically fill in from the "Total Amount" tab.

Total Direct Labor Costs	\$286,937
Total Fringe Benefit Costs	\$105,593
Total Labor Overhead Costs	\$0
Total Subcontract Costs	\$0
Total Consultant Costs	\$0
Total Other Direct Costs	\$22,660
Total Material Handling Costs	\$0
Subtotal Costs	\$420,190
Total G&A Costs	\$150,596
Subtotal Costs	\$570,787
Total Cost of Money	\$0
Total Estimated Costs	\$570,787
Fixed Fee (If proposing a CPFF contract)	\$0
Total Estimated Costs Plus Fixed Fee	\$570,787

Cost Proposal - T-	otal Amount																		
Prime Offeror:	Printe 1 (Fil-in):																		
Subcontractor:																			
	•			BASE			OPTION 1			OPTION 2			OPTION 3			OPTION 4		Т	olal
Direct Labor (DL)				MMUYR to MMUYR			MMYR to MMYR			MM/YR to MM/YR			MMITR to MMUYE	1		MMCYR to MMCYR	2	_	
	Offeror's Labor Category (Note 1)	Name (Last, First)	Base/Hours	Rate	Extended 5	Base/Hours	Rate	Extended 5	BaseiHours	Rate	Extended 5	Base/Hours	Rate	Extended 5	Base/Hours	Rate	Extended 5	Base/Hours	Extended 5
	Principal Scientist	Hoss Norm	520	\$50.29	328 750 00	620	\$59.05	330 187 50	:	SOLID	30.00	p	30.00	30.00	D	Struc	String	1.040	
	Research Scientist	Mendelsonn, Emma	- 240	540.87	\$42 500 00	1,040	\$42.91	\$44 625 20	347	\$42.91	\$14 875 00	Ö	\$0.00	\$0.02	Ö	50.02	50.02	2,427	
	Senior Engineer	Yоыла, Rot	1,040	\$50.48	352 500 00	1,1140	\$53.00	365 125 00	347	363 00	\$16,375,00	p	30 00	30.00	0	String	Smind	2.427	
	Total Direct Labor Costs		2,600	\$146.63	\$123,750.00	2,600	\$153.97	\$129,937.50	693	\$95.91	\$33,250.00	D	\$0.00	\$0.00	D	\$0.00	\$0.00	5,893	
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	Enngo Benefas	See Lagoofting Raies Tab	\$123,750	26 800%	\$45,540,00	\$129.937	36.800%	\$47,817.00	\$30,250	38 800 W	\$12,200,00	50	0.000%	20.00	90	0.00055	50.00	\$266 937 49	\$105,593 (A)
	Total Fringe Benefit Costs				\$45,540,00			\$47,817.00	,		\$12,235,00			\$0.00			\$0.00		5105,593,00
	H) (See Escalation and Indirect Rate Worksheet)		<b>†</b>					,			<b>*</b> · · · <b>,</b> · · · · · · · · · · · · · · · · · · ·						*****		
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	Total Other Direct Costs	000 000 000 000			\$17,184,00			\$10,496.00			\$0.00			\$0.00			\$0.00		\$22,860.D0
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- Costo Pive Fixed Fee | India 3| \$153,279.10 | \$255,279.10 | \$255,279.10 |

  For PR MC submission Subcontractor Name If there are subconfracts, let all subcontractor names

  List Principal I-vestigators and Key Personne working on the project even if they receive no salary support.

  Subconneads/thisningsnam/schora i finanties if Baccopinger place Subcontractor name(s) or interarganizational Name/Code in Column B

  Consultarily II appropriate, place Consultant name(s) in Column B

  Remember that the total of all Bases and Option workshopes should equal the total of the Total Amount workshope.

  Add descriptions in Column B it costs are entered in the following areas. Funge Benefits, Latter Overticad, Other Direct Costs/other types. Material Handling. General and Administrative, and Facilities Gost of Money.
- Any proposed personnel in the technical proposal should be identified with their labor category.

  Profit or fee is not allowed on direct costs for travel. In addition, a DD Form 1547 (DFARS 215.404-70) will be utilized in calculating the Government objective for fee.
  This worksheet does not pull data from other worksheets within the Excel Spreadsheet.
- Note 1: Note 2: Note 3:

LABOR Information	Prime 1 (Fill-in)							
		Base Period	Option	Option II	Option III	Option IV	Supporting Documentation	Supporting Rationale
Labor Category (Note 1)	Name	Direct Labor Rate	Escalation Rate	Escalation Rate	Escalation Rate	Escalation Rate	(Note 2)	(Note 3)
Principal Scientist	Ross, Noam	\$55.29	\$58.05	\$0.00				
Research Scientist	Mendelsohn, Emma	\$40.87	\$42.91	\$42.91				
Senior Engineer	Young, Rob	\$50.48	\$53.00	\$53.00				

Indirect Rates								
	Base Period	Option	Option II	Option III	Option IV	Basis of Rate	Applied Against (Note 5)	Supporting Documentation (Note 6)
Rate Category	Rate	Rate	Rate	Rate	Rate	(Example: FPRA, FPRP, Estimate, etc.)		
Fringe Benefits	36.80%	36.80%	36.80%			Estimate	Direct labor costs	
Labor Overhead								
Material Handling								
General and Administrative	35.84%	35.84%	35.84%			Provisional (PROV)	Total direct costs	
Facilities Cost of Money								

Note 1: Add additional labor categories if needed.

Note 2: Provide the location (section, page) in the proposal's Cost Narrative/Supporting Documentation (BAA para 3.5.2.2.1) to support your price basis (e.g. copy of current payroll record or signed offer letter)

Note 3: Provide the location (section, page) in the proposal's Technical Basis of Estimate (BAA para 3.5.2.1.2) of the rationale for the appropriateness and necessity of the proposed labor categories and the labor hours allocated to

Note 4: Add additional indirect rates as needed.

If the offeror does not have a Forward Pricing Rate Agreement (FPRA), Forward Pricing Rate Recommendation (FPRR), or provisional billing rates; in order to assist the Government in evaluating the reasonableness of your proposed indirect rates, please provide the following information:

- 1. Performance data used to develop your proposed indirect rates. This typically consists of pool costs and base costs that demonstrate how the indirect rates were derived.
- 2. Information regarding your projections for out years, including your assumptions and method for developing these estimates.

#### Subcontractor

# SUBCONTRACTOR COST DATA Prime 1 (Fill-in)

Subcontracts/Interorganizational Transfers – A fully disclosed cost proposal as detailed as the Offeror's cost proposal including support documentation in accordance with FAR 15.404-3(b) and 15.404-3(c), the prime contractor shall perform and provide a cost/price analysis of each subcontractor's cost Certified cost or pricing data may be required for subcontractor proposals over \$2,000,000.00, in accordance with Section 811 of the National Defense Authorization Act for Fiscal Year 2018 (Pub. L. 115-91).

Subcontractor	CAGE Code	Competitive/Sole Source	Cost/price analysis included (Y/N)	Type of Subcontract (i.e., Fixed Price, Time and Materials, etc.)	HIACHMANISTIAN	Total amount exceeds \$750K (Y/N)	Cost/Price Analysis (Note 1)

Note 1: Provide the location (section, page) in the proposal's Cost Narrative/Supporting Documentation (BAA para 3.5.2.2.1) of the cost/price analysis for each individual subcontractor.

Offeror: Page 4 of 9 Pages

CONSULTANTS	Prime 1 (Fill-in)					
Name (Note 1)	Description of effort to be performed by the Consultant or attach Consultant Statement of Work	Number of Hours	Hourly Rate	Total	Supporting Documentation (Note 2)	Supporting Rationale (Note 3)
Base						
				\$0.00		
				\$0.00		
				\$0.00		
Option I						
				\$0.00		
				\$0.00		
				\$0.00		
				\$0.00		

· For Prime - fill out all columns if applicable. Indicate "Not Applicable" in B5 when there is no information to include.

Note 1: Include a separate section in the above table for the Base and each Option.

Note 2: Provide the location (section, page) in the proposal's Cost Narrative/Supporting Documentation (BAA para 3.5.2.2.1) used to support your price basis (e.g. copy of quote or agreement).

Note 3: Provide the location (section, page) in the proposal's Technical Basis of Estimate (BAA para 3.5.2.1.2) of the rationale for the necessity of the proposed consultant(s).

MATERIAL	LS - SUPPLIES (Note 1)		Prime 1 (Fill-	in)				
ltem (Note 2)	Description of Material (include model number)	Qty	Unit Price	Total Price	Competitive /Sole Source	Vendor/Source (If known)	Supporting Documentation (Note 3)	Supporting Rationale (Note 4)
Base								
1	Laptop computers	2	\$2,000.00	\$4,000.00				
Option I								

- · Fill out all columns on the Materials/Supplies Tab for each item.
- · Add additional lines if needed.
- Ensure that descriptions and vendors listed on the Materials/Supplies worksheet match information provided in the backup.
- Note 1: Material is property that may be incorporated into or attached to a deliverable end item or that may be consumed or expended in performing a contract. It includes assemblies, components, parts, raw and processed materials, and small tools and supplies that may be consumed in normal use in performing a contract. Material should be proposed separately from Equipment (see following spreadsheet and definitions).
- Note 2: Include a separate section in the above table for the Base and each Option.
  - Provide the location (section, page) in the proposal's Cost Narrative/Supporting Documentation (BAA para 3.5.2.2.1) of the documentation used to support
- Note 3: your price basis (e.g. copy of quote, page from catalog, recent invoice or PO). If the supporting documentation is located on a website, then please provide a
- Note 4: Provide the location (section, page) in the proposal's Technical Basis of Estimate (BAA para 3.5.2.1.2) for the necessity of the proposed materials/supplies

EQUIPMEN	IT (NOTE 1)	Prime 1 (Fill-in)								
Item (Nate 2)	Description of Equipment (include model number) (Note 1)	Will the equipment be included as part of a deliverable item under the award? (Y/N)	Type of Equipment (special test equipment, special tooling, general purpose equipment, or plant equipment) (Note 3)		Unit of Issue	Unit Price	Total Price	Vendor/Source	Supporting Documentation (Note 4)	Supporting Rationale (Note 5)
Base										
							\$0			
							\$0	·		·
Option I				, i						·
							\$0			

- · Fill out all columns on the Equipment Tab for each item.
- · Add additional lines if needed.
- . Ensure that descriptions and vendors listed on the Equipment worksheet match information provided in the backup.
- Note 1: Contractors are normally required to furnish all equipment and/or facilities necessary to perform Government contracts (see FAR 45.102(a)). The Government may allow
- Note 2: Include a separate section in the above table for the Base and each Option. Add additional lines labeled with Options if needed.
- Note 3: Definitions

performing a contract. It consists of items or assemblies of equipment including standard or general purpose items or components that are interconnected and interdependent for installing special test equipment, and which are of such a specialized nature that without substantial modification or alteration their use is limited to the development or Plant equipment means personal property of a capital nature (including equipment, machine tools, test equipment, furniture, vehicles, and accessory and auxiliary items) for

- Note 4: Provide the location (section, page) in the proposal's Cost Narrative/Supporting Documentation (BAA para 3.5.2.2.1) used to support your price basis (e.g. copy of quote, page from catalog, recent invoice or PO).
- Note 5: Provide the location (section, page) in the proposal's Technical Basis of Estimate (BAA para 3.5.2.1.2) of the rationale for the necessity of the proposed materials/supplies

		Prime 1	(Fill-in)											
	Trip Purpose	Number	Number of	Number	Air/Rail	Per Diem	Per Diem	Rental	Parking	Mileage	Тахі	Other	TOTAL	Supporting
City, State														
New York, NY	Conference Attendance	1	2	5	\$1,500.00	\$155.00	\$196.00	\$0.00	\$0.00	0	\$262.50	\$300.00		
Montpelier, France				Totals	\$3,000.00	\$1,395.00	\$1,568.00	\$0.00	\$0.00	\$0.00	\$525.00	\$600.00	\$7,088.00	
New York, NY	Annual Technical Review	1	1	5	\$350.00	\$71.00	\$161.00	\$0.00	\$0.00	0	\$262.50	\$0.00		
Alexandria, VA				Totals	\$350.00	\$319.50	\$644.00	\$0.00	\$0.00	\$0.00	\$262.50	\$0.00	\$1,576.00	
New York, NY	Conference Attendance	1	2	5	\$500.00	\$55.00	\$225.00	\$0.00	\$0.00	0	\$262.50	\$300.00		
Bozeman, Montana				Totals	\$1,000.00	\$495.00	\$1,800.00	\$0.00	\$0.00	\$0.00	\$525.00	\$600.00	\$4,420.00	
New York, NY	Annual Technical Review	1	1	5	\$350.00	\$71.00	\$161.00	\$0.00	\$0.00	0	\$262.50	\$0.00		
Alexandria, VA				Totals	\$350.00	\$319.50	\$644.00	\$0.00	\$0.00	\$0.00	\$262.50	\$0.00	\$1,576.00	
TOTAL													\$14,660	

- When completing the "From" and "To" information, enter the city and state where travel will originate and end. For example, from Seattle WA to Washington DC. Do not list airport codes.
- Include the trip purpose. It must be referred to in the Technical Proposal. Publications, conference attendance, and presentations are encouraged but must be justified to and approved by the Program Officer.
- · Remember that you cannot stay overnight the same number of days that you travel. There must always be one more day than night.
- · Do not combine any categories.
- · For lodging and M&IE, use federal Per Diem Rates found at http://www.defensetravel.dod.mil/site/perdiemCalc.cfm.
- For M&IE, input the full rate in the white block. The gray block will automatically calculate the total to include 75% for the first and last day of travel.

Estimates and the resultant costs claimed must conform to the applicable Federal cost principles.

Include a separate section in the above table for the base and each option

If there are miscellaneous expenses associated with the trip, provide description and rationale.

Provide the location (section, page) in the proposal's Cost Narrative/Supporting Documentation (BAA para 3.5.2.2.1) used to support your price basis (e.g. copy of quote, page from catalog, recent invoice or PO).

OTHER DIRECT COSTS (Note 1)			Prime 1 (Fill-in	)
Description (Note 2)	Qty	Unit of Issue	Unit Price	Total Price
Base				\$0
Publication costs	1	Publication	\$1,500	\$1,500
Cloud Services and Storage	1	Annual	\$2,000	\$2,000
Communication and Collaboration Services &	1	Annual	\$1,000	\$1,000
Option I				\$0
Publication costs	1	Publication	\$1,500	\$1,500
Cloud Services and Storage	1	Annual	\$2,000	\$2,000
Communication and Collaboration Services &	1	Annual	\$1,000	\$1,000

• If you have any ODC entries, be sure to fill out all columns.

• Ensure that descriptions and vendors listed on the ODC Details works

Examples include rental fees, shipping costs, license fees

Include a separate section in the above table for the Base and each

Note 1:

Note 2:

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

### Attachment 1 - Statement of Work (SOW)

Project Title: Predictin Biothreat Impacts from Early-Stage Data via Transfer Learning

Document Date: February 26, 2021

### **Objective**

The project's objectives are to determine the efficacy of machine learning (ML) techniques in predicting epidemic impacts in early-stage, data-sparse conditions, particularly novel biothreats, both intentional (e.g. novel genetically-modified pathogens) and natural (e.g. zoonotic). We define "early stage" as the initial period in an outbreak when only short time-series data are available, prior to rapid growth and geographic spread, and epidemiological parameters are not well-established. Specifically, we will answer three questions that will enable rapid decision-making based on ML models of epidemic impacts:

**Q1:** How much does partial-pooling improve predictions of epidemic impacts, and what information transfers best across diseases?

Q2: What variables are most predictive of epidemic impacts at early stages of an outbreak?

**Q3:** At what point in epidemic growth do hybrid models that incorporate mechanistic epidemiological components become more accurate than pure ML models?

This work will contribute to fundamental knowledge of properties and variables that contribute most to predictive accuracy at early-stage stages in emerging outbreak prediction and forecasting. It will enable the development of robust and deployable predictive models at early stages of outbreaks, and support informed choices for data collection and model selection.

# Scope

The awardee proposes a two-year study to determine the efficacy of ML techniques in predicting epidemic impacts in early-stage, data-sparse conditions, particularly novel biothreats. The awardee team shall focus on the following major goals and milestones:

- Year 1, Milestone 1 (0-6 months): Create and curate predictor database, project infrastructure, continuous integration/validation, establish data-imputation and automatic predictor workflow (Tasks 1.1-1.2).
- Year 1, Milestone 2 (6-12 months): Bayesian additive regression tree (BART) model fitting and tuning, determination of variable contribution to prediction accuracy (Task 1.3). Document and validate project code and infrastructure (Task 1.4)
- Option Year 1, Milestone 3 (12-18 months): Extend the database to include additional predictors, extend the BART model to ingest genetic data (Tasks 2.1-2.3)
- Option Year, Milestone 4 (18-24 months): Construction of hybrid ML/mechanistic model structure, comparative testing of pure and hybrid version, determination of timing when current data overtakes transfer data (Task 2.4). Document and validate project code and infrastructure (Task 2.5)
- Option Task 1, (12-18 months): Develop a graphical user interface (GUI) for the model, allowing users to input of early-stage outbreak data and generate predictions, uncertainty bounds, and interpretable explanations of influential variables. Document GUI and validate code.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

#### Background

Situational awareness and risk assessment are critical at early stages of disease outbreaks, when interventions can best mitigate adverse outcomes. Yet decision-makers often face a paucity of data at these stages. Traditional modeling methods to forecast disease impact severity rely heavily on prior knowledge or assumptions about mechanisms of disease spread progression, and they frequently do not anticipate the actual mechanisms.<sup>1-3</sup> They also take considerable time and effort to develop.

Recent theory suggests disease dynamics are fundamentally predictable<sup>4</sup>, and ML methods have been successfully applied in emerging disease science. ML refers to a group of methods, ranging from simple nearest-neighbor approaches to function-approximation methods such as boosted trees and penalized splines, to complex deep-learning neural networks. ML has been used to predict the likelihood of disease emergence and the zoonotic potential of various hosts and viruses<sup>5,6</sup>, and deep-learning approaches have been used to infer viral hosts from the comparatively richer data of genetic sequences of viruses.<sup>7,8</sup>

ML methods are generally unstructured, flexible, and offer greater predictive power over traditional statistical approaches. However, these methods are data-intensive, and a key challenge with ML development is data paucity and/or a low signal-to-noise ratio. Surveillance data for even existing, well-studied diseases is rarely rich enough for the data scale of many modern ML techniques. For instance, a ML approach to predict impacts from a new outbreak of Ebola would draw training data from fewer than twenty previous events, each with varying levels of data coverage.

Transfer learning, or data fusion, provides a solution to data paucity issues by using data from other domains to train parts of models that are general, thus limiting new data requirements for more specific components. Hierarchical modeling, or *partial pooling*, is a transfer learning approach for partitioning out which data can be shared between models and which are only applicable to a specific case. For instance, Ebola forecasts for countries outside the historic range of the disease may be improved by calibrating against Cholera outbreaks, which have occurred both within and outside countries with Ebola outbreaks. The degree to which the disease outcomes are correlated is learned, rather than assumed.

An alternative recent development in disease forecasting is the development of hybrid models that incorporate simple mechanistic structures and predictive ML methods to learn model parameters. These approaches have demonstrated high predictive accuracy for near-term forecasting of epidemic growth and impact, particularly during the COVID-19 pandemic<sup>9</sup>. However, these approaches require adequate time-series of data to project forward from current conditions. Their utility in early stages and long-term forecasting accuracy has yet to be determined. In opting to apply these models, it is critical to understand these data requirements and at what point in an outbreak hybrid approaches can overtake pure ML early-stage predictions in accuracy. This project seeks to develop, apply, and compare pure and hybrid-ML techniques to enable the development of robust, rapidly usable predictions at early stages of outbreaks.

Key references (additional references can be found in the Technical Proposal):

1 MacDiarmid, S. C. et al. Handbook On Import Risk Analysis for animals and animal products: quantitative risk assessment. Vol. 2 (Office International des Épizooties, 2004).

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

- Moura, J. A., McManus, C. M., Bernal, F. E. M. & de Melo, C. B. An analysis of the 1978 African swine fever outbreak in Brazil and its eradication. *Rev. Sci. Tech.* **29**, 549-563, doi:10.20506/rst.29.3.1992 (2010).
- Delgado, J. *et al.* U.K. Foot and Mouth Disease: A Systemic Risk Assessment of Existing Controls. *Risk Anal.* 37, 1768-1782, doi:10.1111/risa.12704 (2017).
- Scarpino, S. V. & Petri, G. On the predictability of infectious disease outbreaks. *Nat. Commun.* **10**, 898, doi:10.1038/s41467-019-08616-0 (2019).
- Olival, K. J. *et al.* Host and viral traits predict zoonotic spillover from mammals. *Nature*, doi:10.1038/nature22975 (2017).
- 6 Allen, T. *et al.* Global hotspots and correlates of emerging zoonotic diseases. *Nat. Commun.* **8**, 1124, doi:10.1038/s41467-017-00923-8 (2017).
- Mock, F., Viehweger, A., Barth, E. & Marz, M. VIDHOP, viral host prediction with Deep Learning. *Bioinformatics* (2019).
- 8 Babayan, S. A., Orton, R. J. & Streicker, D. G. Predicting reservoir hosts and arthropod vectors from evolutionary signatures in RNA virus genomes. *Science* **362**, 577-580, doi:10.1126/science.aap9072 (2018).
- 9 Gu, Y. *COVID-19 Projections Using Machine Learning*, <a href="https://covid19-projections.com/about/#historical-performance">https://covid19-projections.com/about/#historical-performance</a> (2020).

#### Data Sources

Fitting and testing models will draw from disease outbreak data (counts, timing, and geography) and predictor data including pathogen traits (disease symptoms, pathogen traits, modes of transmission), pathogen sources (point of origin, geographic range of potential natural hosts), environmental context (climatic conditions, ecosystem traits), and socioeconomic context (local population densities, demographics, transportation networks, healthcare systems). Please see the Technical Proposal for available data sources.

# Tasks/Scientific Goals (Format: Year.Task.Subtask)

#### YEAR 1

# Task 1.1 – Develop and deploy project infrastructure

The awardee shall develop a computational environment that enables continuous model development and testing and allows for research and models to be reproduced and extended by external teams. The computational environment will include continuous maintenance and documentation of project codebase.

#### Subtasks

- 1.1.1 Layout a provision internal project network
- 1.1.2 Set up continuous integration pipelines
- 1.1.3 Implement backup and data verification jobs
- 1.1.4 Implement differential privacy testing

# Task 1.2 – Engineer project database

The awardee shall curate, integrate, cross-link, and validate relevant datasets from various sources into a project database, accounting for differing space and time scales, data formats, and data completeness.

#### Subtasks

- 1.2.1 Design database schema
- 1.2.2 Write pipelines to ingest data
- 1.2.3 Cross-link data sources via common ontology

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

- 1.2.4 Write quality assurance/quality control (QA/QC) tests for data and perform data cleaning
- 1.2.5 Create automatic extraction routines to populate relevant predictors from location information
- 1.2.6 Update data from live sources and clean and maintain data continuously

# Task 1.3 – Machine Learning (ML) model development and testing

The awardee shall design, implement, test, and analyze performance of ML model structures and iteratively improve them.

Subtasks

- 1.3.1 Establish quantitative and qualitative metrics of model performance
- 1.3.2 Implement "practical naïve" model baseline
- 1.3.3 Design Bayesian additive regression tree (BART) model structure
- 1.3.4 Design alternative generalized additive model (GAM) structure
- 1.3.5 Implement within-disease, all-disease, and partial-pooled disease model versions
- 1.3.6 Measure variable contribution by re-fitting
- 1.3.7 Analyze prediction-level variable contributions and model performance
- 1.3.8 Iteratively update models to improve performance

# Task 1.4 – Maintain and document project infrastructure

The awardee shall document the methodology for database schema, data ingestion pipelines,  $Q\Lambda/QC$  tests, extraction routes, and ML models established in Tasks 1.1-1.3. *Subtasks* 

- 1.4.1 Write and maintain initial codebase documentation
- 1.4.2 Perform code review, testing, and validation

# **OPTION YEAR (OPTION I):**

#### Task 2.1 – Extend project database

In the option year, the awardee shall extend the database to include additional predictors and continuously maintain and update to accumulate more data from upstream sources.

Subtasks

- 2.1.1 Extend database schema to include genetic data and auto-encoded variables
- 2.1.2 Write pipelines to ingest genetic data
- 2.1.3 Cross-link data sources via common ontologies
- 2.1.4Write QA/QC tests for data and perform data cleaning
- 2.1.5Update data from live sources and clean and maintain data continuously

## Task 2.2 – Extend BART model

In the option year, the awardee shall extend the BART model to ingest genetic data and test the contribution of this data in enhancing or recapitulating pathogen trait data.

Subtasks

- 2.2.1Develop genetic predictor variable set via neural-network autoencoding
- 2.2.2 Modify BART and GAM structures to incorporate genetic variables
- 2.2.3 Implement within-disease, all-disease, and partial-pooled disease model versions
- 2.2.4 Compare performance of genetic, pathogen-trait, and combined models

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

Task 2.3 – Hybrid ML-mechanistic model development and testing

In the option year, the awardee shall develop the hybrid machine-learning mechanistic model, compare its performance to a pure ML model, and determine the switch-over time when the mechanistic model becomes more predictive in the course of the outbreak.

Subtasks

- 2.4.1 Establish quantitative and qualitative metrics of model performance
- 2.4.2 Design mechanistic model structure
- 2.4.3 Determine mechanistic model prior forms
- 2.4.4 Modify ML model to predict mechanistic parameters rather than outcomes
- 2.4.5Implement within-disease, all-disease, and partial-pooled disease model versions
- 2.4.6Measure variable contribution by re-fitting
- 2.4.7 Analyze prediction-level variable contributions and model performance using criteria developed in Task 1.3
- 2.4.8Compare Hybrid and ML performance along outbreak time-series progression
- 2.4.9Iteratively update models to improve performance

# Task 2.4 – Maintain and document project infrastructure

In the option year, the awardee shall document the methodology for extended database schema, and data ingestion pipelines,  $Q\Lambda/QC$  tests, extraction routes, and ML models established in Tasks 2.1-2.3.

Subtasks

- 2.4.1 Write and maintain initial codebase documentation
- 2.4.2 Perform code review, testing, and validation

# **OPTION TASK (OPTION II):**

# Task 3.1 – Develop Model Graphical User Interface

As an additional optional task, the awardee shall develop a graphical user interface (GUI) for the ML model developed in task 1.3, allowing users to input of early-stage outbreak data and generate predictions, uncertainty bounds, and interpretable explanations of influential variables.

Subtasks

- 3.1.1 Port model into portable framework usable for standalone or web-based interaction
- 3.1.2 Optimize for rapid response time and application size
- 3.1.3 Design input interface
- 3.1.4 Design visualizations for predictions, uncertainty, and variable importance

# Task 3.2 – Maintain and document project infrastructure

In the option task, the awardee shall document usage of the GUI, and perform code review of testing and validation of the GUI codebase created in tasks 3.1

Subtasks

- 3.2.1 Create GUI documentation
- 3.2.2 Perform code review, testing, and validation

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

#### Deliverables

- Reports and Documents
  - Monthly reports on task status
  - Monthly cost statement
  - Periodic program reviews (virtual and/or in person)
  - Annual technical report, which shall include
    - Executive Summary
    - I. Introduction/Background
    - II. Methodology
    - III. Findings
    - IV. Discussion/Future Recommendations
    - V. Conclusion
    - VI. Appendices
      - a. Snapshots of GUI/how to use
      - b. Data worksheet (summarizing list of data sources used, metadata abstracted, links, etc.)
- Source and executable software code
- Project databases
- Technical papers describing algorithm design and performance
- Conference presentations

#### Code Modules and Database Archives

Code and databases will be made available for download and provided electronically via DoD SAFE and/or on-disk to DTRA at the end of the period of performance.

