



SERC Talks "Oceans and Human Health: What We Have Learned About COVID From Cholera" December 7, 2022 | 1:00 PM ET

Rita R. Colwell, Ph.D., D.Sc., Distinguished University Professor, University of Maryland at College Park and Johns Hopkins Bloomberg School of Public Health; President, CosmosID, Inc.

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"Oceans and Human Health: What We Have Learned About COVID From Cholera"





Rita R. Colwell, Ph.D., D.Sc. Distinguished University Professor, University of Maryland at College Park and Johns Hopkins Bloomberg School of Public Health; President, CosmosID, Inc.

Moderator: Dr. William Rouse

SERC Research Council Member; Senior Fellow, Office of the Senior Vice President for Research, McCourt School of Public Policy, Georgetown University



"...whoever wishes to pursue the science of medicine must first investigate the seasons of the year and what occurs in them."

Hippocrates, 4th Century B.C.

Oceans and Human Health: What we have learned about COVID from Cholera



December 7, 2022

Rita R. Colwell, Ph.D., D.Sc. Distinguished University Professor University of Maryland College Park

Pandemics of the modern world

Impacted Region		-	7														-	♥,	\bigcirc	
Broad Pathogen		Bacterium	Virus	Bacterium) Bacteriun	Bacteriur	n E Bacteriur	Bacteriun n	n Bacteriun	Virus n	Bacteriur	Virus n	Virus	Bacteriun	Virus n	Virus	Virus	Virus	Virus	
Years lasted	Unknown	22	7	7	8	26	12	15	15	6	24	3	1	Ongoing	1	2	1	3	Ongoing	
Mortality	Decimated civilizations	75-200M	56M	Unknown	300K	1.5M	17-50M	150K	600К	1.0M	1.0M	25-50M	2M	Ongoing	5 1M	 774 	600K	 11K 	Ongoing	
Possible Origin	Worldwide	Central/ East Asia	Europe	e India	India	India	China	India	India	Uzbekista	an India	Debated	Singapo	re H Indonesi	ong Kor a	og China	Mexico	Africa	China	
Pandemic	 Plague Cholera s Influenza 	Bubonic Plague	Smallpo	First Cholera x	Second Cholera	Third Cholera	Plague	Fourth Cholera	Fifth Cholera	Russian Flu*	Sixth Cholera	Spanish Flu*	Asian Flu	Seventh Cholera		SARS#	Swine Flu	Ebola		
r	Pre-mobilized world	1347	1521	1817	1829	1837	1855	1863	1881	1889	1899	1918	1957	1961	1968	2002	2009	2013	2019	?
Data collected and summarized from Sherman., I. 2007 Twelve Disease that Changed Our World, American Society for Microbiology, USA Zimmerman, B.E. and Zimmerman., D.J 2003 Killer Germs, McGraw Hill, USA *Source of virus debated, hence used prevalent name of disease, #SARS-CoV-1, ^S SARS-CoV-2									Cholera					COVID-19						

Water-related diseases

	Cases per year	Deaths per year
Amoebiasis	48,000,000	110,000
Arsenic	28-35m exposed to drinking water with elevated levels	
Diarrhoeal disease, Including cholera	1.5 billion	1,800,000
Dracunuliasis (guinea worm)	> 5000	-
Fluorosis	26 million (China)	-
Giardiasis	500,000	Low
Hepatitis A	1,500,00	-
Intestinal helminths	133,000,000	9400
Malaria	396,000,000	1,300,000
Schistosomiasis	160,000,000	> 10,000
Trachoma	500,000,000	-
Typhoid	500,000	25,000

Cholera: A Global Disease

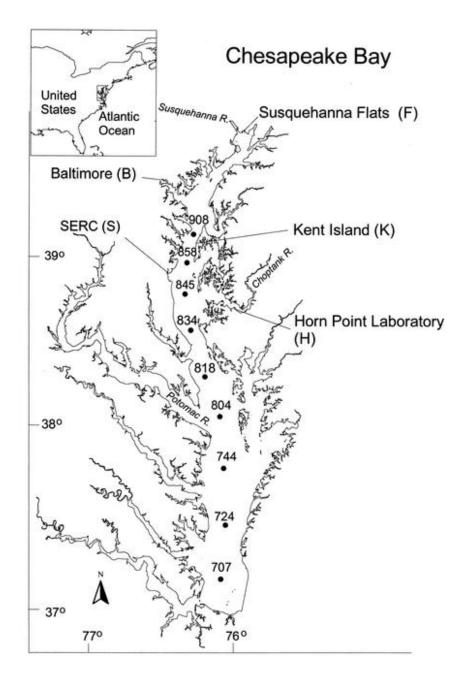
- Acute water-related diarrheal disease
- Seventh pandemic started in 1960s
- Occurs in more than 50 countries affecting approximately 7 million people
- Bengal Delta is known as "native homeland" of cholera outbreaks
- Since cholera bacteria
 - exist naturally in aquatic habitats
 - evidence of new biotypes emerging, it is highly unlikely that cholera will be eradicated but clearly can be controlled by provision of safe drinking water.













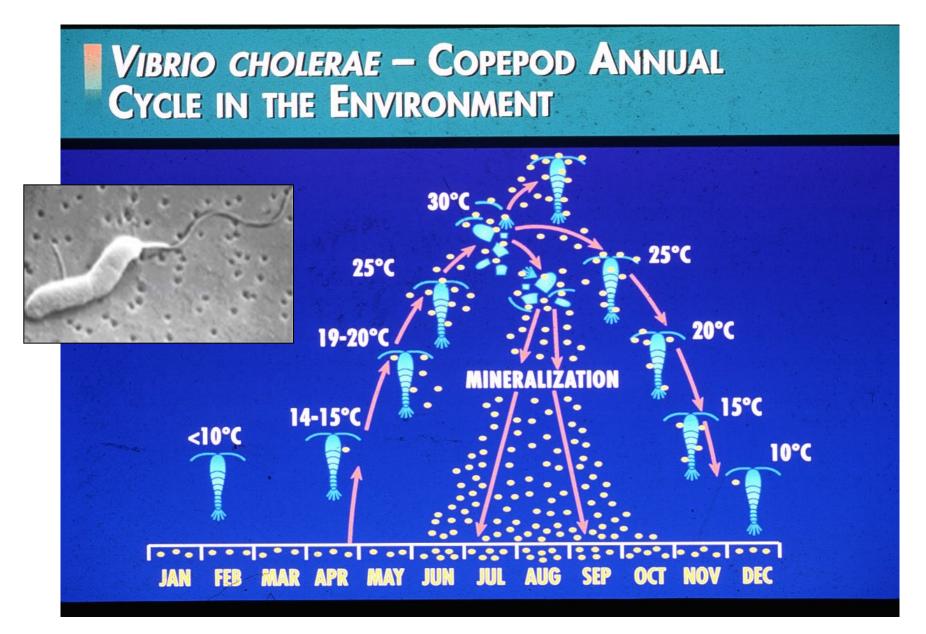


MARYLAN