Note: The data and information collected in this effort is funded on the condition that it will not be released to another nation or international organization without the specific authority from the Defense Threat Reduction Agency. Individual or corporate rights originating in the information, whether patented or not, will be respected; that the recipient government will report promptly to the Defense Threat Reduction Agency any known or suspected compromise and that the information will be provided substantially the same degree of security afforded it by the Department of Defense of the United States. No U.S. Government commitment to sell, lend, lease, co-develop, or co-produce defense articles is implied or intended. Also, regardless of any other markings on the document, it will not be downgraded or declassified without written approval of the originating U.S. agency.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

# Gantt Chart

Description	Y1 Q1-2	Y1 Q3-4	Y2 Q1-2	Y3 Q3-4
Year 1, Milestone 1				
1.1.1 Layout a provision project network				
1.1.2 Set up continious integration pipelines				
1.1.3 Implement backup and data verification jobs				
1.1.4 Implement differential privacy testing				
1.2.1 Design database schema				
1.2.2 Write pipelines to ingest data				
1.2.3 Cross-link data sources via common ontology				
1.2.4 Write quality assurance/quality control (QA/QC) tests for data and perform data cleaning				
1.2.5 Create automatic extraction routines to populate relevant predictors from location information				
1.2.6 Update data from live sources and clean and maintain data continuously				
1.3.1 Establish quantitative and qualitative metrics of model performance				
1.3.2 Implement "practical naïve" model baseline				
1.3.3 Design model structure				
1.3.4 Design alternative structure				
1.3.5 Implement within-disease, all-disease, and partial-pooled disease model versions				
1.3.6 Measure variable contribution by re-fitting				
1.3.7 Analyze prediction-level variable contributions and model performance				
1.3.8 Iteratively update models to improve performance				
1.4.1 Write and maintain initial codebase documentation				
1.4.2 Perform code review, testing, and validation				
Option Year 1 (Option I)				
2.1.1 Extend database schema to include genetic data				
	<del> </del>			
2.1.2 Write pipelines to ingest genetic data 2.1.3 Cross-link data sources via common entologies				
2.1.4 Write QA/QC tests for data and perform data cleaning				
2.1.5 Update data from live sources and clean and maintain data continuously				
2.2.1 Develop genetic predictor variable set via neural-network autoencoding	1			
2.2.2 Modify BART and GAM structures to incorporate genetic variables				
2.2.3 Implement within-disease, all-disease, and partial-pooled disease models				
·				
2.2.4 Compare performance of genetic, pathogen-trait, and combined models				
2.4.1 Establish quantitative and qualitative metrics of model performance				
2.4.2 Design mechanistic model structure				
2.4.3 Determine mechanistic model prior forms     2.4.4 Modify ML model to predict mechanistic parameters rather than outcomes				
2.4.5 Implement within-disease, all-disease, and partial-pooled disease model versions	-			
2.4.6 Measure variable contribution by re-fitting				
2.4.7 Analyze prediction-level variable contributions and model performance				
2.4.8 Compare Hybrid and ML performance along outbreak time-series progression	-			
2.4.9 Iteratively update models to improve performance				
2.5.1 Write and maintain initial codebase documentation				
2.5.2 Perform code review, testing, and validation	<del> </del>			
Option Task (Option II)				
3.1.1 Port model into portable framework usable for standalone or web-based interaction				
3.1.2 Optimize for rapid response time and application size	-			
3.1.3 Design input interface				
3.1.4 Design visualizations for predictions, uncertainty, and variable importance	-			
3.2.1 Create GUI documentation	-			
3.2.2 Perform code review, testing, and validation				

# RESEARCH & RELATED BUDGET - Budget Period 1

OMB Number: 4040-0001 Expiration Date: 12/31/2022

ORGANIZATIO	ONAL DUNS:	077090066	cccc	Enter name of Or	ganization:	Ecolie:	alth Al	liance				
Budget Type:	⊠ Project	Subawa	ard/Consortium	ו		Budget	Period:	1 S	tart Date:	07/01/2020	End Date: 06/30/202	1
A. Senior/Key	/ Person											
Prefix	First	Middle	Last	Suffix	Bas	e Salary (S	s) C	Month al. Acad	•	Requested Salary (\$)	Fringe Benefits (\$)	Funds Requested (\$)
Dr.	Noam		Ross				3.00			25,000.0	9,200.00	34,200.00
Project Role	: PD/PT											
	Robert		Yeung				6.00			52,500.0	0 19,320.00	71,820.00
Project Role	: Senier Engi	neer										
	Emma		Mendelsohn				6.00			42,500.0	0 15,640.00	58,140.00
Project Role	: Research Sc	ientist										
Additional Senior Key Persons: Add Attachment								requested for all Senior sons in the attached file				
										7	otal Senior/Key Person	164,160.00
B. Other Pers	onnel											
Number of Personnel	Project	Role				Cal.	Months Acad.	Sum.		quested	Fringe Benefits (\$)	Funds Requested (\$)
	Post Doctoral	Associates										
	Graduate Stud	dents			ĺ							
	Undergraduate	e Students										
	Secretarial/Cle	erical			[							
	Total Number C	Other Personr	nel								Total Other Personnel	
								Total S	Salary, W	ages and Fri	nge Benefits (A+B)	164,160.00

C. E	Equipment Description		
	items and dollar amount for each item exceeding \$5,000 Equipment item		Funds Requested (\$)
Additi	itional Equipment:	dd Attachment	
	Total funds requested for all equipm	nent listed in the attached file	
		Total Equipment	
D. Tr	[ravel		Funds Requested (\$)
1. [	Domestic Travel Costs (Incl. Canada, Mexico and U.S. Possessions)		1,576.00
2. F	Foreign Travel Costs		7,088.00
		Total Travel Cost	8,664.00
E. Pa	Participant/Trainee Support Costs		Funds Requested (\$)
1, 1	Tuition/Fees/Health Insurance		
2. 9	Stipends		
<b>3</b> . 1	Travel		
4. 5	Subsistence		
5. (	Other		
	Number of Participants/Trainees Total Parti	cipant/Trainee Support Costs	

F.	Other Direct Costs			Funds Requested (\$)
1.	Materials and Supplies			7,000.00
2.	Publication Costs			1,500.00
3.	Consultant Services			
4.	ADP/Computer Services			
5.	Subawards/Consortium/Contractual Costs			
6.	Equipment or Facility Rental/User Fees			
7.	Alterations and Renovations			
8.				
9.				
10.				
			Total Other Direct Costs	8,500.00
G. I	Direct Costs			Funds Requested (\$)
		Total Dir	rect Costs (A thru F)	181,324.00
<u>H. I</u>	ndirect Costs			
	Indirect Cost Type	Indirect Cost Rate (%)	Indirect Cost Base (\$)	Funds Requested (\$)
	EcoHealth Alliance's NICA rate	35.84	181,324.00	64,986.52
_			Total Indirect Costs	64,986.52
	nizant Federal Agency ncy Name, POC Name, and			
POC	Phone Number)  Department of Defense, De	partment of the Navy	, Sharon Gales, (1.763	. עככא. טעט.
<u>I. Te</u>	otal Direct and Indirect Costs			Funds Requested (\$)
	Total Direc	t and Indirect Institu	tional Costs (G + H)	246,310.52
<u>J. F</u>	ee		1	Funds Requested (\$)
<u>K. 1</u>	Total Costs and Fee		<b></b>	Funds Requested (\$)
		Total (	Costs and Fee (I + J)	246,310.52
<u>L. E</u>	Budget Justification			
(Onl	y attach one file.) 1234 DTRA MI Budget Just if ignt	ior.pdf	Delete Attachmer	nt View Attachment

## RESEARCH & RELATED BUDGET - Budget Period 2

OMB Number: 4040-0001 Expiration Date: 12/31/2022

ORGANIZATI	ONAL DUNS:	077090068	cccc	Enter name of Org	janization:	Ecolic	alth All	iance				
Budget Type:	: Project	Subaw	ard/Consortiun	n		Budget	Period:	2 Sta	ırt Date: 67	7/01/2021	End Date: 06/30/2022	
A. Senior/Ke	y Person											
Prefix	First	Middle	Last	Suffix	Base	e Salary (	\$) Ca			Requested Salary (\$)	Fringe Benefits (\$)	Funds Requested (\$)
Dr.	Noam		Ross				3.00			26,250.00	9,660.00	35,910.00
Project Role	e: [PD/PI											
	Robert		Young				6.00			55,125.00	20,286.00	75,411.00
Project Role	Senior Engin	neer										
	Emma		Mendelsohn				6.00			44,605.00	16,422.00	61,047.00
Project Role	: Research Sci	entist							•			
Additional Senio	or Key Persons:			Add	Attachment					Key Pers	ons in the attached file	
		Budget Period: 2 Start Date: 67/61/2621 End Date: 66/36/2622    Months   Requested   Salary (\$)   Requested (\$)   Requeste										
B. Other Per	sonnel											
Number of Personnel	Project F	Role						Sum.				
	Post Doctoral A	Associates			[							
	Graduate Stude	ents			Ī							
	Undergraduate	Students			Ī							
	Secretarial/Cle	rical										
	Total Number O	ther Person	nel								Total Other Personnel	
								Total Sa	alary, Wag	es and Fri	nge Benefits (A+B)	172,368.00

C. Ec	quipment Description		
	items and dollar amount for each item exceeding \$5,000 Equipment item		Funds Requested (\$)
Additi	tional Equipment:	Add Attachment	
	Total funds requested for	all equipment listed in the attached file	
		Total Equipment	
D. Tr	ravel		Funds Requested (\$)
1. [	Domestic Travel Costs (Incl. Canada, Mexico and U.S. Poss	essions)	5,996.00
2. F	Foreign Travel Costs		
		Total Travel Cost	5,996.00
E. Pa	articipant/Trainee Support Costs		Funds Requested (\$)
<b>1</b> . T	Tuition/Fees/Health Insurance		
<b>2.</b> S	Stipends		
<b>3</b> . T	Travel		
4. 5	Subsistence		
5. (	Other		
	Number of Participants/Trainees	Total Participant/Trainee Support Costs	

F.	Other Direct Costs	Funds Requested (\$)
1.	Materials and Supplies	3,000.00
2.	Publication Costs	1,500.00
3.	Consultant Services	
4.	ADP/Computer Services	
5.	Subawards/Consortium/Contractual Costs	
6.	Equipment or Facility Rental/User Fees	
7.	Alterations and Renovations	
8.		
9.		
10.		
	Total Other Direct Costs	4,500.00
<u>G. I</u>	Direct Costs	Funds Requested (\$)
	Total Direct Costs (A thru F)	182,864.00
<u>H. I</u>	Indirect Cost Type Indirect Cost Rate (%) Indirect Cost Base (\$)	Funds Requested (\$)
	EceHealth Alliance's NICA rate 35.84 182,864.00	65,538.46
	Total Indirect Costs	65,538.46
	inizant Federal Agency	
	Phone Number)  Department of Defense, Department of the Navy, Sharon Gales, (1.763)	1.696.8559
I. T	otal Direct and Indirect Costs	Funds Requested (\$)
	Total Direct and Indirect Institutional Costs (G + H)	248,402.46
J. F	Fee	Funds Requested (\$)
		,
κ. :	Total Costs and Fee	Funds Requested (\$)
	Total Costs and Fee (I + J)	248,402.46
<u>L. E</u>	Budget Justification	
(Onl	y attach one file.) 1234 DTRA MI Budger Tust i Cigar ion and C	

# RESEARCH & RELATED BUDGET - Cumulative Budget

Ocation A Oct. Nr	Totals	(\$)
Section A, Senior/Key Person		336,528,00
Section B, Other Personnel		
Total Number Other Personnel		
Total Salary, Wages and Fringe Benefits (A+B)		336,528.00
Section C, Equipment	Ī	333,333,000
Section D, Travel		14,660.00
1. Domestic	7,572.00	
2. Foreign	7,088.00	
Section E, Participant/Trainee Support Costs		
1. Tuition/Fees/Health Insurance		
2. Stipends		
3. Travel		
4. Subsistence		
5. Other		
6. Number of Participants/Trainees		
Section F, Other Direct Costs		13,000.00
1. Materials and Supplies	10,000.00	13,000.00
2. Publication Costs	3,000.00	
3. Consultant Services	3,000.00	
4. ADP/Computer Services		
5. Subawards/Consortium/Contractual Costs		
6. Equipment or Facility Rental/User Fees		
7. Alterations and Renovations		
8. Other 1		
9. Other 2		
10. Other 3		
Section G, Direct Costs (A thru F)		
Section H, Indirect Costs	<u> </u>	364,188.CC
Section I, Total Direct and Indirect Costs (G + H)		130,524.98
Section J, Fee		494,712.98
Section K, Total Costs and Fee (I + J)		
•		494,712.98

## BUDGET JUSTIFICATION FOR ECOHEALTH ALLIANCE

EcoHealth Alliance requests a total of \$4,975,681 over all-years of the proposed project to support personnel, travel, equipment, consortium agreements, and applicable indirect costs.

## A. Key Personnel

Noam Ross, Ph D., Principal Investigator (3.0 calendar months for all years). Dr. Ross will oversee the project and be responsible for project management, research, development, design, analysis, and manuscript writing and conference presentations, as well as communication with DTRA. We request \$25,000 in Y1 due to the high cost of living in New York City and a 5% cost of living increase in OY1.

Emma Mendelsohn, M.Sc, Research Scientist. (6.0 calendar months for all years). Ms. Mendelsohn will lead data engineering and contribute expertise to algorithm development. She will contribute to manuscript writing and conference presentations. We request \$42,500 in Y1 with a 5% cost of living increase in OY1

Robert Young M.Sc, Senior Research Software Engineer. (6.0 calendar months for all years). Mr. Young will lead infrastructure engineering and contribute expertise to algorithm development. He will contribute to manuscript writing and conference presentations. We request \$52,500 in Y1 with a 5% cost of living increase in OY1.

### Fringe Benefits

Fringe benefits are calculated as 36.80% of base salary p.a. with \$44,160 requested in Y1 calculated from the base salary for all personnel. In OY1, we budget for a 5% per year cost of living allowance increase in all salaries.

#### B. Other Personnel

No funds are requested for other personnel.

### C. Equipment

No funds are requested for equipment.

### D. Travel

Per recommendations in the BAA, we request travel funds to Nationally Recognized Conferences and the Annual Technical Review.

## Domestic Travel

Domestic travel is requested for one trip per year from New York City per year to Northern Virginia the Annual Technical Review. Transportation is estimated at \$350 per trip along with the government GSA per diem rates for Northern Virginia (\$161 for accommodation and \$71 for meals and incidentals). Additional domestic travel support will facilitate two EcoHealth Alliance team members presenting at one conference. In OY1, this will be the EEID conference held in Bozeman, MT in 2022, with transportation costs of \$500, \$225 for accommodation and \$55 for meals and incidentals. For each trip, \$25 per day is estimated in taxi costs and for those that

require transport to the airport \$150 RT is included. In total, we are requesting \$1,576 in Y1 and \$5,996 in OY1 for domestic travel

#### International Travel

One international trip for two team members to present at the EEID conference held in Montpelier, France, in 2021 requested for Y1. Flights are estimated at \$1500 and the federal per diem rates of \$196 for lodging and \$155 for meals in Montpelier. \$25 per day is estimated in taxi costs plus \$150 RT trips to airport is included. In total we are requesting \$7,088 in Y1.

## E. Participant/Trainee Support Costs

No funds are requested for participant or trainee support costs.

#### F. Other Direct Costs

Materials and Supplies

PI Ross and Research Scientist Mendelsohn will both need a new computer, equipped with Microsoft software for which we request a total of \$4,000, in only \$11.

## Cloud Computing and Storage Services

We will make use of cloud computer services including spot server rental for compute-intensive jobs, data storage, continuous integration and testing of computer code. We request \$2,000 per year in Y1-OY1.

### Team Communication and Collaboration Software and Services

We request \$1,000 per year for subscriptions for software and services for document sharing, chat, voice, and video communications.

### Publication Costs

To facilitate the dissemination of project findings, \$1,500 per year in only Y1-OY1 is requested for open access publication costs in international peer-reviewed journals.

#### H. Indirect Costs

We are requesting the EcoHealth Alliance federally approved indirect cost rate of 35.84% on all applicable direct costs. The USA Department of Defense's Department of the Navy has approved this rate on 01 April 2020. We request \$ \$64,986.52 in Y1 and \$65,538.46 in OY1.

PI: Noam Ross

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

## Predicting Biothreat Impacts from Early-Stage Data via Transfer Learning

### Phase II Technical Proposal

#### ABSTRACT

Accurately forecasting the impacts of novel biological agents in the early stages of their emergence is critical to planning control measures for warfighters and civilians. However, at the beginning of outbreaks, data about pathogen properties and dynamics are often scarce. Machine learning (ML) techniques generally require large data sets, thus hampering their utility in these contexts. Our proposed work addresses this challenge by using cross-disease prediction via transfer learning utilizing data from previous emergence events to improve early-stage predictions for future events. We will use partial-pooling to determine the degree to which information can be shared across diseases according to hierarchical similarities in pathogen and host properties, and in environmental and population conditions. We will also determine which properties contribute most to predictive accuracy, providing guidance for prioritizing datacollection in early outbreak stages. We will test and compare two approaches – pure ML predictions and hybrid ML/mechanistic models – to determine which provides better predictions of outbreak size and impacts. We will determine at what point in an outbreak current data overtakes historical data as a better predictor of outbreak dynamics. Making use of data sets from multiple host species as well as efficient model-validation techniques, we will expand the size of available training data and thus models' predictive capacity.

#### **SCOPE**

#### Objective

Our primary research objectives are to determine the efficacy of ML techniques in predicting epidemic impacts in early-stage, data-sparse conditions, particularly novel biothreats both intentional (e.g. novel genetically-modified pathogens) and natural (e.g. zoonotic). Specifically, we will answer three questions that will enable rapid decision-making based on ML models of epidemic impacts:

Q1: How much does partial-pooling improve predictions of epidemic impacts, and what information transfers best across diseases?

Q2: What variables are most predictive of epidemic impacts at early stages of an outbreak?

**Q3:** At what point in epidemic growth do hybrid models that incorporate mechanistic epidemiological components become more accurate than pure ML models?

This work will contribute to fundamental knowledge of properties and variables that contribute most to predictive accuracy at early-stage stages in emerging outbreak prediction and forecasting. It will enable the development of robust and deployable predictive models at early stages of outbreaks, and support informed choices for data collection and model selection.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

## **Background**

Situational awareness and risk assessment are critical at early stages of disease outbreaks, when interventions can best mitigate adverse outcomes. Yet decision-makers often face a paucity of data at these stages. Traditional modeling methods to forecast disease impact severity rely heavily on prior knowledge or assumptions on mechanisms of disease spread progression, and they frequently do not anticipate the actual mechanisms.<sup>1-3</sup> They also take considerable time and effort to develop.

Recent theory suggests disease dynamics are fundamentally predictable<sup>4</sup>, and machine learning (ML) methods have demonstrated utility for some aspects of emerging disease science, e.g. to predict the likelihood of disease emergence and zoonotic potential of various hosts and viruses<sup>5,6</sup>, and deep learning has been used to infer viral hosts from the comparatively richer data of genetic sequences of viruses.<sup>7,8</sup>

Machine learning (ML) refers to a group of methods, ranging from simple nearest-neighbor approaches to function-approximation methods such as boosted trees and penalized splines, to complex deep-learning neural networks. These methods are generally unstructured, flexible, and offer greater predictive power over traditional statistical approaches. However, these methods can also be data-intensive; a key challenge with ML methods development is that data are too sparse, or signal/noise ratio too low, to effectively train algorithms. Surveillance data for even existing, well-studied diseases is rarely rich enough for the data scale of many modern machine learning or deep learning techniques. For instance, an ML approach to predict impacts from a new outbreak of Ebola would have training data from to less than twenty previous events for which there is comprehensive epidemiological data from 10 West and Central African nations

Transfer learning, or data fusion, provides a solution for using data from other domains to train parts of models that are general, thus limiting new data requirements for more specific components. Hierarchical modeling, or *partial pooling*, is a transfer learning approach for partitioning out which data can be shared between models and which are only applicable to a specific case. For instance, Ebola forecasts from countries outside of the virus' historic range may be improved by calibrating against influenza outbreaks, which have occurred both within and outside countries with Ebola outbreaks. The degree to which the disease outcomes are correlated (e.g., similarity of modes of transmission) are learned, rather than assumed.

An alternative recent development in disease forecast is the development of hybrid models that incorporate simple mechanistic structures and predictive ML methods to learn model parameters. These approaches have demonstrated high predictive accuracy for near-term forecasting of epidemic growth and impact, particularly during the COVID-19 pandemic<sup>9</sup>. However, these approaches require adequate time-series of data to project forward from current conditions. Their utility in early stages and long-term forecasting accuracy has yet to be determined. In opting to apply these models, it is critical to understand these data requirements and at what point in an outbreak hybrid approaches can overtake pure ML early-stage predictions in accuracy.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

We will develop algorithms to predict the dynamics and impacts of outbreaks of new infectious diseases, based on data available at early stages of outbreaks. In doing so, we will answer the following key questions.

- How much does partial-pooling improve predictions of epidemic impacts, and what information transfers best across diseases?
- What variables are most predictive of epidemic impacts at early stages of an outbreak?
- At what point in epidemic growth do hybrid models that incorporate mechanistic epidemiological components become more accurate than pure ML models?

### Algorithmic Methods:

We will test the ability to ML methods incorporating partial pooling to predict impacts from a new outbreak of disease. Our primary model structure will be built around Bayesian additive regression trees (BARTs)<sup>10</sup>. BARTs provide three features appropriate for this application: First, they can provide both high-quality predictions as well as robustness in relatively data-poor scenarios. Secondly, they have explicit uncertainty structures is better able to characterize error than other regression tree methods that do not. Thirdly, they are able to incorporate hierarchical partial pooling. In some scenarios where data are smooth rather than categorical, generalized additive models (GAMs) can offer similar performance and similar features<sup>11</sup>, and we will test these as an alternate formulation.

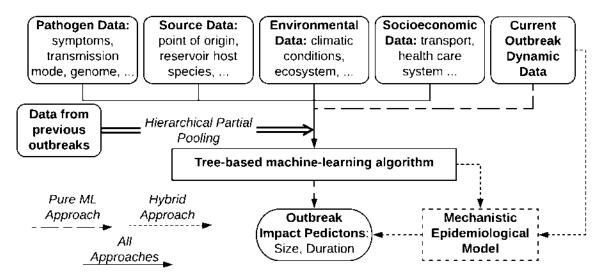
We will parametrize the models using data on pathogen traits (disease symptoms, pathogen traits, modes of transmission), pathogen sources (point of origin, geographic range of potential natural hosts), environmental context (climatic conditions, ecosystem traits), and socioeconomic context (local population densities, demographics, transportation networks, healthcare systems), as well as time-series data of infections from the earliest phase of the outbreaks. Not all these data are typically available at the early stages of a new bioagent outbreak, and are in many cases incomplete even in the case of historical outbreaks. Thus, we will use multiple imputation<sup>12</sup> in all models to allow for model fitting, prediction and assessment of uncertainty under incomplete data.

We will explore multiple model structures, but all formulations will be fit and tested under three conditions: within-disease forecasting where models are fit only on data from a single disease and tested at ability to predict impacts from that disease, all-disease forecasting where models are fit on data from all diseases and tested against ability to predict each disease, and partial-pooled where models are fit all models from all diseases but parameter sharing across diseases are partially pooled. (See 'Model Evaluation and Validation' below)

While ML approaches are best for predictive power, they do not have the flexibility to be used for alternate scenarios or intervention evaluation that traditional compartmental models do. We will compare "pure" ML algorithms to a hybrid approach using ML to generate parameters for mechanistic epidemiological models. In this hybrid case, we will use the ML model to predict prior parameters for compartmental epidemiological model using the same data as used in the

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

pure ML model, excluding early time-series data. The model be structured to have high flexibility under small number of parameters – i.e. SIR-type model with variable contact structure and latent/infectious periods. This model will then be fit to the time series data for the outbreak under the constraint of these prior parameters. Using this approach (see figure), we will determine whether the structural knowledge embedded in the epidemiological model improves outbreak predictions or hinders them due to assumptions. As with the pure ML approach, we will test the hybrid model approach under *within-disease*, *all-disease*, *and partial-pooled* approaches.



**Figure**: Schematic of machine-learning approaches. ML algorithm ingests biothreat condition data and makes predictions based on training from previous outbreak data. In pure ML approach, current outbreak dynamics are used as ML predictors of outbreak impacts. In hybrid approach, ML algorithm predicts parameters of an epidemiological model, which predicts impacts using parameters and current outbreak dynamics.

Model variables will be analyzed at both model-wide and individual case level to determine their contribution to performance. First, to evaluate model-wide predictive performance, we will re-fit models successively dropping each predictor to determine which provide the greatest improvement in performance. At the individual case level, we will calculate Shapley Additive Explanation (SHAP) scores<sup>13</sup> for all variables. This will allow for inspection of the role of variables across major categories (pathogen type, continent), and determine how accuracy varies across these contexts. Clusters of similar SHAP scores will also allow us to identify clusters of diseases or conditions that have similar behavior or performance accuracy. Such information will ultimately support decision-making in selecting models or variables to select for prediction based on the context of new outbreaks.

To measure the capacity of algorithms to predict *new*, emerging diseases, we will score using structured cross-validation in which no disease outbreak prediction is based on an algorithm

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

trained on the same pathogen. For example, for Ebola outbreak predictions, all previous Ebola outbreaks will be held out from training data.

#### Model Evaluation and Validation

Models will be fit and tested against their ability predict (a) size of an outbreak (total number of cases), (b) peak size of outbreak (number of cases at highest point), (c) time to completion of an outbreak, (d) time to peak of an outbreak, and (e) geographic spread of outbreak (distance to farthest case from point of origin). Prediction scores will consist of mean-square-error of each of these prediction against true values, as well as a score of equal-weighted errors of all five measures. All models will also be tested for fidelity of uncertainty measures—the fraction of true data values that fall within prediction confidence intervals—and these metrics will be reported along with predictive performance.

All model performance scores will be compared against meaningful, "practical naïve" baselines. While it is common practice to use a "null" or "zero-knowledge" baseline, this often amounts to inflating the apparent power of the algorithm. We will opt for a more realistic, "practical naïve" baseline, which instead follows what an analyst would approximate in practice. For these models, we will use a practical baseline model of expecting outbreaks to follow the average size and timing of outbreaks of the same disease in the past 25 years. All model scores will be reported as percentage improvements over this baseline.

Algorithm performance will be measured against holdout data to validate true predictive performance. Repeat testing on holdout data can lead to overfitting, with predictions too specific to the hold-out set and not indicative of future performance. Yet increasing holdout reduces the availability of training data. Thus, we will use a *differential privacy* approach drawn from information security to validate prediction results. The hold-out data will be obfuscated, and validation performance will be measured against bootstrapped replicates of validation data, essentially introducing noise to reduce overfitting against the hold-out. This will ensure accurate estimates of predictive performance in an operational environment. 10

#### Data Sources

Data sources for fitting and testing models include output data of number, timing, and geography of disease outbreaks, and predictor data include pathogen traits (disease symptoms, pathogen traits, modes of transmission), pathogen sources (point of origin, geographic range of potential natural hosts), environmental context (climatic conditions, ecosystem traits), and socioeconomic context (local population densities, demographics, transportation networks, healthcare systems). Gathering predictor data is often as large a factor in the time required to create first-pass predictions at the early stage of an outbreak. Thus, we will create automated pipelines for gathering, merging, and updating these data sets, and to extract key predictors from basic location and host variables to minimize manual inputs required. For instance, by linking to high-resolution geospatial databases, we will automatically populate data for population density,

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

ecosystem types, transport infrastructure that can be updated and refined with on-the-ground data if available.

Disease outbreak impact data: We will draw a variety of sources to build a database of several thousand of outbreak time-series to train our models. First, we will use time series from ΛΙDO, the most comprehensive such repository, with approximately 700 series across 40 diseases. Recognizing that ΛΙDO has limits and data gaps, we will supplement this with several other sources including One Health disease datasets compiled and maintained by ΕΗΛ:

- COVID-19 Data: Data on cases, hospitalization, and deaths from COVID-19 are uniquely
  available at national, sub-national and city scales for a broad swath of countries with varying
  conditions and policy responses. While the COVID-19 pandemic is linked across countries,
  in our hierarchical modeling framework we will to separate these out into multiple regional
  outbreaks, shedding more light on the effects of environmental and socioeconomic factors
  under a common host and pathogen.
- Multispecies datasets: We will supplement human disease outbreaks with data on veterinary disease outbreaks. EHA maintains a database on veterinary disease outbreaks derived from OIE and FAO data and supplemented by digitized archival reports (extending back to 1992), with 2900 outbreak records across 122 diseases. This will more than quadruple the available disease time series for training. Partial-pooling will ensure that only information in relevant veterinary data sets will be applied to human outbreaks and vice versa. For instance, effects of population density or health/veterinary capacity are expected to have partial correlation across all diseases. In addition, this will allows the model to predict impacts from these diseases, which has direct defense applications given the food supply, trade, and political stability impacts of veterinary diseases
- Emerging Infections Disease Repository: EHA's Emerging Infectious Disease Repository (eidr.ecohealthalliance.org/)<sup>14</sup> contains data on an additional 350 zoonotic disease emergence outbreaks, giving additional insights in particular to new emergence events from zoonotic sources. Full time-series are incomplete but outbreak size and duration are recorded, allowing them to be integrated into our framework with multiple imputation.

*Predictor variables*: Disease dynamics depend on a complex interplay of pathogen, host, and environmental conditions. We will use data on pathogens (traits, genomes), sources (reservoir host traits, range, relatedness to humans), environmental context (climate, ecosystems traits), and socioeconomic context (local population densities, transportation networks, healthcare systems) as predictors for our models, as well as time-series data of infections from the earliest phase of the outbreaks. As we expect all data will be available for neither our full training set nor in common condition of new outbreaks, use multiple imputation<sup>12</sup> in all models to allow for prediction as well as uncertainty assessment under incomplete data.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

- Pathogen Trait Data consists of information about pathogen taxa, transmission mode, structural, and biological traits. Relevant traits for pathogens, and have been largely aggregated for viruses by our Host-Pathogen Phylogeny Project<sup>5</sup> and parallel projects for bacterial pathogens<sup>15</sup> and non-zoonotic human viruses<sup>16</sup>.
- Pathogen Genetic Data consist of genetic sequence of representative samples of individual pathogen species. Representative sequences of nearly all relevant pathogens are available via genetic repositories GenBank, ICTV, or GISAID. We expect pathogen data at the scale of pathogen species to largely predict similar information as pathogen traits, but genomic data may be faster to use in cases of rapid sequencing of novel pathogens. We will test whether incorporating genetic data into model features via neural-network autoencoders<sup>17</sup> provides improved performance over separate representation of pathogen traits (e.g., taxonomic classification, transmission mode), or other representations of genetic data such as phylogeny. Whether this performance is additive, or whether these two types of information are redundant. Even if redundant, genetic data may improve predictive capacity in the case of new pathogens where pathogen traits have not been fully delineated. Rich inputs from genetic data will be integrated using neural-network encoders.
- Source Data refers to intelligence about the origin of the disease, which includes location and mode of origin (environmental, zoonotic host species, laboratory or accidental release). In many cases source data is unknown, particularly in the early stages of a new outbreak event. For zoonotic-origin diseases, the pool of known host species and their geographic provide probabilistic information on the source. Our own Host-Pathogen Phylogeny Project aggregates host traits for all known mammalian diseases hosts.<sup>5</sup>
- Environmental Data refers to information describing the conditions in which the disease may spread. These will include local climate variables from the BIOCLIM database, land-use coverage (HYDE), and human and livestock population densities (World Bank, FAO).
- Socioeconomic Data these include economic activity, governance, medical and veterinary capacity (World Bank, FAO), and transportation network connectivity in the region of disease spread.

#### **Programmatics**

All work will be undertaken by the project prime, EcoHealth Alliance (EHA). EHA has a proven track record of project management with U.S. government agencies including USAID, DTRA, DHS, NIH, and NSF, ranging from global epidemiological research and surveillance programs to small-to-medium scale technology development.

Our management approach is to bring production-quality software development practices to data science for research and development. We use agile development methodology, with weekly development meetings, monthly project status reviews, and regular organization-wide review seminars to gather feedback from our broader interdisciplinary expertise. All tasks in the

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

statement of work will benefit from the significant experience of the project staff and in-kind contribution from other EHA personnel with combined expertise in global emerging infectious disease epidemiology, public health, veterinary medicine, machine learning, and biostatistics. We also focus on *deployment-ready* research infrastructure to minimize the need to re-engineer models developed in the research process for applied use.

We engage multiple best practices to minimize project disruption risk. *Personnel risk* entails loss of expertise and project or institutional knowledge due to loss of core project staff. EHA maintains project data, documentation, and code in common cross-team repositories. Version-controlled project repositories ensure that project histories are tracked, and all team members can retrieve previous work by others. Through regular code review, we ensure all team members have working knowledge of project modules and the ability to re-task. *Infrastructure risk entails* loss of project infrastructure such a. EHA performs on-site high-performance computing cluster or data storage. We keep code, data, results and documentation on-site only for active experiments, and back up these to off-site cloud infrastructure. Our compute environment is reproducible, being configured by pre-programmed, self-documenting methods such as Dockerfiles, Ansible playbooks, and Bash scripts. It can be re-deployed to off-site cloud resources or new hardware as needed.

#### Relevance

Our research will fundamentally improve infectious disease forecasting by identifying the relative importance of early-stage factors in outbreak prediction. While pathogen-host-environment linkages are fundamental in disease modeling, predictive methods developed to assess emerging biothreats usually focus on each individually. We will also advance knowledge of hybrid ML/mechanistic models, which have not yet been used in applied disease forecasting. Together this research will impact C-WMD science by providing practical guidance for early-stage data-gathering and prototyping an interactive decision support tool for biothreat impact forecasting.

#### **Credentials**

EcoHealth Alliance (EHA) will implement this project, with Noam Ross, Principal Scientist for Computational Research, as project lead. EHA's mission is focused on emerging diseases and cutting-edge research into the critical connections between human, wildlife, and ecosystem health that drive disease emergence and spread.

**PI:** Dr. Ross will direct he project. He is a computational disease ecologist who provides expertise in dynamic epidemiological processes, machine learning and biostatistics, and high-performance computing. He has led work in ecological and epidemiological modeling and data management for EHA's work on USAID PREDICT and NIH NIAID viral surveillance research. He has experience developing livestock disease outbreak and spread simulations in collaboration with FAO as well as private industry. Dr. Ross is PI for EcoHealth's work with DHS on machine learning for veterinary disease outbreaks. He also a leader in open-source scientific computing.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

He is a founding editorial board member of rOpenSci, an organization that defines standards for reproducibility and runs peer review for scientific software, and is a core contributor to the Rocker project, which provides open-source, standardized computing containers for data science. Dr. Ross holds a Ph.D. in computational ecology from the University of California-Davis.

**Organization:** EHA has a proven track record of large project management with U.S. government agencies. We lead the largest global country portfolio, and serve as lead for Modeling and Analytics and Behavioral Risk Analysis teams, for the USAID Emerging Pandemic Threats PREDICT program, a \$118 million program focused on predicting and preventing pandemic diseases. The PREDICT project is building a global early warning system to detect and reduce the impacts of emerging diseases that move between wildlife and people (zoonotic diseases). EHA also leads multiple field epidemiological contracts funded by DTRA, NIH, and NSF.

EHA has proven expertise in surveillance-oriented technology development. We managed a five-year contract to develop applications for DTRA's Chemical Biological Integration Platform (CBIP, formerly the Biosurveillance Ecosystem—BSVE), as well as the development of the Inbound Bioevent Information System (IBIS), and Rapid Evaluation to Prevent Epidemics in Livestock (REPEL) project under DHS's Analytics for Prevention of Biothreat Incidents (APBI) program and Science and Technology Directorate. Through this work, EHA scientists have gained substantial expertise in both basic research and working with government project managers and analysts to develop technologies that are in line with the goals of the funding. The expertise resulting from this effort serves as the foundation for the current proposal.

**Team:** Robert Young, Senior Research Software Engineer, will lead project infrastructure development and the creation ML pipelines. Mr. Young has a background in artificial intelligence and bioinformatics. He has developed and managed laboratory information systems in neuroscience, genomics and performed research in neural-network design for image-recognition in astronomy. Mr. Young currently leads cyperinfrastructure work for EHA's veterinary disease forecasting under DHS as well as the DTRA-Funded Western Asia Bat Research Network.

Emma Mendelsohn, M.Sc., Research Scientist is a data scientist with extensive expertise in dynamic systems modeling, biostatistics, web application development and data engineering. Emma is lead analyst for EHA's work on antimicrobial resistance emergence work as well as projects linking emerging infectious diseases with nutrition and non-communicable diseases. Emma will lead data engineering and contribute expertise to algorithm development and application design.

Peter Daszak, Ph.D., President of EcoHealth Alliance will provide oversight and act as project advisor on infectious disease spread. Dr. Daszak is a leader in the field of emerging diseases and disease ecology.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

William Karesh, DVM, Senior Vice President of EcoHealth Alliance will serve as project advisor on veterinary topics as well as liaise with data-providing organizations, such as OIE and ProMED, as required. Dr. Karesh is President of the Working Group on Animal Health and COVID-19 Task Forces for OIE, Co-Chair of the Wildlife Health Specialist Group of the IUCN.

**Facilities:** EcoHealth Alliance is a 48-year old scientific research NGO that specializes in multidisciplinary research and surveillance of the spread of zoonotic emerging diseases. EcoHealth Alliance is based in New York City with 10,000 ft<sup>2</sup> of office space. The scientific and technology staff of 30 are supported by a core administrative staff of 8 people who are available for work on this project and are funded through core funds.

EcoHealth Alliance is equipped with fiber optic Internet access and video conferencing facilities to facilitate easy communication between collaborators. EcoHealth Alliance employees have around the clock access to servers, VPNs, encryption software, and IT support. We use standard office suite services including Google Apps (Hosted email and web-based collaboration software), Microsoft Office, and Slack, running on Apple macOS, Ubuntu Linux, and Windows Operating Systems. EHA teams work in multiple computer languages (Python, NodeJS, and R programming languages), development frameworks (Agile, continuous integration), and machine-learning toolboxes. Our development operations are based on reproducible, containerized frameworks (Ansible, Docker). We perform high-performance computation across our dedicated cluster of System76 multi-core serversand with NVIDIA deep-learning GPUs and spot cloud somputs instances through Amazon Web Services EC2 and Microsoft Virtual Machines, We use a variety of other cloud services for storage (AWS S3, Azure Blob), continuous integration (GitHub, Gitlab), and other tasks.

## Work to be Performed

Tasks in Year. Task. Subtask format

#### **YEAR 1:**

**Task 1.1 – Develop and deploy project infrastructure:** Infrastructure engineering consists of provisioning a best-in-class computational environment that enables continuous model development and testing and allows for research and models to be reproduced and extended by external teams. It includes continuous maintenance and documentation of project codebase.

- 1.1.1 Layout a provision project network
- 1.1.2 Set up continuous integration pipelines
- 1.1.3 Implement backup and data verification jobs
- 1.1.4 Implement differential privacy testing
- 1.1.5 Write and maintain codebase documentation
- 1.1.6 Perform code review, testing and validation

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

Task 1.2 – Engineer Project Database: Data engineering consists of curating, integrating and cross-linking and validating relevant data sets from various sources into a project database. This allows data with different sources, space and time scales, formats, and completeness to be used in common models establishes a database that can be used as a common resource.

#### Subtasks

- 1.2.1 Design database schema
- 1.2.1 Write pipelines to ingest time-series data
- 1.2.2 Write pipelines to ingest predictor data
- 1.2.3 Cross-link data sources via common ontologies
- 1.2.4 Write quality assurance/quality control (QA/QC) tests for data and perform data cleaning
- 1.2.5 Create automatic extraction routines to populate relevant predictors from location information
- 1.2.6 Update data from live sources and clean and maintain data continuously

**Task 1.3 – ML Model Development and Testing:** The core ML development task consists of designing, implementing, testing, and analyzing performance of ML model structures and iteratively improving them.

#### Subtasks

- 1.3.1 Implement "practical naïve" model baseline
- 1.3.2 Design Bayesian additive regression tree (BART) model structure
- 1.3.3 Design alternative generalized additive model (GAM) structure
- 1.3.4 Implement within-disease, all-disease, and partial-pooled disease model versions
- 1.3.5 Measure variable contribution by re-fitting
- 1.3.6 Analyze prediction-level variable contributions and model performance
- 1.3.7 Iteratively update models to improve performance

#### **OPTION YEAR:**

Task 2.1 – Develop and deploy project infrastructure: In the option year we will continue to perform maintenance and documentation of project codebase to ensure reproducibility and reusability of additional components

#### Subtasks

- 2.1.1 Write and maintain codebase documentation
- 2.1.2 Perform code review, testing and validation
- **Task 2.2 Engineer Project Database:** In the option year we will extend the database to include additional predictors and continuously maintain and update to accumulate more data from upstream sources.

- 2.2.1 Extend database schema to include genetic data and auto-encoded variables
- 2.2.2 Write pipelines to ingest genetic data
- 2.2.3 Cross-link data sources via common ontologies
- 2.2.4 Write QA/QC tests for data and perform data cleaning

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

2.2.5 Update data from live sources and clean and maintain data continuously

Task 2.3 – BART Model Development and Testing: In the option year we will extend the BART model to ingest genetic data and test the contribution of this data in enhancing or recapitulating pathogen trait data.

#### Subtasks

- 2.3.1 Develop genetic predictor variable set via neural-network autoencoding
- 2.3.2 Modify BART and GAM structures to incorporate genetic variables
- 2.3.3 Implement within-disease, all-disease, and partial-pooled disease model versions
- 2.3.4 Compare performance of genetic, pathogen-trait, and combined models

Task 2.4 – Hybrid ML-Mechanistic Model Development and Testing: In the option year will develop the hybrid machine-learning mechanistic model, compare its performance to a pure ML model and determine the switch-over time when the mechanistic model becomes more predictive in the course of the outbreak.

- 2.4.1 Design mechanistic model structure
- 2.4.2 Determine mechanistic model prior forms
- 2.4.3 Modify ML model to predict mechanistic parameters rather than outcomes
- 2.4.4 Implement within-disease, all-disease, and partial-pooled disease model versions
- 2.4.5 Measure variable contribution by re-fitting
- 2.4.6 Analyze prediction-level variable contributions and model performance
- 2.4.7 Compare Hybrid and ML performance along outbreak time-series progression
- 2.4.8 Iteratively update models to improve performance

PI: Noam Ross

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

## Performance Schedule

Description	Y1 Q1-2	Y1 Q3-4	OY 01-2	OY 03-4
·	111041-2	111043-4	01 212	01 23-4
Task 1: Develop and deploy project infrastructure			1	
1.1.1 Layout a provision project network				
1.1.2 Set up continious integration pipelines				
1.1.3 Implement backup and data verification jobs				
1.1.4 Implement differential privacy testing				
1.1.5 Write and maintain codebase documentation				
1.1.6 Code review, testing and validation		1		
2.1.5 Write and maintain codebase documentation				
2.1.6 Perform code review, testing and validation				
Task 2: Engineer Project Database				
1.2.1 Design database schema				
1.2.1 Write pipelines to ingest time-series data				
1.2.2 Write pipelines to ingest predictor data				
1.2.4 Cross-link data sources via common ontologies				
1.2.4 Write QA/QC tests for data and perform data cleaning				
1.2.5 Create automatic extraction routines to populate relevant predictors				
1.2.6 Update data from live sources and clean and maintain data continuously				
2.2.1 Extend database schema to include genetic data				
2.2.2 Write pipelines to ingest genetic data				
2.2.3 Cross-link data sources via common ontologies				
2.2.4 Write QA/QC tests for data and perform data cleaning				
2.2.5 Update data from live sources and clean and maintain data continuously				
Task 3: ML Model Development and Testing				
1.3.1 Implement "practical naïve" model baseline				
1.3.2 Design BART model structure				
1.3.3 Design alternative GAM model structure				
1.3.4 Implement within-disease, all-disease, and partial-pooled disease models versions				
1.3.5 Measure variable contribution by re-fitting				
1.3.6 Analyze prediction-level variable contributions and model performance				
1.3.7 Iteratively update models to improve performance				
2.3.1 Develop genetic predictor variable set via neural-network autoencoding				
2.3.2 Modify BART and GAM structures to incorporate genetic variables				
2.3.3 Implement within-disease, all-disease, and partial-pooled disease models				
2.3.4 Compare performance of genetic, pathogen-trait, and combined models				
Task 4: Hybrid Model Development and Testing				
2.4.1 Design mechanistic model structure				
2.4.2 Determine mechanistic model prior forms				
2.4.3 Modify ML model to predict mechanistic parameters				
2.4.4 Implement within-disease, all-disease, and partial-pooled models				
2.4.5 Measure variable contribution by re-fitting				
2.4.6 Analyze prediction-level variable contributions and model performance				
2.4.7 Compare Hybrid and ML performance along outbreak time-series progression				
2.4.8 Iteratively update models to improve performance				

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

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Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

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Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

## Attachment 1 - Statement of Work (SOW)

Project Title: Predicting Biothreat Impacts from Early-Stage Data via Transfer Learning

Document Date: May 28, 2020

## **Objective**

The project's objectives are to determine the efficacy of machine learning (ML) techniques in predicting epidemic impacts in early-stage, data-sparse conditions, particularly novel biothreats, both intentional (e.g. novel genetically-modified pathogens) and natural (e.g. zoonotic). Specifically, we will answer three questions that will enable rapid decision-making based on ML models of epidemic impacts:

**Q1:** How much does partial-pooling improve predictions of epidemic impacts, and what information transfers best across diseases?

**Q2:** What variables are most predictive of epidemic impacts at early stages of an outbreak?

**Q3:** At what point in epidemic growth do hybrid models that incorporate mechanistic epidemiological components become more accurate than pure ML models?

This work will contribute to fundamental knowledge of properties and variables that contribute most to predictive accuracy at early-stage stages in emerging outbreak prediction and forecasting. It will enable the development of robust and deployable predictive models at early stages of outbreaks, and support informed choices for data collection and model selection.

#### Scope

The awardee proposes a two-year study to determine the efficacy of ML techniques in predicting epidemic impacts in early-stage, data-sparse conditions, particularly novel biothreats. The awardee team shall focus on the following major goals and milestones:

- Year 1, Milestone 1 (6 months): Create and curate predictor database, project infrastructure, continuous integration/validation, establish data-imputation and automatic predictor workflow (Tasks 1.1-1.2).
- Year 1, Milestone 2 (12 months): Bayesian additive regression tree (BART) model fitting and tuning, determination of variable contribution to prediction accuracy (Task 1.3)
- Option Year, Milestone 3 (18 months): Maintain and document project infrastructure, extend the database to include additional predictors, extend the BART model to ingest genetic data (Tasks 2.1-2.3)
- Option Year, Milestone 4 (24 months): Construction of hybrid ML/mechanistic model structure, comparative testing of pure and hybrid version, determination of timing when current data overtakes transfer data (Task 2.4)

## Background

Situational awareness and risk assessment are critical at early stages of disease outbreaks, when interventions can best mitigate adverse outcomes. Yet decision-makers often face a paucity of data at these stages. Traditional modeling methods to forecast disease impact severity rely heavily on prior knowledge or assumptions about mechanisms of disease spread progression, and they frequently do not anticipate the actual mechanisms.<sup>1-3</sup> They also take considerable time and effort to develop.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

Recent theory suggests disease dynamics are fundamentally predictable<sup>4</sup>, and ML methods have been successfully applied in emerging disease science. ML refers to a group of methods, ranging from simple nearest-neighbor approaches to function-approximation methods such as boosted trees and penalized splines, to complex deep-learning neural networks. ML has been used to predict the likelihood of disease emergence and the zoonotic potential of various hosts and viruses<sup>5,6</sup>, and deep-learning approaches have been used to infer viral hosts from the comparatively richer data of genetic sequences of viruses.<sup>7,8</sup>

ML methods are generally unstructured, flexible, and offer greater predictive power over traditional statistical approaches. However, these methods are data-intensive, and a key challenge with ML development is data paucity and/or a low signal-to-noise ratio. Surveillance data for even existing, well-studied diseases is rarely rich enough for the data scale of many modern ML techniques. For instance, a ML approach to predict impacts from a new outbreak of Ebola would draw training data from fewer than twenty previous events, each with varying levels of data coverage.

Transfer learning, or data fusion, provides a solution to data paucity issues by using data from other domains to train parts of models that are general, thus limiting new data requirements for more specific components. Hierarchical modeling, or *partial pooling*, is a transfer learning approach for partitioning out which data can be shared between models and which are only applicable to a specific case. For instance, Ebola forecasts for countries outside the historic range of the disease may be improved by calibrating against Cholera outbreaks, which have occurred both within and outside countries with Ebola outbreaks. The degree to which the disease outcomes are correlated is learned, rather than assumed.

An alternative recent development in disease forecasting is the development of hybrid models that incorporate simple mechanistic structures and predictive ML methods to learn model parameters. These approaches have demonstrated high predictive accuracy for near-term forecasting of epidemic growth and impact, particularly during the COVID-19 pandemic<sup>9</sup>. However, these approaches require adequate time-series of data to project forward from current conditions. Their utility in early stages and long-term forecasting accuracy has yet to be determined. In opting to apply these models, it is critical to understand these data requirements and at what point in an outbreak hybrid approaches can overtake pure ML early-stage predictions in accuracy. This project seeks to develop, apply, and compare pure and hybrid-ML techniques to enable the development of robust, rapidly usable predictions at early stages of outbreaks.

Key references (additional references can be found in the Technical Proposal):

- MacDiarmid, S. C. et al. Handbook On Import Risk Analysis for animals and animal products: quantitative risk assessment. Vol. 2 (Office International des Épizooties, 2004).
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- Delgado, J. *et al.* U.K. Foot and Mouth Disease: A Systemic Risk Assessment of Existing Controls. *Risk Anal.* **37**, 1768-1782, doi:10.1111/risa.12704 (2017).

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

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- Olival, K. J. *et al.* Host and viral traits predict zoonotic spillover from mammals. *Nature*, doi:10.1038/nature22975 (2017).
- Allen, T. *et al.* Global hotspots and correlates of emerging zoonotic diseases. *Nat. Commun.* **8**, 1124, doi:10.1038/s41467-017-00923-8 (2017).
- 7 Mock, F., Viehweger, Λ., Barth, E. & Marz, M. VIDHOP, viral host prediction with Deep Learning. *Bioinformatics* (2019).
- Babayan, S. A., Orton, R. J. & Streicker, D. G. Predicting reservoir hosts and arthropod vectors from evolutionary signatures in RNA virus genomes. *Science* **362**, 577-580, doi:10.1126/science.aap9072 (2018).
- 9 Gu, Y. COVID-19 Projections Using Machine Learning, <a href="https://covid19-projections.com/about/#historical-performance">https://covid19-projections.com/about/#historical-performance</a> (2020).

#### Data Sources

Fitting and testing models will draw from disease outbreak data (counts, timing, and geography) and predictor data including pathogen traits (disease symptoms, pathogen traits, modes of transmission), pathogen sources (point of origin, geographic range of potential natural hosts), environmental context (climatic conditions, ecosystem traits), and socioeconomic context (local population densities, demographics, transportation networks, healthcare systems). Please see the Technical Proposal for available data sources.

## Tasks/Scientific Goals (Format: Year. Task. Subtask)

#### YEAR 1

## Task 1.1 – Develop and deploy project infrastructure

The awardee shall develop a computational environment that enables continuous model development and testing and allows for research and models to be reproduced and extended by external teams. The computational environment will include continuous maintenance and documentation of project codebase.

#### Subtasks

- 1.1.1 Layout a provision project network
- 1.1.2 Set up continuous integration pipelines
- 1.1.3 Implement backup and data verification jobs
- 1.1.4 Implement differential privacy testing
- 1.1.5 Write and maintain codebase documentation
- 1.1.6 Perform code review, testing and validation

## Task 1.2 – Engineer project database

The awardee shall curate, integrate, cross-link, and validate relevant datasets from various sources into a project database, accounting for differing space and time scales, data formats, and data completeness.

- 1.2.1 Design database schema
- 1.2.1 Write pipelines to ingest time-series data
- 1.2.2 Write pipelines to ingest predictor data
- 1.2.3 Cross-link data sources via common ontologies

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

- 1.2.4 Write quality assurance/quality control (QA/QC) tests for data and perform data cleaning
- 1.2.5 Create automatic extraction routines to populate relevant predictors from location information
- 1.2.6 Update data from live sources and clean and maintain data continuously

## Task 1.3 - Machine Learning (ML) model development and testing

The awardee shall design, implement, test, and analyze performance of ML model structures and iteratively improve them.

Subtasks

- 1.3.1 Implement "practical naïve" model baseline
- 1.3.2 Design Bayesian additive regression tree (BART) model structure
- 1.3.3 Design alternative generalized additive model (GAM) structure
- 1.3.4 Implement within-disease, all-disease, and partial-pooled disease model versions
- 1.3.5 Measure variable contribution by re-fitting
- 1.3.6 Analyze prediction-level variable contributions and model performance
- 1.3.7 Iteratively update models to improve performance

#### **OPTION YEAR:**

## Task 2.1 – Maintain and document project infrastructure

In the option year, the awardee shall continue to perform maintenance and documentation of project codebase to ensure reproducibility and reusability of additional components

Subtasks

- 2.1.1 Write and maintain codebase documentation
- 2.1.2 Perform code review, testing and validation

#### Task 2.2 – Extend project database

In the option year, the awardee shall extend the database to include additional predictors and continuously maintain and update to accumulate more data from upstream sources.

Subtasks

- 2.2.1 Extend database schema to include genetic data and auto-encoded variables
- 2.2.2 Write pipelines to ingest genetic data
- 2.2.3 Cross-link data sources via common ontologies
- 2.2.4 Write QA/QC tests for data and perform data cleaning
- 2.2.5 Update data from live sources and clean and maintain data continuously

## Task 2.3 – Extend BART model

In the option year, the awardee shall extend the BART model to ingest genetic data and test the contribution of this data in enhancing or recapitulating pathogen trait data.

Subtasks

- 2.3.1 Develop genetic predictor variable set via neural-network autoencoding
- 2.3.2 Modify BART and GAM structures to incorporate genetic variables
- 2.3.3 Implement within-disease, all-disease, and partial-pooled disease model versions
- 2.3.4 Compare performance of genetic, pathogen-trait, and combined models

## Task 2.4 - Hybrid ML-mechanistic model development and testing

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

In the option year, the awardee shall develop the hybrid machine-learning mechanistic model, compare its performance to a pure ML model, and determine the switch-over time when the mechanistic model becomes more predictive in the course of the outbreak.

#### Subtasks

- 2.4.1 Design mechanistic model structure
- 2.4.2 Determine mechanistic model prior forms
- 2.4.3 Modify ML model to predict mechanistic parameters rather than outcomes
- 2.4.4 Implement within-disease, all-disease, and partial-pooled disease model versions
- 2.4.5 Measure variable contribution by re-fitting
- 2.4.6 Analyze prediction-level variable contributions and model performance
- 2.4.7 Compare Hybrid and ML performance along outbreak time-series progression
- 2.4.8 Iteratively update models to improve performance

#### **Deliverables**

Reports and Documents

- Monthly reports on task status
- Technical papers describing algorithm design and performance
- Conference presentations

#### Code Modules and Database Archives

All code modules and documentation will be continually available on EcoHealth Alliance's GitHub page with access provided to DTRA. Databases will be made available for download and provided on-disk to DTRA per request.

**Cover Sheet** 

Proposal Number FRBAA14-6-2-0502

Phase I Proposal Number

Topic Thrust Area 6

Proposal Title Predicting Biothreat Impacts from Early-Stage Data via Transfer Learning

**Applicant Information** 

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**DUNS** 0770900660000 **State/Province** MY

**CAGE** Zip 10001 - 2317

Website Country USA

POC Name Dr. Noam Ross POC Email ross@ecohealthalliance.org

Cost

**Applicant Certification** 

Organization Type Non-Profit Organization

Are Human Subjects involved? No

Are Vertebrate Animals involved? No

Has a proposal for essentially equivalent work been submitted to other US government agencies or DoD components?

Agency 1 Contract/Grant No.

No

Agency 2 Contract/Grant No.

Agency 3 Contract/Grant No.

Are you a current DoD Contractor or No

Grantee?

Agency Point Of Contact

Phone #

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Proprietary Information (list page numbers)

List a maximum of 8 Key Words or phrases, separated by commas, that describe the Project.

Technical Abstract (Limit your abstract to 200 words with no classified or proprietary information)

Knowingly and willfully making any false-fictitious or fraudulent statements or representations may be a felony under the Federal Criminal False Statement Act (18 USC Sec 1001) punishable by a fine of up to \$10 000, up to five years in prison, or both

PI: Noam Ross

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

## Predicting Biothreat Impacts from Early-Stage Data via Transfer Learning

### Phase II Technical Proposal

#### ABSTRACT

Accurately forecasting the impacts of novel biological agents in the early stages of their emergence is critical to planning control measures for warfighters and civilians. However, at the beginning of outbreaks, data about pathogen properties and dynamics are often scarce. Machine learning (ML) techniques generally require large data sets, thus hampering their utility in these contexts. Our proposed work addresses this challenge by using cross-disease prediction via transfer learning utilizing data from previous emergence events to improve early-stage predictions for future events. We will use partial-pooling to determine the degree to which information can be shared across diseases according to hierarchical similarities in pathogen and host properties, and in environmental and population conditions. We will also determine which properties contribute most to predictive accuracy, providing guidance for prioritizing datacollection in early outbreak stages. We will test and compare two approaches – pure ML predictions and hybrid ML/mechanistic models – to determine which provides better predictions of outbreak size and impacts. We will determine at what point in an outbreak current data overtakes historical data as a better predictor of outbreak dynamics. Making use of data sets from multiple host species as well as efficient model-validation techniques, we will expand the size of available training data and thus models' predictive capacity.

#### **SCOPE**

#### Objective

Our primary research objectives are to determine the efficacy of ML techniques in predicting epidemic impacts in early-stage, data-sparse conditions, particularly novel biothreats both intentional (e.g. novel genetically-modified pathogens) and natural (e.g. zoonotic). Specifically, we will answer three questions that will enable rapid decision-making based on ML models of epidemic impacts:

Q1: How much does partial-pooling improve predictions of epidemic impacts, and what information transfers best across diseases?

Q2: What variables are most predictive of epidemic impacts at early stages of an outbreak?

**Q3:** At what point in epidemic growth do hybrid models that incorporate mechanistic epidemiological components become more accurate than pure ML models?

This work will contribute to fundamental knowledge of properties and variables that contribute most to predictive accuracy at early-stage stages in emerging outbreak prediction and forecasting. It will enable the development of robust and deployable predictive models at early stages of outbreaks, and support informed choices for data collection and model selection.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

## **Background**

Situational awareness and risk assessment are critical at early stages of disease outbreaks, when interventions can best mitigate adverse outcomes. Yet decision-makers often face a paucity of data at these stages. Traditional modeling methods to forecast disease impact severity rely heavily on prior knowledge or assumptions on mechanisms of disease spread progression, and they frequently do not anticipate the actual mechanisms.<sup>1-3</sup> They also take considerable time and effort to develop.

Recent theory suggests disease dynamics are fundamentally predictable<sup>4</sup>, and machine learning (ML) methods have demonstrated utility for some aspects of emerging disease science, e.g. to predict the likelihood of disease emergence and zoonotic potential of various hosts and viruses<sup>5,6</sup>, and deep learning has been used to infer viral hosts from the comparatively richer data of genetic sequences of viruses.<sup>7,8</sup>

Machine learning (ML) refers to a group of methods, ranging from simple nearest-neighbor approaches to function-approximation methods such as boosted trees and penalized splines, to complex deep-learning neural networks. These methods are generally unstructured, flexible, and offer greater predictive power over traditional statistical approaches. However, these methods can also be data-intensive; a key challenge with ML methods development is that data are too sparse, or signal/noise ratio too low, to effectively train algorithms. Surveillance data for even existing, well-studied diseases is rarely rich enough for the data scale of many modern machine learning or deep learning techniques. For instance, an ML approach to predict impacts from a new outbreak of Ebola would have training data from to less than twenty previous events for which there is comprehensive epidemiological data from 10 West and Central African nations

Transfer learning, or data fusion, provides a solution for using data from other domains to train parts of models that are general, thus limiting new data requirements for more specific components. Hierarchical modeling, or *partial pooling*, is a transfer learning approach for partitioning out which data can be shared between models and which are only applicable to a specific case. For instance, Ebola forecasts from countries outside of the virus' historic range may be improved by calibrating against influenza outbreaks, which have occurred both within and outside countries with Ebola outbreaks. The degree to which the disease outcomes are correlated (e.g., similarity of modes of transmission) are learned, rather than assumed.

An alternative recent development in disease forecast is the development of hybrid models that incorporate simple mechanistic structures and predictive ML methods to learn model parameters. These approaches have demonstrated high predictive accuracy for near-term forecasting of epidemic growth and impact, particularly during the COVID-19 pandemic<sup>9</sup>. However, these approaches require adequate time-series of data to project forward from current conditions. Their utility in early stages and long-term forecasting accuracy has yet to be determined. In opting to apply these models, it is critical to understand these data requirements and at what point in an outbreak hybrid approaches can overtake pure ML early-stage predictions in accuracy.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

We will develop algorithms to predict the dynamics and impacts of outbreaks of new infectious diseases, based on data available at early stages of outbreaks. In doing so, we will answer the following key questions.

- How much does partial-pooling improve predictions of epidemic impacts, and what information transfers best across diseases?
- What variables are most predictive of epidemic impacts at early stages of an outbreak?
- At what point in epidemic growth do hybrid models that incorporate mechanistic epidemiological components become more accurate than pure ML models?

### Algorithmic Methods:

We will test the ability to ML methods incorporating partial pooling to predict impacts from a new outbreak of disease. Our primary model structure will be built around Bayesian additive regression trees (BARTs)<sup>10</sup>. BARTs provide three features appropriate for this application: First, they can provide both high-quality predictions as well as robustness in relatively data-poor scenarios. Secondly, they have explicit uncertainty structures is better able to characterize error than other regression tree methods that do not. Thirdly, they are able to incorporate hierarchical partial pooling. In some scenarios where data are smooth rather than categorical, generalized additive models (GAMs) can offer similar performance and similar features<sup>11</sup>, and we will test these as an alternate formulation.

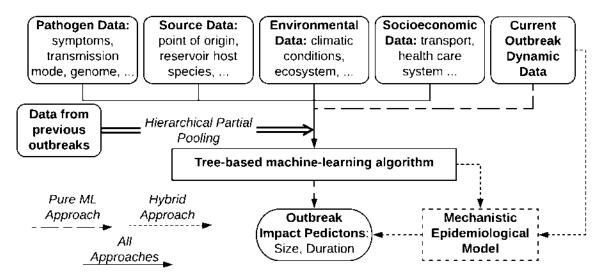
We will parametrize the models using data on pathogen traits (disease symptoms, pathogen traits, modes of transmission), pathogen sources (point of origin, geographic range of potential natural hosts), environmental context (climatic conditions, ecosystem traits), and socioeconomic context (local population densities, demographics, transportation networks, healthcare systems), as well as time-series data of infections from the earliest phase of the outbreaks. Not all these data are typically available at the early stages of a new bioagent outbreak, and are in many cases incomplete even in the case of historical outbreaks. Thus, we will use multiple imputation<sup>12</sup> in all models to allow for model fitting, prediction and assessment of uncertainty under incomplete data.

We will explore multiple model structures, but all formulations will be fit and tested under three conditions: within-disease forecasting where models are fit only on data from a single disease and tested at ability to predict impacts from that disease, all-disease forecasting where models are fit on data from all diseases and tested against ability to predict each disease, and partial-pooled where models are fit all models from all diseases but parameter sharing across diseases are partially pooled. (See 'Model Evaluation and Validation' below)

While ML approaches are best for predictive power, they do not have the flexibility to be used for alternate scenarios or intervention evaluation that traditional compartmental models do. We will compare "pure" ML algorithms to a hybrid approach using ML to generate parameters for mechanistic epidemiological models. In this hybrid case, we will use the ML model to predict prior parameters for compartmental epidemiological model using the same data as used in the

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

pure ML model, excluding early time-series data. The model be structured to have high flexibility under small number of parameters – i.e. SIR-type model with variable contact structure and latent/infectious periods. This model will then be fit to the time series data for the outbreak under the constraint of these prior parameters. Using this approach (see figure), we will determine whether the structural knowledge embedded in the epidemiological model improves outbreak predictions or hinders them due to assumptions. As with the pure ML approach, we will test the hybrid model approach under *within-disease*, *all-disease*, *and partial-pooled* approaches.



**Figure**: Schematic of machine-learning approaches. ML algorithm ingests biothreat condition data and makes predictions based on training from previous outbreak data. In pure ML approach, current outbreak dynamics are used as ML predictors of outbreak impacts. In hybrid approach, ML algorithm predicts parameters of an epidemiological model, which predicts impacts using parameters and current outbreak dynamics.

Model variables will be analyzed at both model-wide and individual case level to determine their contribution to performance. First, to evaluate model-wide predictive performance, we will re-fit models successively dropping each predictor to determine which provide the greatest improvement in performance. At the individual case level, we will calculate Shapley Additive Explanation (SHAP) scores<sup>13</sup> for all variables. This will allow for inspection of the role of variables across major categories (pathogen type, continent), and determine how accuracy varies across these contexts. Clusters of similar SHAP scores will also allow us to identify clusters of diseases or conditions that have similar behavior or performance accuracy. Such information will ultimately support decision-making in selecting models or variables to select for prediction based on the context of new outbreaks.

To measure the capacity of algorithms to predict *new*, emerging diseases, we will score using structured cross-validation in which no disease outbreak prediction is based on an algorithm

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

trained on the same pathogen. For example, for Ebola outbreak predictions, all previous Ebola outbreaks will be held out from training data.

#### Model Evaluation and Validation

Models will be fit and tested against their ability predict (a) size of an outbreak (total number of cases), (b) peak size of outbreak (number of cases at highest point), (c) time to completion of an outbreak, (d) time to peak of an outbreak, and (e) geographic spread of outbreak (distance to farthest case from point of origin). Prediction scores will consist of mean-square-error of each of these prediction against true values, as well as a score of equal-weighted errors of all five measures. All models will also be tested for fidelity of uncertainty measures—the fraction of true data values that fall within prediction confidence intervals—and these metrics will be reported along with predictive performance.

All model performance scores will be compared against meaningful, "practical naïve" baselines. While it is common practice to use a "null" or "zero-knowledge" baseline, this often amounts to inflating the apparent power of the algorithm. We will opt for a more realistic, "practical naïve" baseline, which instead follows what an analyst would approximate in practice. For these models, we will use a practical baseline model of expecting outbreaks to follow the average size and timing of outbreaks of the same disease in the past 25 years. All model scores will be reported as percentage improvements over this baseline.

Algorithm performance will be measured against holdout data to validate true predictive performance. Repeat testing on holdout data can lead to overfitting, with predictions too specific to the hold-out set and not indicative of future performance. Yet increasing holdout reduces the availability of training data. Thus, we will use a *differential privacy* approach drawn from information security to validate prediction results. The hold-out data will be obfuscated, and validation performance will be measured against bootstrapped replicates of validation data, essentially introducing noise to reduce overfitting against the hold-out. This will ensure accurate estimates of predictive performance in an operational environment. 10

#### Data Sources

Data sources for fitting and testing models include output data of number, timing, and geography of disease outbreaks, and predictor data include pathogen traits (disease symptoms, pathogen traits, modes of transmission), pathogen sources (point of origin, geographic range of potential natural hosts), environmental context (climatic conditions, ecosystem traits), and socioeconomic context (local population densities, demographics, transportation networks, healthcare systems). Gathering predictor data is often as large a factor in the time required to create first-pass predictions at the early stage of an outbreak. Thus, we will create automated pipelines for gathering, merging, and updating these data sets, and to extract key predictors from basic location and host variables to minimize manual inputs required. For instance, by linking to high-resolution geospatial databases, we will automatically populate data for population density,

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

ecosystem types, transport infrastructure that can be updated and refined with on-the-ground data if available.

Disease outbreak impact data: We will draw a variety of sources to build a database of several thousand of outbreak time-series to train our models. First, we will use time series from ΛΙDO, the most comprehensive such repository, with approximately 700 series across 40 diseases. Recognizing that ΛΙDO has limits and data gaps, we will supplement this with several other sources including One Health disease datasets compiled and maintained by ΕΗΛ:

- COVID-19 Data: Data on cases, hospitalization, and deaths from COVID-19 are uniquely
  available at national, sub-national and city scales for a broad swath of countries with varying
  conditions and policy responses. While the COVID-19 pandemic is linked across countries,
  in our hierarchical modeling framework we will to separate these out into multiple regional
  outbreaks, shedding more light on the effects of environmental and socioeconomic factors
  under a common host and pathogen.
- Multispecies datasets: We will supplement human disease outbreaks with data on veterinary disease outbreaks. EHA maintains a database on veterinary disease outbreaks derived from OIE and FAO data and supplemented by digitized archival reports (extending back to 1992), with 2900 outbreak records across 122 diseases. This will more than quadruple the available disease time series for training. Partial-pooling will ensure that only information in relevant veterinary data sets will be applied to human outbreaks and vice versa. For instance, effects of population density or health/veterinary capacity are expected to have partial correlation across all diseases. In addition, this will allows the model to predict impacts from these diseases, which has direct defense applications given the food supply, trade, and political stability impacts of veterinary diseases
- Emerging Infections Disease Repository: EHA's Emerging Infectious Disease Repository (eidr.ecohealthalliance.org/)<sup>14</sup> contains data on an additional 350 zoonotic disease emergence outbreaks, giving additional insights in particular to new emergence events from zoonotic sources. Full time-series are incomplete but outbreak size and duration are recorded, allowing them to be integrated into our framework with multiple imputation.

*Predictor variables*: Disease dynamics depend on a complex interplay of pathogen, host, and environmental conditions. We will use data on pathogens (traits, genomes), sources (reservoir host traits, range, relatedness to humans), environmental context (climate, ecosystems traits), and socioeconomic context (local population densities, transportation networks, healthcare systems) as predictors for our models, as well as time-series data of infections from the earliest phase of the outbreaks. As we expect all data will be available for neither our full training set nor in common condition of new outbreaks, use multiple imputation<sup>12</sup> in all models to allow for prediction as well as uncertainty assessment under incomplete data.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

- Pathogen Trait Data consists of information about pathogen taxa, transmission mode, structural, and biological traits. Relevant traits for pathogens, and have been largely aggregated for viruses by our Host-Pathogen Phylogeny Project<sup>5</sup> and parallel projects for bacterial pathogens<sup>15</sup> and non-zoonotic human viruses<sup>16</sup>.
- Pathogen Genetic Data consist of genetic sequence of representative samples of individual pathogen species. Representative sequences of nearly all relevant pathogens are available via genetic repositories GenBank, ICTV, or GISAID. We expect pathogen data at the scale of pathogen species to largely predict similar information as pathogen traits, but genomic data may be faster to use in cases of rapid sequencing of novel pathogens. We will test whether incorporating genetic data into model features via neural-network autoencoders<sup>17</sup> provides improved performance over separate representation of pathogen traits (e.g., taxonomic classification, transmission mode), or other representations of genetic data such as phylogeny. Whether this performance is additive, or whether these two types of information are redundant. Even if redundant, genetic data may improve predictive capacity in the case of new pathogens where pathogen traits have not been fully delineated. Rich inputs from genetic data will be integrated using neural-network encoders.
- Source Data refers to intelligence about the origin of the disease, which includes location and mode of origin (environmental, zoonotic host species, laboratory or accidental release). In many cases source data is unknown, particularly in the early stages of a new outbreak event. For zoonotic-origin diseases, the pool of known host species and their geographic provide probabilistic information on the source. Our own Host-Pathogen Phylogeny Project aggregates host traits for all known mammalian diseases hosts.<sup>5</sup>
- Environmental Data refers to information describing the conditions in which the disease may spread. These will include local climate variables from the BIOCLIM database, land-use coverage (HYDE), and human and livestock population densities (World Bank, FAO).
- Socioeconomic Data these include economic activity, governance, medical and veterinary capacity (World Bank, FAO), and transportation network connectivity in the region of disease spread.

#### **Programmatics**

All work will be undertaken by the project prime, EcoHealth Alliance (EHA). EHA has a proven track record of project management with U.S. government agencies including USAID, DTRA, DHS, NIH, and NSF, ranging from global epidemiological research and surveillance programs to small-to-medium scale technology development.

Our management approach is to bring production-quality software development practices to data science for research and development. We use agile development methodology, with weekly development meetings, monthly project status reviews, and regular organization-wide review seminars to gather feedback from our broader interdisciplinary expertise. All tasks in the

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

statement of work will benefit from the significant experience of the project staff and in-kind contribution from other EHA personnel with combined expertise in global emerging infectious disease epidemiology, public health, veterinary medicine, machine learning, and biostatistics. We also focus on *deployment-ready* research infrastructure to minimize the need to re-engineer models developed in the research process for applied use.

We engage multiple best practices to minimize project disruption risk. *Personnel risk* entails loss of expertise and project or institutional knowledge due to loss of core project staff. EHA maintains project data, documentation, and code in common cross-team repositories. Version-controlled project repositories ensure that project histories are tracked, and all team members can retrieve previous work by others. Through regular code review, we ensure all team members have working knowledge of project modules and the ability to re-task. *Infrastructure risk entails* loss of project infrastructure such a. EHA performs on-site high-performance computing cluster or data storage. We keep code, data, results and documentation on-site only for active experiments, and back up these to off-site cloud infrastructure. Our compute environment is reproducible, being configured by pre-programmed, self-documenting methods such as Dockerfiles, Ansible playbooks, and Bash scripts. It can be re-deployed to off-site cloud resources or new hardware as needed.

#### Relevance

Our research will fundamentally improve infectious disease forecasting by identifying the relative importance of early-stage factors in outbreak prediction. While pathogen-host-environment linkages are fundamental in disease modeling, predictive methods developed to assess emerging biothreats usually focus on each individually. We will also advance knowledge of hybrid ML/mechanistic models, which have not yet been used in applied disease forecasting. Together this research will impact C-WMD science by providing practical guidance for early-stage data-gathering and prototyping an interactive decision support tool for biothreat impact forecasting.

#### **Credentials**

EcoHealth Alliance (EHA) will implement this project, with Noam Ross, Principal Scientist for Computational Research, as project lead. EHA's mission is focused on emerging diseases and cutting-edge research into the critical connections between human, wildlife, and ecosystem health that drive disease emergence and spread.

**PI:** Dr. Ross will direct he project. He is a computational disease ecologist who provides expertise in dynamic epidemiological processes, machine learning and biostatistics, and high-performance computing. He has led work in ecological and epidemiological modeling and data management for EHA's work on USAID PREDICT and NIH NIAID viral surveillance research. He has experience developing livestock disease outbreak and spread simulations in collaboration with FAO as well as private industry. Dr. Ross is PI for EcoHealth's work with DHS on machine learning for veterinary disease outbreaks. He also a leader in open-source scientific computing.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

He is a founding editorial board member of rOpenSci, an organization that defines standards for reproducibility and runs peer review for scientific software, and is a core contributor to the Rocker project, which provides open-source, standardized computing containers for data science. Dr. Ross holds a Ph.D. in computational ecology from the University of California-Davis.

**Organization:** EHA has a proven track record of large project management with U.S. government agencies. We lead the largest global country portfolio, and serve as lead for Modeling and Analytics and Behavioral Risk Analysis teams, for the USAID Emerging Pandemic Threats PREDICT program, a \$118 million program focused on predicting and preventing pandemic diseases. The PREDICT project is building a global early warning system to detect and reduce the impacts of emerging diseases that move between wildlife and people (zoonotic diseases). EHA also leads multiple field epidemiological contracts funded by DTRA, NIH, and NSF.

EHA has proven expertise in surveillance-oriented technology development. We managed a five-year contract to develop applications for DTRA's Chemical Biological Integration Platform (CBIP, formerly the Biosurveillance Ecosystem—BSVE), as well as the development of the Inbound Bioevent Information System (IBIS), and Rapid Evaluation to Prevent Epidemics in Livestock (REPEL) project under DHS's Analytics for Prevention of Biothreat Incidents (APBI) program and Science and Technology Directorate. Through this work, EHA scientists have gained substantial expertise in both basic research and working with government project managers and analysts to develop technologies that are in line with the goals of the funding. The expertise resulting from this effort serves as the foundation for the current proposal.

**Team:** Robert Young, Senior Research Software Engineer, will lead project infrastructure development and the creation ML pipelines. Mr. Young has a background in artificial intelligence and bioinformatics. He has developed and managed laboratory information systems in neuroscience, genomics and performed research in neural-network design for image-recognition in astronomy. Mr. Young currently leads cyperinfrastructure work for EHA's veterinary disease forecasting under DHS as well as the DTRA-Funded Western Asia Bat Research Network.

Emma Mendelsohn, M.Sc., Research Scientist is a data scientist with extensive expertise in dynamic systems modeling, biostatistics, web application development and data engineering. Emma is lead analyst for EHA's work on antimicrobial resistance emergence work as well as projects linking emerging infectious diseases with nutrition and non-communicable diseases. Emma will lead data engineering and contribute expertise to algorithm development and application design.

Peter Daszak, Ph.D., President of EcoHealth Alliance will provide oversight and act as project advisor on infectious disease spread. Dr. Daszak is a leader in the field of emerging diseases and disease ecology.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

William Karesh, DVM, Senior Vice President of EcoHealth Alliance will serve as project advisor on veterinary topics as well as liaise with data-providing organizations, such as OIE and ProMED, as required. Dr. Karesh is President of the Working Group on Animal Health and COVID-19 Task Forces for OIE, Co-Chair of the Wildlife Health Specialist Group of the IUCN.

**Facilities:** EcoHealth Alliance is a 48-year old scientific research NGO that specializes in multidisciplinary research and surveillance of the spread of zoonotic emerging diseases. EcoHealth Alliance is based in New York City with 10,000 ft<sup>2</sup> of office space. The scientific and technology staff of 30 are supported by a core administrative staff of 8 people who are available for work on this project and are funded through core funds.

EcoHealth Alliance is equipped with fiber optic Internet access and video conferencing facilities to facilitate easy communication between collaborators. EcoHealth Alliance employees have around the clock access to servers, VPNs, encryption software, and IT support. We use standard office suite services including Google Apps (Hosted email and web-based collaboration software), Microsoft Office, and Slack, running on Apple macOS, Ubuntu Linux, and Windows Operating Systems. EHA teams work in multiple computer languages (Python, NodeJS, and R programming languages), development frameworks (Agile, continuous integration), and machine-learning toolboxes. Our development operations are based on reproducible, containerized frameworks (Ansible, Docker). We perform high-performance computation across our dedicated cluster of System76 multi-core serversand with NVIDIA deep-learning GPUs and spot cloud somputs instances through Amazon Web Services EC2 and Microsoft Virtual Machines, We use a variety of other cloud services for storage (AWS S3, Azure Blob), continuous integration (GitHub, Gitlab), and other tasks.

#### Work to be Performed

Tasks in Year. Task. Subtask format

#### **YEAR 1:**

**Task 1.1 – Develop and deploy project infrastructure:** Infrastructure engineering consists of provisioning a best-in-class computational environment that enables continuous model development and testing and allows for research and models to be reproduced and extended by external teams. It includes continuous maintenance and documentation of project codebase.

- 1.1.1 Layout a provision project network
- 1.1.2 Set up continuous integration pipelines
- 1.1.3 Implement backup and data verification jobs
- 1.1.4 Implement differential privacy testing
- 1.1.5 Write and maintain codebase documentation
- 1.1.6 Perform code review, testing and validation

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

Task 1.2 – Engineer Project Database: Data engineering consists of curating, integrating and cross-linking and validating relevant data sets from various sources into a project database. This allows data with different sources, space and time scales, formats, and completeness to be used in common models establishes a database that can be used as a common resource.

#### Subtasks

- 1.2.1 Design database schema
- 1.2.1 Write pipelines to ingest time-series data
- 1.2.2 Write pipelines to ingest predictor data
- 1.2.3 Cross-link data sources via common ontologies
- 1.2.4 Write quality assurance/quality control (QA/QC) tests for data and perform data cleaning
- 1.2.5 Create automatic extraction routines to populate relevant predictors from location information
- 1.2.6 Update data from live sources and clean and maintain data continuously

**Task 1.3 – ML Model Development and Testing:** The core ML development task consists of designing, implementing, testing, and analyzing performance of ML model structures and iteratively improving them.

#### Subtasks

- 1.3.1 Implement "practical naïve" model baseline
- 1.3.2 Design Bayesian additive regression tree (BART) model structure
- 1.3.3 Design alternative generalized additive model (GAM) structure
- 1.3.4 Implement within-disease, all-disease, and partial-pooled disease model versions
- 1.3.5 Measure variable contribution by re-fitting
- 1.3.6 Analyze prediction-level variable contributions and model performance
- 1.3.7 Iteratively update models to improve performance

#### **OPTION YEAR:**

Task 2.1 – Develop and deploy project infrastructure: In the option year we will continue to perform maintenance and documentation of project codebase to ensure reproducibility and reusability of additional components

#### Subtasks

- 2.1.1 Write and maintain codebase documentation
- 2.1.2 Perform code review, testing and validation
- **Task 2.2 Engineer Project Database:** In the option year we will extend the database to include additional predictors and continuously maintain and update to accumulate more data from upstream sources.

- 2.2.1 Extend database schema to include genetic data and auto-encoded variables
- 2.2.2 Write pipelines to ingest genetic data
- 2.2.3 Cross-link data sources via common ontologies
- 2.2.4 Write QA/QC tests for data and perform data cleaning

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

2.2.5 Update data from live sources and clean and maintain data continuously

Task 2.3 – BART Model Development and Testing: In the option year we will extend the BART model to ingest genetic data and test the contribution of this data in enhancing or recapitulating pathogen trait data.

#### Subtasks

- 2.3.1 Develop genetic predictor variable set via neural-network autoencoding
- 2.3.2 Modify BART and GAM structures to incorporate genetic variables
- 2.3.3 Implement within-disease, all-disease, and partial-pooled disease model versions
- 2.3.4 Compare performance of genetic, pathogen-trait, and combined models

Task 2.4 – Hybrid ML-Mechanistic Model Development and Testing: In the option year will develop the hybrid machine-learning mechanistic model, compare its performance to a pure ML model and determine the switch-over time when the mechanistic model becomes more predictive in the course of the outbreak.

- 2.4.1 Design mechanistic model structure
- 2.4.2 Determine mechanistic model prior forms
- 2.4.3 Modify ML model to predict mechanistic parameters rather than outcomes
- 2.4.4 Implement within-disease, all-disease, and partial-pooled disease model versions
- 2.4.5 Measure variable contribution by re-fitting
- 2.4.6 Analyze prediction-level variable contributions and model performance
- 2.4.7 Compare Hybrid and ML performance along outbreak time-series progression
- 2.4.8 Iteratively update models to improve performance

PI: Noam Ross

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

# Performance Schedule

Description	Y1 Q1-2	Y1 Q3-4	OY Q1-2	OY Q3-4
Task 1: Develop and deploy project infrastructure				
1.1.1 Layout a provision project network				
1.1.2 Set up continious integration pipelines				
1.1.3 Implement backup and data verification jobs				
1.1.4 Implement differential privacy testing				
1.1.5 Write and maintain codebase documentation				
1.1.6 Code review, testing and validation				
2.1.5 Write and maintain codebase documentation				
2.1.6 Perform code review, testing and validation				
Task 2: Engineer Project Database				
1.2.1 Design database schema				
1.2.1 Write pipelines to ingest time-series data				
1.2.2 Write pipelines to ingest predictor data				
1.2.4 Cross-link data sources via common ontologies				
1.2.4 Write QA/QC tests for data and perform data cleaning				
1.2.5 Create automatic extraction routines to populate relevant predictors				
1.2.6 Update data from live sources and clean and maintain data continuously				
2.2.1 Extend database schema to include genetic data				
2.2.2 Write pipelines to ingest genetic data				
2.2.3 Cross-link data sources via common ontologies				
2.2.4 Write QA/QC tests for data and perform data cleaning				
2.2.5 Update data from live sources and clean and maintain data continuously				
Task 3: ML Model Development and Testing				
1.3.1 Implement "practical naïve" model baseline				
1.3.2 Design BART model structure				
1.3.3 Design alternative GAM model structure				
1.3.4 Implement within-disease, all-disease, and partial-pooled disease models versions				
1.3.5 Measure variable contribution by re-fitting				
1.3.6 Analyze prediction-level variable contributions and model performance				
1.3.7 Iteratively update models to improve performance				
2.3.1 Develop genetic predictor variable set via neural-network autoencoding				
2.3.2 Modify BART and GAM structures to incorporate genetic variables				
2.3.3 Implement within-disease, all-disease, and partial-pooled disease models				
2.3.4 Compare performance of genetic, pathogen-trait, and combined models				
Task 4: Hybrid Model Development and Testing	ı	1		· · · · · · · · · · · · · · · · · · ·
2.4.1 Design mechanistic model structure				
2.4.2 Determine mechanistic model prior forms				
2.4.3 Modify ML model to predict mechanistic parameters				
2.4.4 Implement within-disease, all-disease, and partial-pooled models				
2.4.5 Measure variable contribution by re-fitting				
2.4.6 Analyze prediction-level variable contributions and model performance				
2.4.7 Compare Hybrid and ML performance along outbreak time-series progression				
2.4.8 Iteratively update models to improve performance				

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

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# RESEARCH & RELATED BUDGET - Budget Period 1

OMB Number: 4040-0001 Expiration Date: 12/31/2022

ORGANIZATIO	ONAL DUNS:	077090066	cccc	Enter name of Or	ganization:	Ecolie:	alth Al	liance				
Budget Type:	⊠ Project	Subawa	ard/Consortium	ו		Budget	Period:	1 S	tart Date:	07/01/2020	End Date: 06/30/202	1
A. Senior/Key	/ Person											
Prefix	First	Middle	Last	Suffix	Bas	e Salary (\$	s) C	Month al. Acad	-	Requested Salary (\$)	Fringe Benefits (\$)	Funds Requested (\$)
Dr.	Noam		Ross				3.00			25,000.0	9,200.00	34,200.00
Project Role	: PD/PT											
	Robert		Yeung				6.00			52,500.0	0 19,320.00	71,820.00
Project Role	: Senier Engi	neer										
	Emma		Mendelsohn				6.00			42,500.0	0 15,640.00	58,140.00
Project Role	: Research Sc	ientist										
Additional Senio	or Key Persons:			Add	Attachment						requested for all Senior sons in the attached file	
										7	otal Senior/Key Person	164,160.00
B. Other Pers	onnel											
Number of Personnel	Project	Role				Cal.	Months Acad.	Sum.		quested	Fringe Benefits (\$)	Funds Requested (\$)
	Post Doctoral	Associates										
	Graduate Stud	dents			Ī							
	Undergraduate	e Students			Ī							
	Secretarial/Cle	erical			[							
	Total Number 0	Other Personr	nel								Total Other Personnel	
								Total S	Salary, W	ages and Fri	nge Benefits (A+B)	164,160.00

C. E	Equipment Description		
	items and dollar amount for each item exceeding \$5,000 Equipment item		Funds Requested (\$)
Addit	itional Equipment:	Add Attachment	
	Total funds requested for all equip	oment listed in the attached file	
		Total Equipment	
D. Tı	<b>Fravel</b>		Funds Requested (\$)
1. [	Domestic Travel Costs (Incl. Canada, Mexico and U.S. Possessions	)	1,576.00
2. I	Foreign Travel Costs		7,088.00
		Total Travel Cost	8,664.00
E. Pa	Participant/Trainee Support Costs		Funds Requested (\$)
1.	Tuition/Fees/Health Insurance		
2. 5	Stipends		
3.	Travel		
4. 5	Subsistence		
5. (	Other		
	Number of Participants/Trainees Total Par	ticipant/Trainee Support Costs	

F.	Other Direct Costs			Funds Requested (\$)
1.	Materials and Supplies			7,000.00
2.	Publication Costs			1,500.00
3.	Consultant Services			
4.	ADP/Computer Services			
5.	Subawards/Consortium/Contractual Costs			
6.	Equipment or Facility Rental/User Fees			
7.	Alterations and Renovations			
8.				
9.				
10.				
			Total Other Direct Costs	8,500.00
G. I	Direct Costs			Funds Requested (\$)
		Total Dir	rect Costs (A thru F)	181,324.00
<u>H. I</u>	ndirect Costs			
	Indirect Cost Type	Indirect Cost Rate (%)	Indirect Cost Base (\$)	Funds Requested (\$)
	EceHealth Alliance's NICA rate	35.84	181,324.00	64,986.52
_			Total Indirect Costs	64,986.52
	nizant Federal Agency ncy Name, POC Name, and			
POC	Phone Number)  Department of Defense, De	partment of the Navy	, Sharon Gales, (1.763	. עככא. טעט.
<u>I. Te</u>	otal Direct and Indirect Costs			Funds Requested (\$)
	Total Direc	t and Indirect Institu	tional Costs (G + H)	246,310.52
<u>J. F</u>	ee		1	Funds Requested (\$)
<u>K. 1</u>	Total Costs and Fee		<b></b>	Funds Requested (\$)
		Total (	Costs and Fee (I + J)	246,310.52
<u>L. E</u>	Budget Justification			
(Onl	y attach one file.) 1234 DTRA MI Budget Just if ignt	ior.pdf	Delete Attachmer	nt View Attachment

# RESEARCH & RELATED BUDGET - Budget Period 2

OMB Number: 4040-0001 Expiration Date: 12/31/2022

ORGANIZATIO	ONAL DUNS:	077090066	cccc	Enter name of Orga	ınization:	Ecolleal	lth All	iance				
Budget Type:	∑ Project	Subawa	ard/Consortiun	n	В	udget F	Period:	2 S	tart Date:	07/01/2021	End Date: 06/30/202	2
A. Senior/Key	y Person											
Prefix	First	Middle	Last	Suffix	Base S	alary (\$)	Ca	Month	-	Requested Salary (\$)	Fringe Benefits (\$)	Funds Requested (\$)
Dm.	Noam		Ross				3.00			26,250.00	9,640.00	35,910.00
Project Role	: PD/PI											
	Robert		Young				6.00			55,125.00	20,286.00	75,411.00
Project Role	: Senior Engi	neer										
	Emma		Mendelsohn				6.00			44,625.00	16,422.00	61,047.00
Project Role	: Research Sc	ientist										
Additional Senio	or Key Persons:			Add At	tachment						requested for all Senior cons in the attached file	
										Т	otal Senior/Key Person	172,368.00
B. Other Pers	sonnel											
Number of Personnel	Project	Role			c		lonths \cad.	Sum.		quested lary (\$)	Fringe Benefits (\$)	Funds Requested (\$)
	Post Doctoral	Associates										
	Graduate Stud	dents										
	Undergraduate	e Students										
	Secretarial/Cle	erical										
	Total Number (	Other Personr	nel								Total Other Personnel	
								Total S	Salary, W	ages and Frii	nge Benefits (A+B)	172,368.00

C. Eq	uipment Description		
	ems and dollar amount for each item exceed quipment item	ing \$5,000	Funds Requested (\$)
Additio	onal Equipment:	Add Attachment	
	Total funds re	equested for all equipment listed in the attached file	
		Total Equipment	
D. Tra	avel		Funds Requested (\$)
<b>1.</b> D	omestic Travel Costs (Incl. Canada, Mexico an	d U.S. Possessions)	5,996.00
2. F	oreign Travel Costs		
		Total Travel Cost	5,996.00
E. Pa	rticipant/Trainee Support Costs		Funds Requested (\$)
1. T	uition/Fees/Health Insurance		
<b>2.</b> S	stipends		
<b>3</b> . T	ravel		
<b>4.</b> S	Subsistence		
<b>5.</b> O	Other		
	Number of Participants/Trainees	Total Participant/Trainee Support Costs	

F.	Other Direct Costs	Funds Requested (\$)
1.	Materials and Supplies	3,000.00
2.	Publication Costs	1,500.00
3.	Consultant Services	
4.	ADP/Computer Services	
5.	Subawards/Consortium/Contractual Costs	
6.	Equipment or Facility Rental/User Fees	
7.	Alterations and Renovations	
8.		
9.		
10.		
	Total Other Direct Costs	4,500.00
G.	Direct Costs	Funds Requested (\$)
	Total Direct Costs (A thru F)	182,864.00
H. I	Indirect Cost Type Indirect Cost Rate (%) Indirect Cost Base (\$)	Funds Requested (\$)
	EcoHealth Alliance's NICA rate 35.84 182,864.00	65,538.46
	Total Indirect Costs	65,538.46
	nizant Federal Agency	-
	Department of Defense, Department of the Navy, Sharon Gales, (1.76)	3.696.8559
. т	otal Direct and Indirect Costs	Funds Requested (\$)
	Total Direct and Indirect Institutional Costs (G + H)	248,402.46
J. F	Fee	Funds Requested (\$)
<u>۷.</u> '	Total Costs and Fee	Funds Requested (\$)
	Total Costs and Fee (I + J)	248,402.46
E	Budget Justification	
Onl	y attach one file.) 1234 DTRA MI Budger Tust if ignt jon, pdf	

# RESEARCH & RELATED BUDGET - Cumulative Budget

Ocation A Oct. Nr	Totals	(\$)
Section A, Senior/Key Person		336,528,00
Section B, Other Personnel		
Total Number Other Personnel		
Total Salary, Wages and Fringe Benefits (A+B)		336,528.00
Section C, Equipment	Ī	333,333,000
Section D, Travel		14,660.00
1. Domestic	7,572.00	
2. Foreign	7,088.00	
Section E, Participant/Trainee Support Costs		
1. Tuition/Fees/Health Insurance		
2. Stipends		
3. Travel		
4. Subsistence		
5. Other		
6. Number of Participants/Trainees		
Section F, Other Direct Costs		13,000.00
1. Materials and Supplies	10,000.00	13,000.00
2. Publication Costs	3,000.00	
3. Consultant Services	3,000.00	
4. ADP/Computer Services		
5. Subawards/Consortium/Contractual Costs		
6. Equipment or Facility Rental/User Fees		
7. Alterations and Renovations		
8. Other 1		
9. Other 2		
10. Other 3		
Section G, Direct Costs (A thru F)		
Section H, Indirect Costs	<u> </u>	364,188.CC
Section I, Total Direct and Indirect Costs (G + H)		130,524.98
Section J, Fee		494,712.98
Section K, Total Costs and Fee (I + J)		
•		494,712.98

#### BUDGET JUSTIFICATION FOR ECOHEALTH ALLIANCE

EcoHealth Alliance requests a total of \$4,975,681 over all-years of the proposed project to support personnel, travel, equipment, consortium agreements, and applicable indirect costs.

#### A. Key Personnel

Noam Ross, Ph D., Principal Investigator (3.0 calendar months for all years). Dr. Ross will oversee the project and be responsible for project management, research, development, design, analysis, and manuscript writing and conference presentations, as well as communication with DTRA. We request \$25,000 in Y1 due to the high cost of living in New York City and a 5% cost of living increase in OY1.

Emma Mendelsohn, M.Sc, Research Scientist. (6.0 calendar months for all years). Ms. Mendelsohn will lead data engineering and contribute expertise to algorithm development. She will contribute to manuscript writing and conference presentations. We request \$42,500 in Y1 with a 5% cost of living increase in OY1

Robert Young M.Sc, Senior Research Software Engineer. (6.0 calendar months for all years). Mr. Young will lead infrastructure engineering and contribute expertise to algorithm development. He will contribute to manuscript writing and conference presentations. We request \$52,500 in Y1 with a 5% cost of living increase in OY1.

#### Fringe Benefits

Fringe benefits are calculated as 36.80% of base salary p.a. with \$44,160 requested in Y1 calculated from the base salary for all personnel. In OY1, we budget for a 5% per year cost of living allowance increase in all salaries.

#### B. Other Personnel

No funds are requested for other personnel.

#### C. Equipment

No funds are requested for equipment.

#### D. Travel

Per recommendations in the BAA, we request travel funds to Nationally Recognized Conferences and the Annual Technical Review.

#### Domestic Travel

Domestic travel is requested for one trip per year from New York City per year to Northern Virginia the Annual Technical Review. Transportation is estimated at \$350 per trip along with the government GSA per diem rates for Northern Virginia (\$161 for accommodation and \$71 for meals and incidentals). Additional domestic travel support will facilitate two EcoHealth Alliance team members presenting at one conference. In OY1, this will be the EEID conference held in Bozeman, MT in 2022, with transportation costs of \$500, \$225 for accommodation and \$55 for meals and incidentals. For each trip, \$25 per day is estimated in taxi costs and for those that

require transport to the airport \$150 RT is included. In total, we are requesting \$1,576 in Y1 and \$5,996 in OY1 for domestic travel

#### International Travel

One international trip for two team members to present at the EEID conference held in Montpelier, France, in 2021 requested for Y1. Flights are estimated at \$1500 and the federal per diem rates of \$196 for lodging and \$155 for meals in Montpelier. \$25 per day is estimated in taxi costs plus \$150 RT trips to airport is included. In total we are requesting \$7,088 in Y1.

#### E. Participant/Trainee Support Costs

No funds are requested for participant or trainee support costs.

#### F. Other Direct Costs

Materials and Supplies

PI Ross and Research Scientist Mendelsohn will both need a new computer, equipped with Microsoft software for which we request a total of \$4,000, in only \$11.

# Cloud Computing and Storage Services

We will make use of cloud computer services including spot server rental for compute-intensive jobs, data storage, continuous integration and testing of computer code. We request \$2,000 per year in Y1-OY1.

#### Team Communication and Collaboration Software and Services

We request \$1,000 per year for subscriptions for software and services for document sharing, chat, voice, and video communications.

#### Publication Costs

To facilitate the dissemination of project findings, \$1,500 per year in only Y1-OY1 is requested for open access publication costs in international peer-reviewed journals.

#### H. Indirect Costs

We are requesting the EcoHealth Alliance federally approved indirect cost rate of 35.84% on all applicable direct costs. The USA Department of Defense's Department of the Navy has approved this rate on 01 April 2020. We request \$ \$64,986.52 in Y1 and \$65,538.46 in OY1.



# Predicting Biothreat Impacts from Early-Stage Data via Transfer Learning Dr. Noam Ross, EcoHealth Alliance, HDTRA1-14-24c-FRCWMD-BAA

**Objective:** Test new ML approaches - transfer learning by partial pooling, and hybrid mechanistic-ML models, for predicting impacts of novel biothreats at early, data-sparse stage of outbreak.

#### Method:

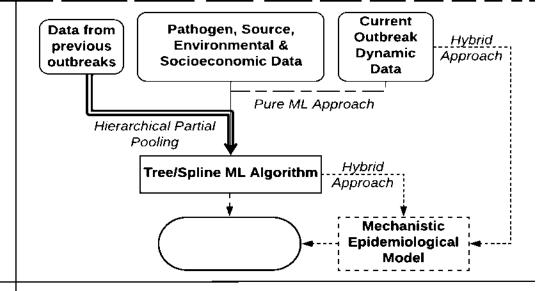
- Quantify gain in prediction by training models to learn information across diseases (partial pooling)
- Assess variables most predictive of epidemic impacts at early stages outbreak.
- Quantify gain in power from using model to predict parameters of mechanistic compartmental model.
- Determine point in epidemic growth where hybrid models become more accurate than pure ML models.

**Benefits of Technology:** Disease forecasts in early stages of outbreaks, especially novel outbreaks, require strong assumptions in highly structured models. Hierarchical partial pooling learns what factors from previous outbreaks may be transferred to apply to new predictions, improving early-stage assessments of severity and risk.

# Challenges:

- Missing data in early-stage outbreak conditions → Learn most important variables to prioritize data collection
- ML models non-interpretable and poor at extrapolation →
  Use hybrid approach with ML parameterizing structured
  mechanistic model capable of simulating scenarios

Readiness: TRL1



#### Milestones:

- Y1Q2: Engineer project database & test infrastructure
- Y1Q4: Build, fit, and compare ML model structures to
- OY1Q2: Extend models to incorporate genetic data
- OY1Q4: Compare pure ML models to ML-mechanistic hybrids

# **Funding Profile:**

\$246,211 Year 1, \$248,402 Option Year

# **Contact information:**

Noam Ross, Senior Research Scientist, ross@ecohealthalliance.org, 1.212.380.4471



Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

#### Attachment 1 - Statement of Work (SOW)

Project Title: Predicting Biothreat Impacts from Early-Stage Data via Transfer Learning

Document Date: May 28, 2020

#### **Objective**

The project's objectives are to determine the efficacy of machine learning (ML) techniques in predicting epidemic impacts in early-stage, data-sparse conditions, particularly novel biothreats, both intentional (e.g. novel genetically-modified pathogens) and natural (e.g. zoonotic). Specifically, we will answer three questions that will enable rapid decision-making based on ML models of epidemic impacts:

**Q1:** How much does partial-pooling improve predictions of epidemic impacts, and what information transfers best across diseases?

**Q2:** What variables are most predictive of epidemic impacts at early stages of an outbreak?

**Q3:** At what point in epidemic growth do hybrid models that incorporate mechanistic epidemiological components become more accurate than pure ML models?

This work will contribute to fundamental knowledge of properties and variables that contribute most to predictive accuracy at early-stage stages in emerging outbreak prediction and forecasting. It will enable the development of robust and deployable predictive models at early stages of outbreaks, and support informed choices for data collection and model selection.

#### Scope

The awardee proposes a two-year study to determine the efficacy of ML techniques in predicting epidemic impacts in early-stage, data-sparse conditions, particularly novel biothreats. The awardee team shall focus on the following major goals and milestones:

- Year 1, Milestone 1 (6 months): Create and curate predictor database, project infrastructure, continuous integration/validation, establish data-imputation and automatic predictor workflow (Tasks 1.1-1.2).
- Year 1, Milestone 2 (12 months): Bayesian additive regression tree (BART) model fitting and tuning, determination of variable contribution to prediction accuracy (Task 1.3)
- Option Year, Milestone 3 (18 months): Maintain and document project infrastructure, extend the database to include additional predictors, extend the BART model to ingest genetic data (Tasks 2.1-2.3)
- Option Year, Milestone 4 (24 months): Construction of hybrid ML/mechanistic model structure, comparative testing of pure and hybrid version, determination of timing when current data overtakes transfer data (Task 2.4)

# Background

Situational awareness and risk assessment are critical at early stages of disease outbreaks, when interventions can best mitigate adverse outcomes. Yet decision-makers often face a paucity of data at these stages. Traditional modeling methods to forecast disease impact severity rely heavily on prior knowledge or assumptions about mechanisms of disease spread progression, and they frequently do not anticipate the actual mechanisms.<sup>1-3</sup> They also take considerable time and effort to develop.

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

Recent theory suggests disease dynamics are fundamentally predictable<sup>4</sup>, and ML methods have been successfully applied in emerging disease science. ML refers to a group of methods, ranging from simple nearest-neighbor approaches to function-approximation methods such as boosted trees and penalized splines, to complex deep-learning neural networks. ML has been used to predict the likelihood of disease emergence and the zoonotic potential of various hosts and viruses<sup>5,6</sup>, and deep-learning approaches have been used to infer viral hosts from the comparatively richer data of genetic sequences of viruses.<sup>7,8</sup>

ML methods are generally unstructured, flexible, and offer greater predictive power over traditional statistical approaches. However, these methods are data-intensive, and a key challenge with ML development is data paucity and/or a low signal-to-noise ratio. Surveillance data for even existing, well-studied diseases is rarely rich enough for the data scale of many modern ML techniques. For instance, a ML approach to predict impacts from a new outbreak of Ebola would draw training data from fewer than twenty previous events, each with varying levels of data coverage.

Transfer learning, or data fusion, provides a solution to data paucity issues by using data from other domains to train parts of models that are general, thus limiting new data requirements for more specific components. Hierarchical modeling, or *partial pooling*, is a transfer learning approach for partitioning out which data can be shared between models and which are only applicable to a specific case. For instance, Ebola forecasts for countries outside the historic range of the disease may be improved by calibrating against Cholera outbreaks, which have occurred both within and outside countries with Ebola outbreaks. The degree to which the disease outcomes are correlated is learned, rather than assumed.

An alternative recent development in disease forecasting is the development of hybrid models that incorporate simple mechanistic structures and predictive ML methods to learn model parameters. These approaches have demonstrated high predictive accuracy for near-term forecasting of epidemic growth and impact, particularly during the COVID-19 pandemic<sup>9</sup>. However, these approaches require adequate time-series of data to project forward from current conditions. Their utility in early stages and long-term forecasting accuracy has yet to be determined. In opting to apply these models, it is critical to understand these data requirements and at what point in an outbreak hybrid approaches can overtake pure ML early-stage predictions in accuracy. This project seeks to develop, apply, and compare pure and hybrid-ML techniques to enable the development of robust, rapidly usable predictions at early stages of outbreaks.

Key references (additional references can be found in the Technical Proposal):

- 1 MacDiarmid, S. C. et al. Handbook On Import Risk Analysis for animals and animal products: quantitative risk assessment. Vol. 2 (Office International des Épizooties, 2004).
- Moura, J. A., McManus, C. M., Bernal, F. E. M. & de Melo, C. B. An analysis of the 1978 African swine fever outbreak in Brazil and its eradication. *Rev. Sci. Tech.* **29**, 549-563, doi:10.20506/rst.29.3.1992 (2010).
- Delgado, J. *et al.* U.K. Foot and Mouth Disease: A Systemic Risk Assessment of Existing Controls. *Risk Anal.* **37**, 1768-1782, doi:10.1111/risa.12704 (2017).

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

- Scarpino, S. V. & Petri, G. On the predictability of infectious disease outbreaks. *Nat. Commun.* **10**, 898, doi:10.1038/s41467-019-08616-0 (2019).
- Olival, K. J. *et al.* Host and viral traits predict zoonotic spillover from mammals. *Nature*, doi:10.1038/nature22975 (2017).
- Allen, T. *et al.* Global hotspots and correlates of emerging zoonotic diseases. *Nat. Commun.* **8**, 1124, doi:10.1038/s41467-017-00923-8 (2017).
- 7 Mock, F., Viehweger, Λ., Barth, E. & Marz, M. VIDHOP, viral host prediction with Deep Learning. *Bioinformatics* (2019).
- Babayan, S. A., Orton, R. J. & Streicker, D. G. Predicting reservoir hosts and arthropod vectors from evolutionary signatures in RNA virus genomes. *Science* **362**, 577-580, doi:10.1126/science.aap9072 (2018).
- 9 Gu, Y. COVID-19 Projections Using Machine Learning, <a href="https://covid19-projections.com/about/#historical-performance">https://covid19-projections.com/about/#historical-performance</a> (2020).

#### Data Sources

Fitting and testing models will draw from disease outbreak data (counts, timing, and geography) and predictor data including pathogen traits (disease symptoms, pathogen traits, modes of transmission), pathogen sources (point of origin, geographic range of potential natural hosts), environmental context (climatic conditions, ecosystem traits), and socioeconomic context (local population densities, demographics, transportation networks, healthcare systems). Please see the Technical Proposal for available data sources.

# Tasks/Scientific Goals (Format: Year. Task. Subtask)

#### YEAR 1

# Task 1.1 – Develop and deploy project infrastructure

The awardee shall develop a computational environment that enables continuous model development and testing and allows for research and models to be reproduced and extended by external teams. The computational environment will include continuous maintenance and documentation of project codebase.

#### Subtasks

- 1.1.1 Layout a provision project network
- 1.1.2 Set up continuous integration pipelines
- 1.1.3 Implement backup and data verification jobs
- 1.1.4 Implement differential privacy testing
- 1.1.5 Write and maintain codebase documentation
- 1.1.6 Perform code review, testing and validation

# Task 1.2 – Engineer project database

The awardee shall curate, integrate, cross-link, and validate relevant datasets from various sources into a project database, accounting for differing space and time scales, data formats, and data completeness.

- 1.2.1 Design database schema
- 1.2.1 Write pipelines to ingest time-series data
- 1.2.2 Write pipelines to ingest predictor data
- 1.2.3 Cross-link data sources via common ontologies

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

- 1.2.4 Write quality assurance/quality control (QA/QC) tests for data and perform data cleaning
- 1.2.5 Create automatic extraction routines to populate relevant predictors from location information
- 1.2.6 Update data from live sources and clean and maintain data continuously

# Task 1.3 - Machine Learning (ML) model development and testing

The awardee shall design, implement, test, and analyze performance of ML model structures and iteratively improve them.

Subtasks

- 1.3.1 Implement "practical naïve" model baseline
- 1.3.2 Design Bayesian additive regression tree (BART) model structure
- 1.3.3 Design alternative generalized additive model (GAM) structure
- 1.3.4 Implement within-disease, all-disease, and partial-pooled disease model versions
- 1.3.5 Measure variable contribution by re-fitting
- 1.3.6 Analyze prediction-level variable contributions and model performance
- 1.3.7 Iteratively update models to improve performance

#### **OPTION YEAR:**

## Task 2.1 – Maintain and document project infrastructure

In the option year, the awardee shall continue to perform maintenance and documentation of project codebase to ensure reproducibility and reusability of additional components

Subtasks

- 2.1.1 Write and maintain codebase documentation
- 2.1.2 Perform code review, testing and validation

#### Task 2.2 – Extend project database

In the option year, the awardee shall extend the database to include additional predictors and continuously maintain and update to accumulate more data from upstream sources.

Subtasks

- 2.2.1 Extend database schema to include genetic data and auto-encoded variables
- 2.2.2 Write pipelines to ingest genetic data
- 2.2.3 Cross-link data sources via common ontologies
- 2.2.4 Write QA/QC tests for data and perform data cleaning
- 2.2.5 Update data from live sources and clean and maintain data continuously

#### Task 2.3 – Extend BART model

In the option year, the awardee shall extend the BART model to ingest genetic data and test the contribution of this data in enhancing or recapitulating pathogen trait data.

Subtasks

- 2.3.1 Develop genetic predictor variable set via neural-network autoencoding
- 2.3.2 Modify BART and GAM structures to incorporate genetic variables
- 2.3.3 Implement within-disease, all-disease, and partial-pooled disease model versions
- 2.3.4 Compare performance of genetic, pathogen-trait, and combined models

## Task 2.4 - Hybrid ML-mechanistic model development and testing

Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

In the option year, the awardee shall develop the hybrid machine-learning mechanistic model, compare its performance to a pure ML model, and determine the switch-over time when the mechanistic model becomes more predictive in the course of the outbreak.

#### Subtasks

- 2.4.1 Design mechanistic model structure
- 2.4.2 Determine mechanistic model prior forms
- 2.4.3 Modify ML model to predict mechanistic parameters rather than outcomes
- 2.4.4 Implement within-disease, all-disease, and partial-pooled disease model versions
- 2.4.5 Measure variable contribution by re-fitting
- 2.4.6 Analyze prediction-level variable contributions and model performance
- 2.4.7 Compare Hybrid and ML performance along outbreak time-series progression
- 2.4.8 Iteratively update models to improve performance

#### **Deliverables**

Reports and Documents

- Monthly reports on task status
- Technical papers describing algorithm design and performance
- Conference presentations

#### Code Modules and Database Archives

All code modules and documentation will be continually available on EcoHealth Alliance's GitHub page with access provided to DTRA. Databases will be made available for download and provided on-disk to DTRA per request.

OMB Number: 4040-0001 Expiration Date: 12/31/2022

APPLICATION FOR FEDERAL ASSISTANCE	3. DATE RECEIVED BY STATE State Application Identifier
SF 424 (R&R)	
1. TYPE OF SUBMISSION	4. a. Federal Identifier
Pre-application Application Changed/Corrected Application	b. Agency Routing Identifier
2. DATE SUBMITTED Applicant Identifier	
	c. Previous Grants.gov Tracking ID
5. APPLICANT INFORMATION	Organizational DUNS: 0770900660000
Legal Name: Ecollealth Alliance	
Department: Division:	
Street1: 460 West 34th Street	
Street2: Suite 1701	
City: New York County / Paris	<b>µ</b> : ZĀ
State: NY: New York	Province:
Country: USA: UNITED STATES	ZIP / Postal Code: 19991-2317
Person to be contacted on matters involving this application	
Prefix: Dr. First Name: Noam	Middle Name:
Last Name: Ross	Suffix:
Position/Title: Principal Scientist, Computational Research	7
Street1: 460 West 34th Street	
Street2: Suite 1701	
City: New York County / Paris	sh:
State: NY: New York	Province:
Country: USA: UNITED STATES	ZIP / Postal Code: 10001 2317
Phone Number: 2123804460	804465
Email: ress@ecchealthalliance.org	
6. EMPLOYER IDENTIFICATION (EIN) or (TIN): 211728494	
7. TYPE OF APPLICANT: M: Nonprofit with 501C3 IRS	Status (Other than Institution of Higher Education)
Other (Specify):	
Small Business Organization Type  Women Owned  Social	lly and Economically Disadvantaged
8. TYPE OF APPLICATION: If Revision, mark a	ppropriate box(es).
New Resubmission A. Increase At	ward B. Decrease Award C. Increase Duration D. Decrease Duration
Renewal Continuation Revision E. Other (spec	
Is this application being submitted to other agencies? Yes No W	/hat other Agencies?
	OG OF FEDERAL DOMESTIC ASSISTANCE NUMBER: 12,351
Defense Threat Reduction Agency	cientific Research - Combating Weapons of Mass Destruction
11. DESCRIPTIVE TITLE OF APPLICANT'S PROJECT:	
Predicting Biothreat Impacts from Early-Stage Data via '	Transfer Learning
12. PROPOSED PROJECT: 13. CONGRESSIONAL DISTRICT	OF APPLICANT
Start Date Ending Date	
07/01/2020 06/30/2022 NY-010	

14. PROJECT DIRECTOR/PRI	NCIPAL INVESTIGA	TOR CONTACT II	NFORMATION							
Prefix: Dr.	First Name: Noam				Middle Na	me: [				
Last Name: Ross					Suffix:					
Position/Title: Principal s	Scientist, Compu	tational Resea	arch							
Organization Name: Ecollea	ilth Alliance									
Department:		Division:								
Street1: 460 West 34th S	Street									
Street2: Suite 1701										
City: New York		County /	Parish: NY							
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Phone Number: 2123804460	0	Fax Number: 2	123804465							
Email: ross@ecohealthal	liance.org									
15. ESTIMATED PROJECT FUNDING  16. IS APPLICATION SUBJECT TO REVIEW BY STATE EXECUTIVE ORDER 12372 PROCESS?										
a. Total Federal Funds Requesi	ted	494,712.98	a.   Lo		APPLICATI					
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19. Authorized Representativ	re									
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Phone Number: 212380446	0	Fax Number:	123804465							
Email: chmura@ccchealth	alliance.org									
Signature of	Authorized Represe	entative					Date S	igned		
	Aleksei Chmura						05/28	72020		
20. Pre-application				hA	ld Attachme	nt				
21. Cover Letter Attachment				<b>⊣</b>	ld Attachme	=======================================			]	
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Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

This publicly releasable abstract is provided to DTRA for use in fulfillment of Section 8123 of the Defense Appropriations Act and future versions of the same.

#### Predicting Biothreat Impacts from Early-Stage Data via Transfer Learning

#### **Project Summary**

Accurately forecasting the impacts of new disease outbreaks is key to decision-making and planning. However, at the early stage of outbreaks, little data is available to to make predictions. Modern "Big Data" machine learning (ML) techniques generally require large data sets, reducing their usability in these situations. Our proposed work addresses this challenge by using models that can learn how much data from previous events is useful for making predictions about current conditions. We will use this approach to understand what aspects of diseases, their hosts, or environments can be used for prediction and how much they can contribute to making accurate forecasts.

# **RESEARCH & RELATED Other Project Information**

OMB Number: 4040-0001 Expiration Date: 12/31/2022

1. Are Human Subjects Involved? Yes No
1.a. If YES to Human Subjects
Is the Project Exempt from Federal regulations? Yes No
If yes, check appropriate exemption number.
If no, is the IRB review Pending? Yes No
IRB Approval Date:
Human Subject Assurance Number:
2. Are Vertebrate Animals Used? Yes No
2.a. If YES to Vertebrate Animals
Is the IACUC review Pending? Yes No
IACUC Approval Date:
Animal Welfare Assurance Number:
3. Is proprietary/privileged information included in the application?
4.a. Does this Project Have an Actual or Potential Impact - positive or negative - on the environment?
4.b. If yes, please explain:
4.c. If this project has an actual or potential impact on the environment, has an exemption been authorized or an environmental assessment (EA) or environmental impact statement (EIS) been performed?  Yes No
4.d. If yes, please explain:
5. Is the research performance site designated, or eligible to be designated, as a historic place? Yes No
5.a. If yes, please explain:
6. Does this project involve activities outside of the United States or partnerships with international collaborators?  Yes  No
6.a. If yes, identify countries:
6.b. Optional Explanation:
7. Project Summary/Abstract [1245-DTRA-ML_Project-Summary_2020-05-]   Delete Attachment   View Attachment
8. Project Narrative 1246-DTRA-MITechnical Proposal-2020- Delete Attachment View Attachment
9. Bibliography & References Cited Add Attachment
10. Facilities & Other Resources Add Attachment
11. Equipment Add Attachment
12. Other Attachments Add Attachments

OMB Number: 4040-0001 Expiration Date: 12/31/2022

# RESEARCH & RELATED Senior/Key Person Profile (Expanded)

		PROFILE - F	Project Director/Pr	incipal Invest	tigator				
Prefix: Dr.	* First Name	3: Noam			Middle N	ame:			
* Last Name: Ro	:55				S	Suffix:			
Position/Title: Pr	incipal Scientist	, Computation	al Research	Department:					
Organization Nar	m <b>e</b> :EcoHealth Allia	ince				Division:			
* Street1: 460 1	West 34th Street								
Street2: Suite	e 1701								
* City: New Y	York		County/ Parish:	NY					
* State: NY:	New York				Province:				
* Country: USA:	UNITED STATES				* Zip / Post	al Code: 100	001-2317		
* Phone Number:	2123804460	Fa	x Number: 212380	)4465					
* E-Mail: ross@e	ecohealthalliance.	org							
Credential, e.g.	, agency login:								]
* Project Role:	PD/PI		Other Project	Role Categor	y:				
Degree Type:	PhD								
Degree Year:	2015								
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			ROFILE - Senior/K	ey Person 1					
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* Last Name: Yo						Suffix:			
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	m <b>e</b> : EcoHealth Allis	ince				Division:			
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Credential, e.g. * Project Role:	agency login:	a l	Other Project	Role Categor	r <b>y:</b> Senior	Engineer			
_		al .	Other Project	Role Categor	r <b>y:</b> Senior	Engineer			

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RESEARCH & RELATED Senior/Key Person Profile (Expanded)			
PROFILE - Senior/Key Person 2			
Prefix: First Name: Emma	Middle Name:		
* Last Name: Mendelsohn	Suffix:		
Position/Title: Research Scientist. Department:			
Organization Name: Ecoffealth Alliance	Division:		
*Street1: 460 West 34th Street			
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* State: NY: New York	Province:		
* Country: USA: UNITED STATES	* Zip / Postal Code: 10001-2317		
* Phone Number: 2123804460 Fax Number: 2123804465			
*E-Mail: nendelsohn@ecohealthalliance.org			
Credential, e.g., agency login:			
* Project Role: Other Professional Other Project Role Category	y: Research Scientist		
Degree Type: MSc			
Degree Year: 2015			
Attach Biographical Sketch 1239-Mendelsohn_Biosketch.pdf	Delete Attachment View Attachment		
Attach Current & Pending Support 1240-Mendelsohn_049.pdf	Delete Attachment View Attachment		
PROFILE - Seniar/Key Person 3			
Prefix: Dr. * First Name: Peter	Middle Name:		
* Last Name: Daszak	Suffix:		
Position/Title: President Department:			
Organization Name: EcoRealth Alliance	Division:		
* Street1: 460 West 34th Street			
Street2: Site 1701			
* City: New York County/ Parish:			
* State: NY: New York	Province:		
* Country: USA: UNITED STATES	* Zip / Postal Code: 10001 2317		
* Phone Number: 12123804460 Fax Number: 12123804465			

Credential, e.g., agency login: daszak

PED

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Other Professional

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Project Role:

Degree Type:

Degree Year:

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Other Project Role Category: Senior Advisor

1241 Daszak\_BioSketch.pdf

# RESEARCH & RELATED Senior/Key Person Profile (Expanded)

PROFILE - Senior/Key Person 4			
Prefix: Dr. * First Name: Billy	Middle Name:		
* Last Name: Karesh	Suffix:		
Position/Title: Executive VP for Health and Policy Departme	nt:		
Organization Name: Ecoffealth Alliance	Division:		
*Street1: 460 West 34th Street			
Street2: Size 1701			
* City: New York County/ Parish:			
* State: NY: New York	Province:		
*Country: USA: UNITED STATES	* Zip / Postal Code: 10001-2317		
* Phone Number: 12134804460 Fax Number: 12023084465			
* E-Mail: karesh@ecohealthalliance.org			
Credential, e.g., agency login: wkaresh			
* Project Role: Other Professional Other Project Role Category: Senior Advisor			
Degree Type: DVM			
Degree Year: 1982			
Attach Biographical Sketch [1243-Karesh_BioSketch.pdf] Delete Attachment View Attachment			
Attach Current & Pending Support 1244-Karesh_069.pdf	Delete Attachment View Attachment		

#### **Noam Ross**

EcoHealth Alliance, 460 W 34th St, New York, NY 10001

E-mail: ross@ccohealthalliance.org

#### **Professional Preparation**

Brown University

University of California-Davis	Ecology	Ph.D.	2015
Appointments			
Principal Scientist, Computational F	Research, EcoHealth Alliance		2020 - present
Senior Research Scientist Ecologist,	EcoHealth Alliance		2016 - 2019
Disease Ecologist, EcoHealth Allian	nce		2015 - 2016
Graduate Researcher, University of	California-Davis		2010 - 2015
Senior Analyst, Environmental Strat	tegy and Markets, GreenOrder		2007 - 2009
Analyst, Environmental Strategy and	d Markets, GreenOrder		2006 - 2007
Contract Researcher: Energy Efficie	ent Products Initiative, Wal-Mart		2006

**Environmental Science** 

B.S.

2006

#### **Publications**

- Albery, G.F., Eskew, E.A., Ross, N. et al. Predicting the global mammalian viral sharing network using phylogeography. **Nature Communications 11**, 2260 (2020).
- Carlson CJ, Albery GF, Merow C, Trisos CH, Zipfel CM, Eskew EA, Ross Net al. Climate change will drive novel cross-species viral transmission. **bioRxiv**, Cold Spring Harbor Laboratory; (2020)
- Eskew EΛ, White ΛM, Ross N, Smith KM, Smith KF, Rodríguez JP, et al. United States wildlife and wildlife product imports from 2000 2014. Scientific Data 16;7(1).
- Li, H., Mendelsohn, E., Zong, C., Zhang, W., Hagan, E., Wang, N., ... Ross N., ... Daszak, P. (2019). Human-animal interactions and bat coronavirus spillover potential among rural residents in Southern China. **Biosafety and Health**, 1(2), 84–90.
- Ross, N., Eskew, E., & Ray, N. (2019). citesdb: An R package to support analysis of CITES Trade Database shipment-level data. **Journal of Open Source Software**, 4(37), 1483.
- Pedersen, E. J., Miller, D. L., Simpson, G. L., & Ross, N. (2019). Hierarchical generalized additive models in ecology: an introduction with mgcv. **PeerJ**, 7, e6876.
- Carlson CJ, Kracalik I, Ross N, Alexander K, Hugh-Jones ME, Fegan M, Elkin B, Epp T, Shury T, Bagirova M, Getz WM, Blackburn JK. (2019). The global distribution of Bacillus anthracis and associated anthrax risk to humans, livestock, and wildlife. **Nature Microbiology**. 4(8):1337-1343 a
- Rostal MK, Ross N, Machalaba C, Cordel C, Paweska JT, Karesh WB. (2018) Benefits of a One Health approach: An example using Rift Valley fever. **One Health** 5:34-36.
- Olival KJ, Hosseini P, Zambrana-Torrelio C, Ross N, Bogich T, Daszak P. (2017) Host and viral traits predict zoonotic spillover from mammals. **Nature** 546:646–650.
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- Schreiber S, Ross N. (2016) Individual-based integral projection models: The role of size-structure on extinction risk and establishment success. **Methods in Ecology and Evolution** 7(7):867-74.
- Boettiger C, Ross N, Hastings A. (2013) Early warning signals: The charted and uncharted territories. **Theoretical Ecology** 6(3):255-64.

Current and Pending Support (See GPG Section II.D.8 for guidance on information to include on this form.)

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The following information should be provided for each investigator and other senior personnel. Failure to provide this information may delay consideration of this proposal.			
Other agencies (including NSF) to which this proposal has			
Investigator: Noam Ross	Other agencies (including Nor ) to F	mich this proposarnas	
Support: 🛛 Current 🔲 Pending 🔲	Submission Planned in Near Future	*Transfer of Support	
Project/Proposal Title:			
Expanding Software Peer Review			
Expanding Software Feet Review			
Source of Support: Sloan Foundation			
Total Award Amount: \$677,784 Total Aw	vard Period Covered: 09/01/2019 - 09	9/30/2021	
Location of Project: EcoHealth Alliance, subcontract f	from UC Berekeley (prime)		
Person-Months Per Year Committed to the	<b>*</b> (. ,	Cumr	
		Sumr:	
Support:	Submission Planned in Near Future	*Transfer of Support	
Project/Proposal Title:			
Rapid Evaluation of Pathogens to prevent Epidemics	in Livestock		
Source of Support: DHS Science and Technology Di	rectorate		
		04/0004	
, , ,	vard Period Covered: 09/05/2019-09/	04/2021	
Location of Project: EcoHealth Alliance			
Person-Months Per Year Committed to the	Cal: 3 Acad:	Sumr:	
Support:	Submission Planned in Near Future	☐ *Transfer of Support	
Project/Proposal Title:		•	
Software Containers for Science			
Software Goritainers for Objetice			
Source of Support: Chan-Zuckerbug Foundation			
	vard Period Covered: 01/01/2020 – 1	2/31/2020	
·	vald Fellod Coveled. 0 1/01/2020 = 1	2/31/2020	
Location of Project: Berkeley, CA and New York, NY			
Person-Months Per Year Committed to the	Cal: 1 Acad:	Sumr:	
Support:	Submission Planned in Near Future	*Transfer of Support	
Project/Proposal Title:			
Understanding Risk of Zoonotic Virus Emergence in E	FID Hotsnots of Southeast Asia		
Shadratang Nok of Zoonata Vilab Emorgonae in E	218 (10topoto di Codinodot / tota		
Source of Support: NIH			
	word Davied Covered: 2/1/20 2/29/2	E	
· · · · · ·	vard Period Covered: 3/1/20 – 2/28/2	5	
Location of Project: Global			
Person-Months Per Year Committed to the	Cal: 1 Acad:	Sumr:	
Support:	Submission Planned in Near Future		
Project/Proposal Title:			
Reducing the threat of Rift Valley Fever through Ecok	agy Enidomiology and Socia Econor	nice	
Reducing the threat of Kilt Valley Fever through Ecolo	ogy, Epiderniology and Socio-Econor	nics	
Course of Cumports DTDA			
Source of Support: DTRA			
	vard Period Covered: 06/01/2019 - 05	5/31/2024	
Location of Project: USA, South Africa			
Person-Months Per Year Committed to the	Cal: 1 Acad:	Sumr:	
*If this project has previously been funded by another		nation for immediately pre-	
ceding funding period.	<b>3</b> ,,	,	

#### Robert J. Young

520 Eighth Avenue, Ste. 1201, New York, NY 10018

E-mail: young@ecohealthalliance.org

#### **Professional Preparation**

University of Minnesota	Computer Science	BS	1999
University of New Mexico	Computer Science	MS	2007
Appointments			
Senior Research Software Engineer, EcoHealth Alliance, New York, NY			2020 - present
Bioinformatics Programmer, Institute for Genomic Medicine, CUMC			2016 - 2020
Computer Analyst, Laboratory for Computational Vision, HHMI			2007 - 2016
Research Assistant, CTI-II/NESSI Project, University of New Mexico			2005 - 2007
Research Assistant, University of New Mexico			2004 - 2005
Bioinformatics Specialist, Visual Neuroscience Laboratory, HHMI			2000 - 2002

#### **Publications and Presentations**

- Healy, M.J., Olinger, R. D., <u>Young, R.J.</u>, Taylor, S.E., Caudell, T., Larson, K.W., (2009) Applying Category Theory to Improve the Performance of a Neural Architecture, **Neurocomputing**, 72(13-15):3158-3173.
- Young, R.J., Ritthaler, M., Zimmer, P., McGraw, J., Healy, M.J., Caudell, T.P., (2007) Comparison of Adaptive Resonance Theory Neural Networks for Astronomical Region of Interest Detection and Noise Characterization, International Joint Conference on Neural Networks, 2123-2128.
- Young, R.J., Ritthaler, M., Healy, M., Caudell, T.P., Zimmer, P. and McGraw, J. (2006) Adaptive Resonance Theory Neural Networks for Astronomical Region of Interest Detection and Noise Characterization, Astronomical Data Analysis Software and Systems (ADASS) XVI, ASP Conference Series, Vol. XXX, Phoenix, AZ (Poster).
- Ritthaler, M., Luger, G. F., <u>Young, R.</u>, McGraw, J. and Zimmer, P. (2006) Bayesian Belief Networks for Astronomical Object Recognition and Classification in CTI-II, **Astronomical Data Analysis Software and Systems XVI, ASP Conference Series, Vol XXX**, Phoenix, ΔZ (Poster).
- Young, R.J. & Williams, L.R., Psychophysical Demonstration of (An)isotropy in Illusory Surface Computation, **Second Annual CS UNM Student Conference (CSUSC 2006)**, Albuquerque, NM (Poster).
- Healy, M.J., Olinger, R., Young, R.J., Caudell, T.P., and Larson, K., (2005) Modification of the ART-1 Architecture Based on Category Theoretic Design Principles, **Special Session in the INNS/IEEE International Joint Conference on Neural Networks**, 457-462.
- Young, R.J. & Williams, L.R., (2005) Psychophysical Measurement of Illusory Contour Shape, First Annual CS UNM Student Conference (CSUSC 2005), Albuquerque, NM (Poster).

#### Peter Daszak

EcoHealth Alliance, 520 Eighth Avenue, Suite 1201, New York, NY 10018, USA E-mail: daszak@ecohealthalliance.org

#### **Professional Preparation**

Bangor University, UK	Zoology	BS	1986
Univ. of East London, UK	Infectious Diseases	PhD	1993
Appointments			
President & Chief Scientist, EcoHealth Alliance New York			2009- present
Executive Director, Consortium for Conservation Medicine		2001-2009	
Faculty Research Scientist, University of Georgia		1999- 2001	
Guest Researcher, Centers for Disease Control and Prevention (CDC)			1999
Faculty Research Scientist, Kingston University, UK		1993- 1998	
Research Assistant, University of East London, UK		1989- 1992	

#### **Publications**

- Nikolay B, Salje H, Hossain MJ, Khan AKMD, Sazzad HMS, Rahman M, <u>Daszak P</u>, Ströher U, Pulliam JRC, Kilpatrick AM, Nichol ST, Klena JD, Sultana S, Afroj S, Luby SP, Cauchemez S & Gurley ES. (2019). Transmission of Nipah Virus 14 Years of Investigations in Bangladesh. **New Engl J Med** 380:1804-1814
- Carroll D, <u>Daszak P\*</u>, Wolfe ND, Gao GF, Morel C, Morzaria S, Pablos-Méndez A, Tomori O, Mazet JAK (2018). The global virome project. **Science** 359: 872-874
- Zhou P, Fan H, Lan T, Yang X-L, Shi W-F, Zhang W, Zhu Y, Zhang Y-W, Xie Q-M, Mani S, Zheng X-S, Li B, Li J-M, Guo H, Pei G-Q, An X-P, Chen J-W, Zhou L, Mai K-J, Wu Z-X, Li D, Anderson DE, Zhang L-B, Li S-Y, Mi Z-Q, He T-T, Cong F, Fuo P-J, Huang R, Luo Y, Liu X-L, Chen J, Huang Y, Sun Q, Zhang X-L-L, Wang Y-Y, Xing S-Z, Chen Y-S, Sun Y, Li J, <u>Daszak P\*</u>, Wang L-F\*, Shi Z-L\*, Tong Y-G\*, Ma J-Y\* (2018). Fatal Swine Acute Diarrhea Syndrome caused by an HKU2-related Coronavirus of Bat Origin. **Nature** 556: 255-258
- Olival KJ\*, Hosseini PR, Zambrana-Torrelio C, Ross N, Bogich TL, <u>Daszak P\*</u> (2017). Host and viral traits predict zoonotic spillover from mammals. **Nature** 546, 646–650
- Allen T, Murray KA, Zambrana-Torrelio C, Morse SS, Rondinini C, Di Marco M, Breit N, Olival KJ, <u>Daszak P\*</u> (2017). Global hotspots and correlates of emerging zoonotic diseases. **Nature Comm** 8: 1124
- Ge X-Y, Li J-L, Yang X-L, Chmura ΛΛ, Zhu G, Epstein JH, Mazet JK, Hu B, Zhang W, Peng C, Zhang Y-J, Luo C-M, Tan B, Wang N, Zhu Y, Crameri G, Zhang S-Y, Wang L-F, <u>Daszak P\*</u>, Shi Z-L\* (2013). Isolation and characterization of a bat SARS-like Coronavirus that uses the ΛCE2 receptor. **Nature** 503: 535-538
- Daszak P\*, Zambrana-Torellio C, Bogich TL, Fernandez M, Epstein JH, Murray KΛ, Hamilton H (2013). Interdisciplinary approaches to understanding disease emergence: The past, present and future drivers of Nipah virus emergence. **PNAS** 110: 3681-3688
- Li W, Shi Z, Yu M, Ren W, Smith C, Epstein JH, Wang H, Crameri G, Hu Z, Zhang H, Zhang J, McEachern J, Field H, <u>Daszak P</u>, Eaton BT, Zhang S, Wang L-F. (2005). Bats are natural reservoirs of SARS-like coronaviruses. **Science** 310: 676-679
- <u>Daszak P\*</u>, Cunningham AA, Hyatt AD (2000). Emerging infectious diseases of wildlife threats to biodiversity and human health. **Science** 287: 443-449

#### Emma Mendelsohn

EcoHealth Alliance, 520 Eighth Avenue, Ste. 1201, New York City, NY 10001, USA E-mail: mendelsohn@ecohealthalliance.org

#### **Professional Preparation**

Wesleyan University, CT	Earth and	BA	2010
	Environmental Science		
Duke University, NC	Ecotoxicology and	MEM	2015
	Environmental Health		

#### **Appointments**

Research Scientist, EcoHealth Alliance	2018 - present
Open-source software peer reviewer, rOpenSci	2019 - present
Project Scientist, Integral Consulting	2015 - 2018
Statistics Teaching Assistant, Duke University	2014
Environmental Health Science Intern, Cardo ChemRisk	2014

#### **Publications**

- Li, H., E. Mendelsohn, C. Zong, E. Hagan, W. Zhang, N. Wang, S. Li, H. Yan, H. Huang, G. Zhu, N. Ross, Λ. Chmura, P. Terry, M. Fielder, M. Miller, Z. Shi, P. Daszak. (2019). Human-animal interaction and bat coronavirus spillover potential among rural residents in Southern China. Biosafety and Health. 1(2):84-90.
- Iwai, H., Hoberman A.M., Goodrum, P.E., Mendelsohn, E., and Anderson, J.K. (2019). Addendum to Iwai and Hoberman (2014) Reassessment of Developmental Toxicity of PFHxA in Mice. **International Journal of Toxicology**. 38(3):183-191.
- Mendelsohn, E., A. Hagopian, K. Hoffman, C.M. Butt, A. Lorenzo, J. Congleton, T.F. Webster, and H.M. Stapleton. (2016). Nail polish as a source of exposure to triphenyl phosphate. **Environment International**. 86:45–51.
- Hoffman, K., C.M. Butt, T.F. Webster, E.V. Preston, S.C. Hammel, C. Makey, A. Lorenzo, E.M. Cooper, C. Carignan, J.D. Meeker, R. Hauser, A. Soubry, S.K. Murphy, T.M. Price, C. Hoyo, E. Mendelsohn, J. Congleton, J.L. Daniels, and H.M. Stapleton. (2017). Temporal trends in exposure to organophosphate flame retardants in the United States. Environmental Science and Technology Letters. 4(3):112-118
- Kopelovich, L., A.L. Perez, N. Jacobs, <u>E. Mendelsohn</u>, and J.J. Keenan. (2015). Screening level human health risk assessment of toluene and dibutyl phthalate in nail lacquers. **Food and Chemical Toxicology**. 81:46–53.

#### William B. Karesh

EcoHealth Alliance, 460 W. 34th St, New York City, NY 10001, USA E-mail: karesh@ccohealthalliance.org

Pro	fession	nal Prei	paration

Clemson University	Biology	BS	1977
Univ. of Georgia	Veterinary Medicine	DVM	1982
Zool. Society of San Diego	Residency Wildlife Health		1982 - 1984
Appointments			
Life Member, Council on Foreign	Relations		2016 -
present			
Emerging Pandemic Threats Partne	er Liaison, USAID EPT PRE	DICT-2	2014 - present
Advisor, WHO Expert Panel on M	ERS-CoV		2013 - present
Expert, WHO International Health	Regulation Roster of Expert	S	2013 - present
Executive Vice President for Healt	2010 - present		
President, Working Group on Wild	llife Diseases, OIE, France		2008 - present
Co-Chair, Wildlife Health Speciali	st Group, IUCN, Switzerland	<del>j</del>	2001 - present
Chief Technical Officer, USAID E	PT PREDICT		2009 - 2014
Chief of Party, USAID Global Avi	an Influenza Network for Su	rveillance	2006 - 2009
Vice President & Director, Global	Health Programs, Wildlife C	ons. Society	2001 - 2010

- Kandeil Λ, Gomaa M, Shehata M, El Taweel ΛN, Mahmoud SH, Bagato O, Moatasim Y, Kutkat O, Kayed ΛS, *Dawson P*, Oui X, Bahl J, Webby RJ, <u>Karesh WB</u>, *Kayali G*. (2019) Isolation and characterization of a distinct influenza A virus from Egyptian bats. **Journal of Virology** 93(2).
- Anthony SJ, Johnson CK, Grieg D, Kramer S, Che X, Wells HL, Hicks AL, Joly D, Wolfe ND, Daszak P, <u>Karesh WB</u>, Lipkin WI, Morse SS, Predict Consortium, Mazet J, Goldstein T. (2019) Global patterns in coronavirus diversity. **Virus Evolution** 3(1).
- *Machalaba CC*, Elwood SE, Forcella S, <u>Karesh WB</u>. Global avian influenza surveillance in wild birds: Λ strategy to capture viral diversity. (2015) **Emerging Infectious Diseases** 21(4):e1-7.
- Alagaili AN, Briese T, Mishra N, Kapoor V, Sameroff SC, de Wit E, Munster VJ, Hensley LE, Zalmout IS, Kapoor A, Epstein JH, <u>Karesh WB</u>, Daszak P, Mohammed OB, Lipkin WI. (2014) Middle East Respiratory Syndrome Coronavirus infection in dromedary camels in Saudi Arabia. **mBio** 5(2):e00884-14.
- <u>Karesh WB</u>, Dobson Λ, Lloyd-Smith JO, Lubroth J, Dixon MΛ, Bennett M, Aldrich S, Harrington T, Formenty P, Loh EH, *Machalaba CC*, Thomas MJ, Heymann DL. (2012) Ecology of zoonoses: Natural and unnatural histories. **The Lancet** 380(9857):1936-45.
- Gaidet N, Caron A, Cappelle J, Cumming GS, Balanca G, Hammoumi S, Cattoli G, Abolnik C, De Almeida RS, Gil P, Fereidouni SR, Grosbois V, Tran A, Mundava J, Fofana B, El Mamy ABO, Ndlovu M, Mondain-Monval JY, Triplet P, Hagemeijer W, <u>Karesh WB</u>, Newman SH, Dodman T. (2012) Understanding the ecological drivers of avian influenza virus infection in wildfowl: A continental-scale study across Africa. **Proceedings of the Royal Society B: Biological Sciences** 279(1731):1131-41.
- Morse SS, Mazet JAK, Woolhouse M, Parrish CR, Carroll D, <u>Karesh WB</u>, Zambrana-Torrelio C, Lipkin WI, Daszak P. (2012) Prediction and prevention of the next pandemic zoonosis. The Lancet 380:1956-65.

(See GPG Section II.D.8 for guidance on information to include on this form.)

Other agencies (including NSF) to which this proposal has: none Investigator: Emma Mendelsohn  Support: Current Pending Submission Planned in Near Future Transfer of Support Project/Proposal Title:  Investigating Impact of Nutritional Deficiencies on Pandemics
Project/Proposal Title:
Investigating Impact of Nutritional Deficiencies on Pandemics
Source of Support: Eleanor Crooks Foundation
Total Award Amount: \$70,000 Total Award Period Covered: 05/01/2020 – 10/31/2020
Location of Project: Global
Person-Months Per Year Committed to the Cal: 2 Acad: Sumr:
Support:
Project/Proposal Title:
REPEL: Rapid Evaluation of Pathogens to prevent Epidemics in Livestock
Source of Support: DHS
Total Award Amount: \$566,274.26 Total Award Period Covered: 09/05/2019-09/04/2021
Location of Project: Global
Person-Months Per Year Committed to the Cal: 6 Acad: Sumr:
Support:   Current Pending Submission Planned in Near Future *Transfer of Support *Transfer o
Project/Proposal Title:
Source of Support:
Total Award Amount: Total Award Period Covered:
Location of Project:
Person-Months Per Year Committed to the Cal: Acad: Sumr:
Support:   Current Pending Submission Planned in Near Future *Transfer of Support *Transfer o
Project/Proposal Title:
Source of Support:
Total Award Amount: Total Award Period Covered:
Location of Project:
ceding funding period.
Person-Months Per Year Committed to the Cal: Acad: Sumr:  *If this project has previously been funded by another agency, please list and furnish information for immediately pre-

NSF Form 1239 (10/99)

USE ADDITIONAL SHEETS AS NECESSARY



(See GPG Section II.D.8 for guidance on information to include on this form.)

The following information should be provided information may delay consideration of this p	roposal.		•	·
Investigator: Robert Young		Other agencies (inclu	uding NSF) to wh	ich this proposal has: none
Support:		Submission Planned	in Near Future	*Transfer of Support
Project/Proposal Title:				
REPEL: Rapid Evaluation of Pathogens to pr	revent Epi	demics in Livestock		
Source of Support: DTRA				
Total Award Amount:  Location of Project: Global	Total Awa	ard Period Covered:	04/09/2019-03/09	9/2021
Person-Months Per Year Committed to the		Cal: 6	Acad:	Sumr:
Support: Current Pending	По	Submission Planned		*Transfer of Support
Project/Proposal Title:		ubmission Flamed	iii Neai Future	Transier of Support
Source of Support: DHS				
Total Award Amount:	Total Awa	ard Period Covered:		
Location of Project:				
Person-Months Per Year Committed to the		Cal:	Acad:	Sumr:
Support: Current Pending	Пѕ	Submission Planned	in Near Future	*Transfer of Support
Project/Proposal Title:	_			
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Person-Months Per Year Committed to the		Cal:	Acad:	Sumr:
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NSF Form 1239 (10/99)

USE ADDITIONAL SHEETS AS NECESSARY



The following information should be provided for eac information may delay consideration of this proposal		senior personne	el. Failure to provide this
The proposition of the propositi	Other agencies (includi	ing NSF) to whi	ch this proposal has
Investigator: William Karesh		- '	, ,
Support: ☐ Current ☐ Pending ☐ Project/Proposal Title:	Submission Planned in	Near Future	☐ *Transfer of Support
Reducing the Threat of Middle East Respiratory Syn- Strengthening Regional Disease Surveillance Capac		RS) and Avian I	nfluenza (AI) in Jordan &
Source of Support: DTRA			
Total Award Amount: \$4,975,681.29 Total A	ward Period Covered: 01	/01/ <mark>2020 - 1</mark> 2/3	31/2024
Location of Project: USA, Jordan			
Person-Months Per Year Committed to the	Cal: 2	Acad:	Sumr:
Support:	Submission Planned in	Near Future	
Project/Proposal Title:	5	<b>A</b> - ! -	
Understanding the Risk of Bat-Bourne Zoonotic Dise	ase Emergence in vveste	ern Asia	
Source of Support: DTRA			
• •	ward Period Covered: 10	/02/201 <b>7</b> - 10/0	11/2022
Location of Project: Global	walu Fellou Coveleu. To	1/02/2017 - 10/0	11/2022
Person-Months Per Year Committed to the	Cal: 1.56	Acad:	Curr
Support: Current Pending	Submission Planned in		Sumr:  *Transfer of Support
Project/Proposal Title:	Cubinission Filanneu III	ivear i didic	Transier of Support
Ground Truth Network			
Gradia Trail Network			
Source of Support: DHS			
• •	ward Period Covered: 09	/30/2016 - 09/2	9/2021
Location of Project: USA			
Person-Months Per Year Committed to the	Cal: 1.56	Acad:	Sumr:
Support:	Submission Planned in		*Transfer of Support
Project/Proposal Title:			
READY: Augmenting Capacity for Humanitarian Eme	ergencies of Infectious Di	iseases with Ep	oidemic or Pandemic
Potential		•	
Source of Support: USAID			
Total Award Amount: \$143,605 Total A	ward Period Covered: 09	/25/2018 - 09/3	30/2021
Location of Project: Global			
Person-Months Per Year Committed to the	Cal: 1.56	Acad:	Sumr:
Support:	Submission Planned in	Near Future	
Project/Proposal Title:			
Emerging Pandemic Threats PREDICT-2			
Source of Support: USAID			
• • •	ward Period Covered: 10	//01/2014 - 09/3	30/2019
Location of Project: Global	_		_
Person-Months Per Year Committed to the	Cal: 4.56	Acad:	Sumr:
*If this project has previously been funded by anothe ceding funding period.	r agency, please list and	turnish informa	ation for immediately pre-

	The following information should be provided for each investigator and other senior personnel. Failure to provide this information may delay consideration of this proposal.				
Other agencies (including NSF) to which this proposal has					
Investigator: William Karesh					
Support: ☐ Current ☐ Pending ☐ Submission Planned in Near Future ☐ *Transfer of Supp	ort				
Project/Proposal Title:					
Crimean-Congo hemorrhagic fever: Reducing an emerging health threat in Tanzania					
Source of Support: DTRA					
Total Award Amount: \$4,848,483					
Location of Project: USA, Tanzania					
Person-Months Per Year Committed to the Cal: 1.0 Acad: Sumr:					
Support:	ort				
Project/Proposal Title:	JI I				
Reducing the threat of Rift Valley Fever through Ecology, Epidemiology and Socio-Economics					
Reducing the threat of Kitt valley rever through Ecology, Epiderhiology and Socio-Economics					
Source of Support: DTRA					
Total Award Amount: \$4,988,526 Total Award Period Covered: 06/01/2019 - 05/31/2024					
Location of Project: USA, South Africa					
•					
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sumr:  Support: □ Current □ Pending □ Submission Planned in Near Future □ *Transfer of Supp	ort				
Support: □ Current □ □ Fending □ Submission Flatined in Near Future □ Transfer of Supp   Project/Proposal Title:	JII				
Understanding filovirus diversity in bat reservoirs and implications for human outbreaks in West Africa					
Source of Support: NIH					
Total Award Amount: \$7,307,869 Total Award Period Covered: 03/01/2020 - 02/28/2025					
Τοταί / Walta / Tilba συνσίου, συνσίου σε					
Location of Project: Global					
Location of Project: Global  Person Menths Per Year Committed to the					
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Person-Months Per Year Committed to the       Cal: 2.0       Acad:       Sumr:         Support:       ☐ Current       ☐ Pending       ☐ Submission Planned in Near Future       ☐ *Transfer of Support:	ort				
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sumr:  Support: Current Pending Submission Planned in Near Future *Transfer of Support/Proposal Title:  Reducing the Threat from high-risk pathogens causing febrile illness in Liberia	ort				
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sumr:  Support: Current Pending Submission Planned in Near Future *Transfer of Support: Project/Proposal Title:  Reducing the Threat from high-risk pathogens causing febrile illness in Liberia  Source of Support: DTRA	ort				
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sumr:  Support: Current Pending Submission Planned in Near Future *Transfer of Support: Project/Proposal Title:  Reducing the Threat from high-risk pathogens causing febrile illness in Liberia  Source of Support: DTRA  Total Award Amount: \$4,912,818.06 Total Award Period Covered: 01/01/2020 - 12/31/2024	ort				
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sumr:  Support: Current Pending Submission Planned in Near Future *Transfer of Support: Project/Proposal Title:  Reducing the Threat from high-risk pathogens causing febrile illness in Liberia  Source of Support: DTRA  Total Award Amount: \$4,912,818.06 Total Award Period Covered: 01/01/2020 - 12/31/2024  Location of Project: USA, Liberia	ort				
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sumr:  Support: Current Pending Submission Planned in Near Future *Transfer of Support: Project/Proposal Title:  Reducing the Threat from high-risk pathogens causing febrile illness in Liberia  Source of Support: DTRA  Total Award Amount: \$4,912,818.06 Total Award Period Covered: 01/01/2020 - 12/31/2024  Location of Project: USA, Liberia  Person-Months Per Year Committed to the Cal: 1.5 Acad: Sumr:					
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sumr:  Support: Current Pending Submission Planned in Near Future *Transfer of Support: Project/Proposal Title:  Reducing the Threat from high-risk pathogens causing febrile illness in Liberia  Source of Support: DTRA  Total Award Amount: \$4,912,818.06 Total Award Period Covered: 01/01/2020 - 12/31/2024  Location of Project: USA, Liberia  Person-Months Per Year Committed to the Cal: 1.5 Acad: Sumr:  Support: Current Pending Submission Planned in Near Future *Transfer of Support: Transfer of Supp					
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Person-Months Per Year Committed to the Support: Current Pending Submission Planned in Near Future *Transfer of Support: Project/Proposal Title:  Reducing the Threat from high-risk pathogens causing febrile illness in Liberia  Source of Support: DTRA  Total Award Amount: \$4,912,818.06 Total Award Period Covered: 01/01/2020 - 12/31/2024  Location of Project: USA, Liberia  Person-Months Per Year Committed to the Cal: 1.5 Acad: Sumr:  Support: Current Pending Submission Planned in Near Future *Transfer of Support: Total Award Amount: Total Award Period Covered:  Source of Support: Total Award Amount: Total Award Period Covered:  Location of Project:  Person-Months Per Year Committed to the Cal: Acad: Sumr:	ort				



(See GPG Section II.D.8 for guidance on information to include on this form.)

The following information should be provided for each investigator and other senior personnel. Fa information may delay consideration of this proposal.	ailure to provide this
Other agencies (including NSF) to which this	s proposal has
Investigator: Peter Daszak	
	Transfer of Support
Project/Proposal Title:	
Reducing the Threat of Middle East Respiratory Syndrome Coronavirus (MERS) and Avian Influen Strengthening Regional Disease Surveillance Capacity (current proposal)	nza (AI) in Jordan &
Source of Support: DTRA	
Total Award Amount: \$4,975,681.29 Total Award Period Covered: 01/01/2020 - 12/31/202	24
Location of Project: USA, Jordan	
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sumr	г:
Support:	Transfer of Support
Project/Proposal Title:	
CNH2: Land-use change and microbial spillover as a coupled human-Resilience	
Source of Support: NSF	
Total Award Amount: \$1,599,738 Total Award Period Covered:	
Location of Project: EcoHealth Alliance and Malaysia	
Person-Months Per Year Committed to the Cal: 0.50 Acad: Summ	
Support:   Current Pending Submission Planned in Near Future *T Project/Proposal Title:	Transfer of Support
Understanding Risk of Zoonotic Virus Emergence in EID Hotspots of South East Asia	
Orderstanding Kisk of Zoonotic Virus Emergence in Elb Hotspots of South East Asia	
Source of Support: NIH	
Total Award Amount: \$7,573,721.35 Total Award Period Covered: 3/1/2020 – 2/28/2025	
Location of Project: EcoHealth Alliance and International Field and Lab Locations	
Person-Months Per Year Committed to the Cal: 3.0 Acad: Sumr	r:
Support: ☐ Current ☐ Pending ☐ Submission Planned in Near Future ☐ *T	Transfer of Support
Project/Proposal Title:	
One Health Workforce – Next Generation	
Source of Support: USAID	
Total Award Amount: \$5,000,000 Total Award Period Covered: 10/1/2019 – 9/30/2020	
Location of Project: EcoHealth Alliance and International Field and Lab Locations	
Person-Months Per Year Committed to the Cal: 1.44 Acad: Sumr	
''	Transfer of Support
Project/Proposal Title:	
Strategies to Prevent (STOP) Spillover	
Source of Support: USAID	
Total Award Amount: \$100,000,000 Total Award Period Covered: 8/31/2020 – 8/30/2025	
Location of Project: EcoHealth Alliance and International Field and Lab Locations	
Person-Months Per Year Committed to the Cal: 3.0 Acad: Sumr	r:
*If this project has previously been funded by another agency, please list and furnish information for	or immediately pre-
ceding funding period.	



# Predicting Biothreat Impacts from Early-Stage Data via Transfer Learning Dr. Noam Ross, EcoHealth Alliance, HDTRA1-14-24c-FRCWMD-BAA

**Objective:** Test new ML approaches - transfer learning by partial pooling, and hybrid mechanistic-ML models, for predicting impacts of novel biothreats at early, data-sparse stage of outbreak.

#### Method:

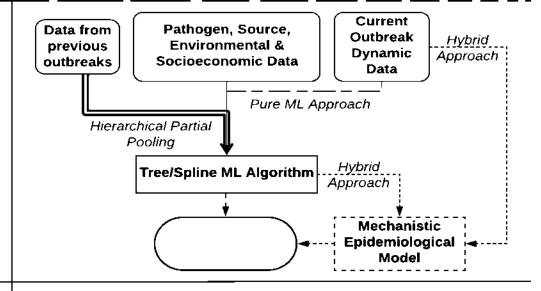
- Quantify gain in prediction by training models to learn information across diseases (partial pooling)
- Assess variables most predictive of epidemic impacts at early stages outbreak.
- Quantify gain in power from using model to predict parameters of mechanistic compartmental model.
- Determine point in epidemic growth where hybrid models become more accurate than pure ML models.

**Benefits of Technology:** Disease forecasts in early stages of outbreaks, especially novel outbreaks, require strong assumptions in highly structured models. Hierarchical partial pooling learns what factors from previous outbreaks may be transferred to apply to new predictions, improving early-stage assessments of severity and risk.

### Challenges:

- Missing data in early-stage outbreak conditions → Learn most important variables to prioritize data collection
- ML models non-interpretable and poor at extrapolation →
  Use hybrid approach with ML parameterizing structured
  mechanistic model capable of simulating scenarios

Readiness: TRL1



### Milestones:

- Y1Q2: Engineer project database & test infrastructure
- Y1Q4: Build, fit, and compare ML model structures to
- OY1Q2: Extend models to incorporate genetic data
- OY1Q4: Compare pure ML models to ML-mechanistic hybrids

## **Funding Profile:**

\$246,211 Year 1, \$248,402 Option Year

### **Contact information:**

Noam Ross, Senior Research Scientist, ross@ecohealthalliance.org, 1.212.380.4471



OMB Number: 4040-0001 Expiration Date: 12/31/2022

APPLICATION FOR FEDERAL ASSISTANCE	3. DATE RECEIVED BY STATE State Application Identifier
SF 424 (R&R)	
1. TYPE OF SUBMISSION	4. a. Federal Identifier
Pre-application Application Changed/Corrected Application	b. Agency Routing Identifier
2. DATE SUBMITTED Applicant Identifier	
	c. Previous Grants.gov Tracking ID
5. APPLICANT INFORMATION	Organizational DUNS: 0770900660000
Legal Name: Ecollealth Alliance	
Department: Division:	
Street1: 460 West 34th Street	
Street2: Suite 1701	
City: New York County / Paris	<b>µ</b> : ZĀ
State: NY: New York	Province:
Country: USA: UNITED STATES	ZIP / Postal Code: 19991-2317
Person to be contacted on matters involving this application	
Prefix: Dr. First Name: Noam	Middle Name:
Last Name: Ross	Suffix:
Position/Title: Principal Scientist, Computational Research	7
Street1: 460 West 34th Street	
Street2: Suite 1701	
City: New York County / Paris	sh:
State: NY: New York	Province:
Country: USA: UNITED STATES	ZIP / Postal Code: 10001 2317
Phone Number: 2123804460	804465
Email: ress@ecchealthalliance.org	
6. EMPLOYER IDENTIFICATION (EIN) or (TIN): 211728494	
7. TYPE OF APPLICANT: M: Nonprofit with 501C3 IRS	Status (Other than Institution of Higher Education)
Other (Specify):	
Small Business Organization Type  Women Owned  Social	lly and Economically Disadvantaged
8. TYPE OF APPLICATION: If Revision, mark a	ppropriate box(es).
New Resubmission A. Increase At	ward B. Decrease Award C. Increase Duration D. Decrease Duration
Renewal Continuation Revision E. Other (spec	
Is this application being submitted to other agencies? Yes No W	/hat other Agencies?
	OG OF FEDERAL DOMESTIC ASSISTANCE NUMBER: 12,351
Defense Threat Reduction Agency	cientific Research - Combating Weapons of Mass Destruction
11. DESCRIPTIVE TITLE OF APPLICANT'S PROJECT:	
Predicting Biothreat Impacts from Early-Stage Data via '	Transfer Learning
12. PROPOSED PROJECT: 13. CONGRESSIONAL DISTRICT	OF APPLICANT
Start Date Ending Date	
07/01/2020 06/30/2022 NY-010	

14. PROJECT DIRECTOR/PRI	NCIPAL INVESTIGA	TOR CONTACT II	NFORMATION						
Prefix: Dr.	First Name: Noam				Middle Na	me: [			
Last Name: Ross					Suffix:				
Position/Title: Principal s	Scientist, Compu	tational Resea	arch						
Organization Name: Ecollea	ilth Alliance								
Department:		Division:							
Street1: 460 West 34th S	Street								
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City: New York		County /	Parish: NY						
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Country:	USA: UNITE	STATES		ZIP	/ Postal Cod	de: 🔯	0001-23	1.7	
Phone Number: 2123804460	0	Fax Number: 2	123804465						
Email: ross@ecohealthal	liance.org								
15. ESTIMATED PROJECT FU	JNDING		16. IS APPLICA 12372 PROCES		UBJECT TO	REV	IEW BY	STATE E	XECUTIVE ORDER
a. Total Federal Funds Requesi	ted	494,712.98	a.   Lo		APPLICATI				
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19. Authorized Representativ	re								
Prefix: Dr.	First Name: Alekse	<u> </u>			Middle I	Name:			
Last Name: Ohmura					Suffix:		<b>L</b>		
	Organizational R	epresentative			'				
Organization: Ecoficatth A					$\neg$				
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Street1: 460 West 34	th Street								
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Phone Number: 212380446	0	Fax Number:	123804465						
Email: chmura@ocohoalthallianco.org									
Signature of	Authorized Represe	entative					Date S	igned	
	Aleksei Chmura						05/28	72020	
20. Pre-application				hA	ld Attachme	nt			
21. Cover Letter Attachment				$\dashv \vdash \vdash$	ld Attachme	=======================================			]
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Thrust Area 7, J5-A: Machine Learning for Infectious Biological Agent Forecasting

This publicly releasable abstract is provided to DTRA for use in fulfillment of Section 8123 of the Defense Appropriations Act and future versions of the same.

### Predicting Biothreat Impacts from Early-Stage Data via Transfer Learning

### **Project Summary**

Accurately forecasting the impacts of new disease outbreaks is key to decision-making and planning. However, at the early stage of outbreaks, little data is available to to make predictions. Modern "Big Data" machine learning (ML) techniques generally require large data sets, reducing their usability in these situations. Our proposed work addresses this challenge by using models that can learn how much data from previous events is useful for making predictions about current conditions. We will use this approach to understand what aspects of diseases, their hosts, or environments can be used for prediction and how much they can contribute to making accurate forecasts.

# **RESEARCH & RELATED Other Project Information**

OMB Number: 4040-0001 Expiration Date: 12/31/2022

1. Are Human Subjects Involved? Yes No
1.a. If YES to Human Subjects
Is the Project Exempt from Federal regulations? Yes No
If yes, check appropriate exemption number.
If no, is the IRB review Pending? Yes No
IRB Approval Date:
Human Subject Assurance Number:
2. Are Vertebrate Animals Used? Yes No
2.a. If YES to Vertebrate Animals
Is the IACUC review Pending? Yes No
IACUC Approval Date:
Animal Welfare Assurance Number:
3. Is proprietary/privileged information included in the application?
4.a. Does this Project Have an Actual or Potential Impact - positive or negative - on the environment?
4.b. If yes, please explain:
4.c. If this project has an actual or potential impact on the environment, has an exemption been authorized or an environmental assessment (EA) or environmental impact statement (EIS) been performed?  Yes No
4.d. If yes, please explain:
5. Is the research performance site designated, or eligible to be designated, as a historic place? Yes No
5.a. If yes, please explain:
6. Does this project involve activities outside of the United States or partnerships with international collaborators? Yes No
6.a. If yes, identify countries:
6.b. Optional Explanation:
7. Project Summary/Abstract [1245-DTRA-ML_Project-Summary_2020-05-]   Delete Attachment   View Attachment
8. Project Narrative 1246-DTRA-MITechnical Proposal-2020- Delete Attachment View Attachment
9. Bibliography & References Cited Add Attachment
10. Facilities & Other Resources Add Attachment
11. Equipment Add Attachment
12. Other Attachments Add Attachments

OMB Number: 4040-0001 Expiration Date: 12/31/2022

# RESEARCH & RELATED Senior/Key Person Profile (Expanded)

		PROFILE - Project Director/	Principal Inves	stigator			
Prefix: Dr.	* First Name	_	•	Middle N	ame:		
* Last Name: Ro		<u> </u>		s	uffix:		
Position/Title: Pu	rincipal Scientist	, Computational Research	Department	<u> </u>			
Organization Na	me: EcoHealth Allia	ande			Division:		
* Street1: 460	West 34th Street						
Street2: Suit	e 1701						
* City: New	York	County/ Parish	u Ny				
* State: NY:	New York			Province:			
* Country: USA	: UNITED STATES			] * Zip / Post	al Code: 10001-2317		
* Phone Number	12123804460	Fax Number: 21238	304465				
* E-Mail: rossa	ecohealthalliance.	org					
Credential, e.g.	., agency login:						
* Project Role:	PC/PI	Other Project	t Role Catego	ory:			
Degree Type:	PhD			i			
Degree Year:	2015						
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		PROFILE - Senior/	Key Person 1				
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* Last Name: Yes Position/Title: Re Organization Na * Street1: 460 Street2: Suit * City: New * State: NY: * Country: USA * Phone Number * E-Mail: Young	oung esearch Software E me: EcoRealth Allie West 34th Street e 1701 York New York : UNITED STATES	ngineer ande  County/ Parish  Fax Number: 2123	Department	S::	Uffix: Division:		
* Last Name: Yes Position/Title: Re Organization Na * Street1: 460 Street2: Suit * City: New * State: NY: * Country: USA * Phone Number * E-Mail: Young	cung esearch Software E me: EcoRealtH Allis West 34th Street e 1701 York New York : UNITED STATES :: 2123804460 @ecohealthalliance	ngineer arroe  County/ Parish Fax Number: 2123	Department	Province:	uffix: Division: al Code: 10001-2317		
* Last Name: Yes Position/Title: Res Organization Na * Street1: 460 Street2: Suit * City: New * State: NY: * Country: USA * Phone Number * E-Mail: Young Oredential, e.g	pung esearch Software E me: EcoRealth Allie West 34th Street e 1701 York New York : UNITED STATES : 2123804460 @ecohealthalliance ., agency login:	ngineer arroe  County/ Parish Fax Number: 2123	Department	Province:	uffix: Division: al Code: 10001-2317		
* Last Name: Yes Position/Title: Res Organization Na * Street1: 460 Street2: Suit * City: New * State: NY: * Country: USA * Phone Number * E-Mail: Young Credential, e.g * Project Role:	pung esearch Software E me: DecKealth Allis West 34th Street e 1701 York New York : UNITED STATES : 2123804460 @ecohealthalliance ., agency login: Other Professiona	ngineer arroe  County/ Parish Fax Number: 2123	Department	Province:	uffix: Division: al Code: 10001-2317		

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RESEARCH & RELATED Senior/Key Person Profile (Expanded)						
PROFILE - Senior/Key Person 2						
Prefix: First Name: Emma	Middle Name:					
* Last Name: Mendelsohn	Suffix:					
Position/Title: Research Scientist Department:						
Organization Name: Ecoffealth Alliance	Division:					
*Street1: 460 West 34th Street						
Street2: Suite 1701						
* City: New York County/ Parish:						
*State: NY: New York	Province:					
* Country: USA: UNITED STATES	* Zip / Postal Code: 10001-2317					
* Phone Number: 2123804460 Fax Number: 2123804465						
*E-Mail: nendelsohn@ecohealthalliance.org						
Credential, e.g., agency login:						
* Project Role: Other Professional Other Project Role Category	y: Research Scientist					
Degree Type: MSC						
Degree Year: 2015						
Attach Biographical Sketch 1239-Mendelsohn_Biosketch.pdf	Delete Attachment View Attachment					
Attach Current & Pending Support 1240-Mendelsohn_049.pdf	Delete Attachment View Attachment					
PROFILE - Senior/Key Person 3						
Prefix: Dr. * First Name: Peter	Middle Name:					
* Last Name: Daszak	Suffix:					
Position/Title: President Department:						
Organization Name: EcoRealth Alliance	Division:					
* Street1: 460 West 34th Street						
Street2: Ste 1701						
* City: New York County/ Parish:						
* State: NY: New York	Province:					
* Country: USA: UNITED STATES	* Zip / Postal Code: 10001 2317					
* Phone Number: 12123804460 Fax Number: 12123804465						

Credential, e.g., agency login: daszak

PED

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Other Professional

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Project Role:

Degree Type:

Degree Year:

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Other Project Role Category: Senior Advisor

1241 Daszak\_BioSketch.pdf

# RESEARCH & RELATED Senior/Key Person Profile (Expanded)

PROFILE - Senior/Key Person 4			
Prefix: Dr. * First Name: Billy	Middle Name:		
* Last Name: Karesh	Suffix:		
Position/Title: Executive VP for Health and Policy Departme	nt:		
Organization Name: Ecoffealth Alliance	Division:		
*Street1: 460 West 34th Street			
Street2: Size 1701			
* City: New York County/ Parish:			
* State: NY: New York	Province:		
* Country: USA: UNITED STATES	* Zip / Postal Code: 10001-2317		
* Phone Number: 12134804460 Fax Number: 12023084465			
* E-Mail: karesh@ecohealthalliance.org			
Credential, e.g., agency login: wkaresin			
* Project Role: Other Professional Other Project Role Cate	gory: Senior Advisor		
Degree Type: DVM			
Degree Year: 1982			
Attach Biographical Sketch [1243-Karesh_BioSketch.pdf] Delete Attachment View Attachment			
Attach Current & Pending Support 1244-Karesh_069.pdf	Delete Attachment View Attachment		

#### **Noam Ross**

EcoHealth Alliance, 460 W 34th St, New York, NY 10001

E-mail: ross@ccohealthalliance.org

#### **Professional Preparation**

Brown University

University of California-Davis	Ecology	Ph.D.	2015
Appointments			
Principal Scientist, Computational F	Research, EcoHealth Alliance		2020 - present
Senior Research Scientist Ecologist,	EcoHealth Alliance		2016 - 2019
Disease Ecologist, EcoHealth Allian	nce		2015 - 2016
Graduate Researcher, University of	California-Davis		2010 - 2015
Senior Analyst, Environmental Strat	tegy and Markets, GreenOrder		2007 - 2009
Analyst, Environmental Strategy and	d Markets, GreenOrder		2006 - 2007
Contract Researcher: Energy Efficie	ent Products Initiative, Wal-Mart		2006

**Environmental Science** 

B.S.

2006

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- Boettiger C, Ross N, Hastings A. (2013) Early warning signals: The charted and uncharted territories. **Theoretical Ecology** 6(3):255-64.

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The following information should be provided for each investigator and other senior personnel. Failure to provide this information may delay consideration of this proposal.			
	Other agencies (including NSF) to w	high this proposal has	
Investigator: Noam Ross	Other agencies (including Nor ) to w	men mis proposarnas	
Support: 🛛 Current 🔲 Pending 🔲	Submission Planned in Near Future	*Transfer of Support	
Project/Proposal Title:			
Expanding Software Peer Review			
Expanding Soltware Feel Review			
Source of Support: Sloan Foundation			
Total Award Amount: \$677,784 Total Av	ward Period Covered: 09/01/2019 - 09	/30/2021	
Location of Project: EcoHealth Alliance, subcontract	from UC Berekeley (prime)		
Person-Months Per Year Committed to the	* " ,	Cumr	
		Sumr:	
Support:	Submission Planned in Near Future	*Transfer of Support	
Project/Proposal Title:			
Rapid Evaluation of Pathogens to prevent Epidemics	in Livestock		
Source of Support: DHS Science and Technology D	irectorate		
		14/0004	
	ward Period Covered: 09/05/2019-09/0	J4/2021	
Location of Project: EcoHealth Alliance			
Person-Months Per Year Committed to the	Cal: 3 Acad:	Sumr:	
Support:	Submission Planned in Near Future	*Transfer of Support	
Project/Proposal Title:			
Software Containers for Science			
Software Containers to Objetice			
Source of Support: Chan-Zuckerbug Foundation			
Total Award Amount: \$75,000 Total Award Period Covered: 01/01/2020 – 12/31/2020			
·		131/2020	
Location of Project: Berkeley, CA and New York, NY			
Person-Months Per Year Committed to the	Cal: 1 Acad:	Sumr:	
Support:	Submission Planned in Near Future	*Transfer of Support	
Project/Proposal Title:			
Understanding Risk of Zoonotic Virus Emergence in	FID Hotsnots of Southeast Asia		
Shadrotanding Note of Zoonotto Viras Emergence in	Ele Hotopote of Couthouse Hota		
Source of Support: NIH			
· · · ·	ward Period Covered: 3/1/20 – 2/28/25	)	
Location of Project: Global			
Person-Months Per Year Committed to the	Cal: 1 Acad:	Sumr:	
Support:	Submission Planned in Near Future	☐ *Transfer of Support	
Project/Proposal Title:			
	logy Enidomiology and Socia Foonem	sion.	
Reducing the threat of Rift Valley Fever through Ecol	logy, ⊑pidemiology and Socio-Econom	iics	
0 (0 (0.70)			
Source of Support: DTRA			
Total Award Amount: \$4,988,526 Total Av	ward Period Covered: 06/01/2019 - 05	/31/2024	
Location of Project: USA, South Africa			
Person-Months Per Year Committed to the	Cal: 1 Acad:	Sumr:	
*If this project has previously been funded by anothe			
ceding funding period.	. againey, produce not arial farmor fillotti	sir isi iiiinodidaay pro-	

#### Robert J. Young

520 Eighth Avenue, Ste. 1201, New York, NY 10018

E-mail: young@ecohealthalliance.org

#### **Professional Preparation**

University of Minnesota	Computer Science	BS	1999
University of New Mexico	Computer Science	MS	2007
Appointments			
Senior Research Software Engine	2020 - present		
Bioinformatics Programmer, Insti	2016 - 2020		
Computer Analyst, Laboratory for	2007 - 2016		
Research Assistant, CTI-II/NESS	2005 - 2007		
Research Assistant, University of	2004 - 2005		
Bioinformatics Specialist, Visual Neuroscience Laboratory, HHMI			2000 - 2002

#### **Publications and Presentations**

- Healy, M.J., Olinger, R. D., <u>Young, R.J.</u>, Taylor, S.E., Caudell, T., Larson, K.W., (2009) Applying Category Theory to Improve the Performance of a Neural Architecture, **Neurocomputing**, 72(13-15):3158-3173.
- Young, R.J., Ritthaler, M., Zimmer, P., McGraw, J., Healy, M.J., Caudell, T.P., (2007) Comparison of Adaptive Resonance Theory Neural Networks for Astronomical Region of Interest Detection and Noise Characterization, International Joint Conference on Neural Networks, 2123-2128.
- Young, R.J., Ritthaler, M., Healy, M., Caudell, T.P., Zimmer, P. and McGraw, J. (2006) Adaptive Resonance Theory Neural Networks for Astronomical Region of Interest Detection and Noise Characterization, Astronomical Data Analysis Software and Systems (ADASS) XVI, ASP Conference Series, Vol. XXX, Phoenix, AZ (Poster).
- Ritthaler, M., Luger, G. F., <u>Young, R.</u>, McGraw, J. and Zimmer, P. (2006) Bayesian Belief Networks for Astronomical Object Recognition and Classification in CTI-II, **Astronomical Data Analysis Software and Systems XVI, ASP Conference Series, Vol XXX**, Phoenix, ΔZ (Poster).
- Young, R.J. & Williams, L.R., Psychophysical Demonstration of (An)isotropy in Illusory Surface Computation, **Second Annual CS UNM Student Conference (CSUSC 2006)**, Albuquerque, NM (Poster).
- Healy, M.J., Olinger, R., <u>Young, R.J.</u>, Caudell, T.P., and Larson, K., (2005) Modification of the ART-1 Architecture Based on Category Theoretic Design Principles, **Special Session in the INNS/IEEE International Joint Conference on Neural Networks**, 457-462.
- Young, R.J. & Williams, L.R., (2005) Psychophysical Measurement of Illusory Contour Shape, First Annual CS UNM Student Conference (CSUSC 2005), Albuquerque, NM (Poster).

#### Peter Daszak

EcoHealth Alliance, 520 Eighth Avenue, Suite 1201, New York, NY 10018, USA E-mail: daszak@ecohealthalliance.org

#### **Professional Preparation**

Bangor University, UK	Zoology	BS	1986
Univ. of East London, UK	Infectious Diseases	PhD	1993
Appointments			
President & Chief Scientist, Ec	oHealth Alliance New York		2009- present
Executive Director, Consortium for Conservation Medicine			2001-2009
Faculty Research Scientist, University of Georgia			1999- 2001
Guest Researcher, Centers for l	1999		
Faculty Research Scientist, Kir	1993- 1998		
Research Assistant, University	of East London, UK		1989- 1992

- Nikolay B, Salje H, Hossain MJ, Khan AKMD, Sazzad HMS, Rahman M, <u>Daszak P</u>, Ströher U, Pulliam JRC, Kilpatrick AM, Nichol ST, Klena JD, Sultana S, Afroj S, Luby SP, Cauchemez S & Gurley ES. (2019). Transmission of Nipah Virus 14 Years of Investigations in Bangladesh. **New Engl J Med** 380:1804-1814
- Carroll D, <u>Daszak P\*</u>, Wolfe ND, Gao GF, Morel C, Morzaria S, Pablos-Méndez A, Tomori O, Mazet JAK (2018). The global virome project. **Science** 359: 872-874
- Zhou P, Fan H, Lan T, Yang X-L, Shi W-F, Zhang W, Zhu Y, Zhang Y-W, Xie Q-M, Mani S, Zheng X-S, Li B, Li J-M, Guo H, Pei G-Q, An X-P, Chen J-W, Zhou L, Mai K-J, Wu Z-X, Li D, Anderson DE, Zhang L-B, Li S-Y, Mi Z-Q, He T-T, Cong F, Fuo P-J, Huang R, Luo Y, Liu X-L, Chen J, Huang Y, Sun Q, Zhang X-L-L, Wang Y-Y, Xing S-Z, Chen Y-S, Sun Y, Li J, <u>Daszak P\*</u>, Wang L-F\*, Shi Z-L\*, Tong Y-G\*, Ma J-Y\* (2018). Fatal Swine Acute Diarrhea Syndrome caused by an HKU2-related Coronavirus of Bat Origin. **Nature** 556: 255-258
- Olival KJ\*, Hosseini PR, Zambrana-Torrelio C, Ross N, Bogich TL, <u>Daszak P\*</u> (2017). Host and viral traits predict zoonotic spillover from mammals. **Nature** 546, 646–650
- Allen T, Murray KA, Zambrana-Torrelio C, Morse SS, Rondinini C, Di Marco M, Breit N, Olival KJ, <u>Daszak P\*</u> (2017). Global hotspots and correlates of emerging zoonotic diseases. **Nature Comm** 8: 1124
- Ge X-Y, Li J-L, Yang X-L, Chmura ΛΛ, Zhu G, Epstein JH, Mazet JK, Hu B, Zhang W, Peng C, Zhang Y-J, Luo C-M, Tan B, Wang N, Zhu Y, Crameri G, Zhang S-Y, Wang L-F, <u>Daszak P\*</u>, Shi Z-L\* (2013). Isolation and characterization of a bat SARS-like Coronavirus that uses the ΛCE2 receptor. **Nature** 503: 535-538
- Daszak P\*, Zambrana-Torellio C, Bogich TL, Fernandez M, Epstein JH, Murray KΛ, Hamilton H (2013). Interdisciplinary approaches to understanding disease emergence: The past, present and future drivers of Nipah virus emergence. **PNAS** 110: 3681-3688
- Li W, Shi Z, Yu M, Ren W, Smith C, Epstein JH, Wang H, Crameri G, Hu Z, Zhang H, Zhang J, McEachern J, Field H, <u>Daszak P</u>, Eaton BT, Zhang S, Wang L-F. (2005). Bats are natural reservoirs of SARS-like coronaviruses. **Science** 310: 676-679
- <u>Daszak P\*</u>, Cunningham AA, Hyatt AD (2000). Emerging infectious diseases of wildlife threats to biodiversity and human health. **Science** 287: 443-449

#### Emma Mendelsohn

EcoHealth Alliance, 520 Eighth Avenue, Ste. 1201, New York City, NY 10001, USA E-mail: mendelsohn@ecohealthalliance.org

#### **Professional Preparation**

Wesleyan University, CT	Earth and	BA	2010
	Environmental Science		
Duke University, NC	Ecotoxicology and	MEM	2015
	Environmental Health		

### **Appointments**

Research Scientist, EcoHealth Alliance	2018 - present
Open-source software peer reviewer, rOpenSci	2019 - present
Project Scientist, Integral Consulting	2015 - 2018
Statistics Teaching Assistant, Duke University	2014
Environmental Health Science Intern, Cardo ChemRisk	2014

- Li, H., E. Mendelsohn, C. Zong, E. Hagan, W. Zhang, N. Wang, S. Li, H. Yan, H. Huang, G. Zhu, N. Ross, Λ. Chmura, P. Terry, M. Fielder, M. Miller, Z. Shi, P. Daszak. (2019). Human-animal interaction and bat coronavirus spillover potential among rural residents in Southern China. Biosafety and Health. 1(2):84-90.
- Iwai, H., Hoberman A.M., Goodrum, P.E., Mendelsohn, E., and Anderson, J.K. (2019). Addendum to Iwai and Hoberman (2014) Reassessment of Developmental Toxicity of PFHxA in Mice. **International Journal of Toxicology**. 38(3):183-191.
- Mendelsohn, E., A. Hagopian, K. Hoffman, C.M. Butt, A. Lorenzo, J. Congleton, T.F. Webster, and H.M. Stapleton. (2016). Nail polish as a source of exposure to triphenyl phosphate. **Environment International**. 86:45–51.
- Hoffman, K., C.M. Butt, T.F. Webster, E.V. Preston, S.C. Hammel, C. Makey, A. Lorenzo, E.M. Cooper, C. Carignan, J.D. Meeker, R. Hauser, A. Soubry, S.K. Murphy, T.M. Price, C. Hoyo, E. Mendelsohn, J. Congleton, J.L. Daniels, and H.M. Stapleton. (2017). Temporal trends in exposure to organophosphate flame retardants in the United States. Environmental Science and Technology Letters. 4(3):112-118
- Kopelovich, L., A.L. Perez, N. Jacobs, <u>E. Mendelsohn</u>, and J.J. Keenan. (2015). Screening level human health risk assessment of toluene and dibutyl phthalate in nail lacquers. **Food and Chemical Toxicology**. 81:46–53.

#### William B. Karesh

EcoHealth Alliance, 460 W. 34th St, New York City, NY 10001, USA E-mail: karesh@ccohealthalliance.org

Pro	fession	nal Prei	paration

Clemson University	Biology	BS	1977
Univ. of Georgia	Veterinary Medicine	DVM	1982
Zool. Society of San Diego	Residency Wildlife Health		1982 - 1984
Appointments			
Life Member, Council on Foreign	Relations		2016 -
present			
Emerging Pandemic Threats Partne	2014 - present		
Advisor, WHO Expert Panel on M	2013 - present		
Expert, WHO International Health	2013 - present		
Executive Vice President for Healt	2010 - present		
President, Working Group on Wild	2008 - present		
Co-Chair, Wildlife Health Speciali	2001 - present		
Chief Technical Officer, USAID E	2009 - 2014		
Chief of Party, USAID Global Avi	2006 - 2009		
Vice President & Director, Global	Health Programs, Wildlife C	ons. Society	2001 - 2010

- Kandeil Λ, Gomaa M, Shehata M, El Taweel ΛN, Mahmoud SH, Bagato O, Moatasim Y, Kutkat O, Kayed ΛS, *Dawson P*, Oui X, Bahl J, Webby RJ, <u>Karesh WB</u>, *Kayali G*. (2019) Isolation and characterization of a distinct influenza A virus from Egyptian bats. **Journal of Virology** 93(2).
- Anthony SJ, Johnson CK, Grieg D, Kramer S, Che X, Wells HL, Hicks AL, Joly D, Wolfe ND, Daszak P, <u>Karesh WB</u>, Lipkin WI, Morse SS, Predict Consortium, Mazet J, Goldstein T. (2019) Global patterns in coronavirus diversity. **Virus Evolution** 3(1).
- *Machalaba CC*, Elwood SE, Forcella S, <u>Karesh WB</u>. Global avian influenza surveillance in wild birds: Λ strategy to capture viral diversity. (2015) **Emerging Infectious Diseases** 21(4):e1-7.
- Alagaili AN, Briese T, Mishra N, Kapoor V, Sameroff SC, de Wit E, Munster VJ, Hensley LE, Zalmout IS, Kapoor A, Epstein JH, <u>Karesh WB</u>, Daszak P, Mohammed OB, Lipkin WI. (2014) Middle East Respiratory Syndrome Coronavirus infection in dromedary camels in Saudi Arabia. **mBio** 5(2):e00884-14.
- <u>Karesh WB</u>, Dobson Λ, Lloyd-Smith JO, Lubroth J, Dixon MΛ, Bennett M, Aldrich S, Harrington T, Formenty P, Loh EH, *Machalaba CC*, Thomas MJ, Heymann DL. (2012) Ecology of zoonoses: Natural and unnatural histories. **The Lancet** 380(9857):1936-45.
- Gaidet N, Caron A, Cappelle J, Cumming GS, Balanca G, Hammoumi S, Cattoli G, Abolnik C, De Almeida RS, Gil P, Fereidouni SR, Grosbois V, Tran A, Mundava J, Fofana B, El Mamy ABO, Ndlovu M, Mondain-Monval JY, Triplet P, Hagemeijer W, <u>Karesh WB</u>, Newman SH, Dodman T. (2012) Understanding the ecological drivers of avian influenza virus infection in wildfowl: A continental-scale study across Africa. **Proceedings of the Royal Society B: Biological Sciences** 279(1731):1131-41.
- Morse SS, Mazet JAK, Woolhouse M, Parrish CR, Carroll D, <u>Karesh WB</u>, Zambrana-Torrelio C, Lipkin WI, Daszak P. (2012) Prediction and prevention of the next pandemic zoonosis. The Lancet 380:1956-65.

(See GPG Section II.D.8 for guidance on information to include on this form.)

Other agencies (including NSF) to which this proposal has: none Investigator: Emma Mendelsohn  Support: Current Pending Submission Planned in Near Future Transfer of Support Project/Proposal Title:  Investigating Impact of Nutritional Deficiencies on Pandemics
Project/Proposal Title:
Investigating Impact of Nutritional Deficiencies on Pandemics
Source of Support: Eleanor Crooks Foundation
Total Award Amount: \$70,000 Total Award Period Covered: 05/01/2020 – 10/31/2020
Location of Project: Global
Person-Months Per Year Committed to the Cal: 2 Acad: Sumr:
Support:
Project/Proposal Title:
REPEL: Rapid Evaluation of Pathogens to prevent Epidemics in Livestock
Source of Support: DHS
Total Award Amount: \$566,274.26 Total Award Period Covered: 09/05/2019-09/04/2021
Location of Project: Global
Person-Months Per Year Committed to the Cal: 6 Acad: Sumr:
Support:   Current Pending Submission Planned in Near Future *Transfer of Support *Transfer o
Project/Proposal Title:
Source of Support:
Total Award Amount: Total Award Period Covered:
Location of Project:
Person-Months Per Year Committed to the Cal: Acad: Sumr:
Support:   Current Pending Submission Planned in Near Future *Transfer of Support *Transfer o
Project/Proposal Title:
Source of Support:
Total Award Amount: Total Award Period Covered:
Location of Project:
ceding funding period.
Person-Months Per Year Committed to the Cal: Acad: Sumr:  *If this project has previously been funded by another agency, please list and furnish information for immediately pre-

NSF Form 1239 (10/99)

USE ADDITIONAL SHEETS AS NECESSARY



(See GPG Section II.D.8 for guidance on information to include on this form.)

The following information should be provided for each investigator and other senior personnel. Failure to provide this information may delay consideration of this proposal.				
Investigator: Robert Young		Other agencies (inclu	uding NSF) to wh	ich this proposal has: none
Support:		Submission Planned	in Near Future	*Transfer of Support
Project/Proposal Title:				
REPEL: Rapid Evaluation of Pathogens to pr	revent Epi	demics in Livestock		
Source of Support: DTRA				
Total Award Amount:  Location of Project: Global	Total Awa	ard Period Covered:	04/09/2019-03/09	9/2021
Person-Months Per Year Committed to the		Cal: 6	Acad:	Sumr:
Support: Current Pending	По	Submission Planned		
Project/Proposal Title:		ubmission Flamed	iii Neai Future	Transier of Support
Source of Support: DHS				
Total Award Amount:	Total Awa	ard Period Covered:		
Location of Project:				
Person-Months Per Year Committed to the		Cal:	Acad:	Sumr:
Support: Current Pending	Пѕ	Submission Planned	in Near Future	*Transfer of Support
Project/Proposal Title:	_			
Source of Support:				
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Person-Months Per Year Committed to the		Cal:	Acad:	Sumr:
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Source of Support:				
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NSF Form 1239 (10/99)

USE ADDITIONAL SHEETS AS NECESSARY



The following information should be provided for eac information may delay consideration of this proposal		senior personne	el. Failure to provide this		
The proposition of the propositi	Other agencies (includi	ing NSF) to whi	ch this proposal has		
Investigator: William Karesh		- '	, ,		
Support: ☐ Current ☐ Pending ☐ Project/Proposal Title:	Submission Planned in	Near Future	☐ *Transfer of Support		
Reducing the Threat of Middle East Respiratory Syn- Strengthening Regional Disease Surveillance Capac		RS) and Avian I	nfluenza (AI) in Jordan &		
Source of Support: DTRA					
Total Award Amount: \$4,975,681.29 Total A	ward Period Covered: 01	/01/ <mark>2020 - 1</mark> 2/3	31/2024		
Location of Project: USA, Jordan					
Person-Months Per Year Committed to the	Cal: 2	Acad:	Sumr:		
Support:	Submission Planned in	Near Future			
Project/Proposal Title:	5	<b>A</b> - ! -			
Understanding the Risk of Bat-Bourne Zoonotic Dise	ase Emergence in vveste	ern Asia			
Source of Support: DTRA					
• •	ward Period Covered: 10	/02/201 <b>7</b> - 10/0	11/2022		
Location of Project: Global	walu Fellou Coveleu. To	1/02/2017 - 10/0	11/2022		
Person-Months Per Year Committed to the	Cal: 1.56	Acad:	Curr		
Support: Current Pending	Submission Planned in		Sumr:  *Transfer of Support		
Project/Proposal Title:	Cubinission Filanneum	ivear i didic	Transier of Support		
Ground Truth Network					
Gradia Trail Network					
Source of Support: DHS					
• •	ward Period Covered: 09	/30/2016 - 09/2	9/2021		
Location of Project: USA					
Person-Months Per Year Committed to the	Cal: 1.56	Acad:	Sumr:		
Support:	Submission Planned in		*Transfer of Support		
Project/Proposal Title:					
READY: Augmenting Capacity for Humanitarian Emergencies of Infectious Diseases with Epidemic or Pandemic					
Potential		•			
Source of Support: USAID					
Total Award Amount: \$143,605 Total A	ward Period Covered: 09	/25/2018 - 09/3	30/2021		
Location of Project: Global					
Person-Months Per Year Committed to the	Cal: 1.56	Acad:	Sumr:		
Support:	Submission Planned in	Near Future			
Project/Proposal Title:					
Emerging Pandemic Threats PREDICT-2					
Source of Support: USAID					
• • •	ward Period Covered: 10	//01/2014 - 09/3	30/2019		
Location of Project: Global	_		_		
Person-Months Per Year Committed to the	Cal: 4.56	Acad:	Sumr:		
*If this project has previously been funded by anothe ceding funding period.	r agency, please list and	turnish informa	ation for immediately pre-		

The following information should be provided for each investigator and other senior personnel. Failure to provide this information may delay consideration of this proposal.		
Other agencies (including NSF) to which this proposal has		
Investigator: William Karesh		
Support:   Current   Pending  Submission Planned in Near Future  *Transfer of Support	rt	
Project/Proposal Title:		
Crimean-Congo hemorrhagic fever: Reducing an emerging health threat in Tanzania		
Source of Support: DTRA		
Total Award Amount: \$4,848,483		
Location of Project: USA, Tanzania		
Person-Months Per Year Committed to the Cal: 1.0 Acad: Sumr:		
Support:	-t	
Project/Proposal Title:	•	
Reducing the threat of Rift Valley Fever through Ecology, Epidemiology and Socio-Economics		
Source of Support: DTRA		
Total Award Amount: \$4,988,526 Total Award Period Covered: 06/01/2019 - 05/31/2024		
Location of Project: USA, South Africa		
·		
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sumr:  Support: □ Current □ Pending □ Submission Planned in Near Future □ *Transfer of Support	rt .	
Project/Proposal Title:	١	
Understanding filovirus diversity in bat reservoirs and implications for human outbreaks in West Africa		
Source of Support: NIH		
Total Award Amount: \$7,307,869 Total Award Period Covered: 03/01/2020 - 02/28/2025		
Location of Project: Global		
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sumr:		
Support:	-t	
Project/Proposal Title:	•	
Reducing the Threat from high-risk pathogens causing febrile illness in Liberia		
Treducing the Threat from high-risk pathogens causing lebrile timess in Liberta		
Source of Support: DTRA		
Total Award Amount: \$4,912,818.06 Total Award Period Covered: 01/01/2020 - 12/31/2024		
Location of Project: USA, Liberia		
Person-Months Per Year Committed to the Cal: 1.5 Acad: Sumr:		
Support:	τ	
Project/Proposal Title:		
Source of Support:		
Source of Support:  Total Award Amount:  Total Award Period Covered:		
Total Award Amount: Total Award Period Covered:		
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Total Award Amount: Total Award Period Covered:  Location of Project:  Person-Months Per Year Committed to the Cal: Acad: Sumr:		



(See GPG Section II.D.8 for guidance on information to include on this form.)

The following information should be provided for each investigator and other senior personnel. Failure to provide this information may delay consideration of this proposal.		
Other agencies (including NSF) to which this proposal has		
Investigator: Peter Daszak		
	Transfer of Support	
Project/Proposal Title:		
Reducing the Threat of Middle East Respiratory Syndrome Coronavirus (MERS) and Avian Influenza (AI) in Jordan & Strengthening Regional Disease Surveillance Capacity (current proposal)		
Source of Support: DTRA		
Total Award Amount: \$4,975,681.29 Total Award Period Covered: 01/01/2020 - 12/31/2024		
Location of Project: USA, Jordan		
Person-Months Per Year Committed to the Cal: 2.0 Acad: Sum	nr:	
Support: ☐ Current ☐ Pending ☐ Submission Planned in Near Future ☐ *	Transfer of Support	
Project/Proposal Title:		
CNH2: Land-use change and microbial spillover as a coupled human-Resilience		
Source of Support: NSF		
Total Award Amount: \$1,599,738 Total Award Period Covered:		
Location of Project: EcoHealth Alliance and Malaysia		
Person-Months Per Year Committed to the Cal: 0.50 Acad: Sum	• •	
Support: ☐ Current ☐ Pending ☐ Submission Planned in Near Future ☐ ** Project/Proposal Title:	Transfer of Support	
Understanding Risk of Zoonotic Virus Emergence in EID Hotspots of South East Asia		
Officerstanding Kisk of Zoonotic Virus Emergence in Etb Hotspots of South East Asia		
Source of Support: NIH		
Total Award Amount: \$7,573,721.35 Total Award Period Covered: 3/1/2020 – 2/28/2025		
Location of Project: EcoHealth Alliance and International Field and Lab Locations		
Person-Months Per Year Committed to the Cal: 3.0 Acad: Sum	nr:	
Support: ☐ Current ☐ Pending ☐ Submission Planned in Near Future ☐ **	Transfer of Support	
Project/Proposal Title:		
One Health Workforce – Next Generation		
Source of Support: USAID		
Total Award Amount: \$5,000,000 Total Award Period Covered: 10/1/2019 – 9/30/2020		
Location of Project: EcoHealth Alliance and International Field and Lab Locations		
Person-Months Per Year Committed to the Cal: 1.44 Acad: Sum		
	Transfer of Support	
Project/Proposal Title:		
Strategies to Prevent (STOP) Spillover		
Source of Support: USAID		
Total Award Amount: \$100,000,000 Total Award Period Covered: 8/31/2020 – 8/30/2025		
Location of Project: EcoHealth Alliance and International Field and Lab Locations		
Person-Months Per Year Committed to the Cal: 3.0 Acad: Sum	nr:	
*If this project has previously been funded by another agency, please list and furnish information f	for immediately pre-	
ceding funding period.		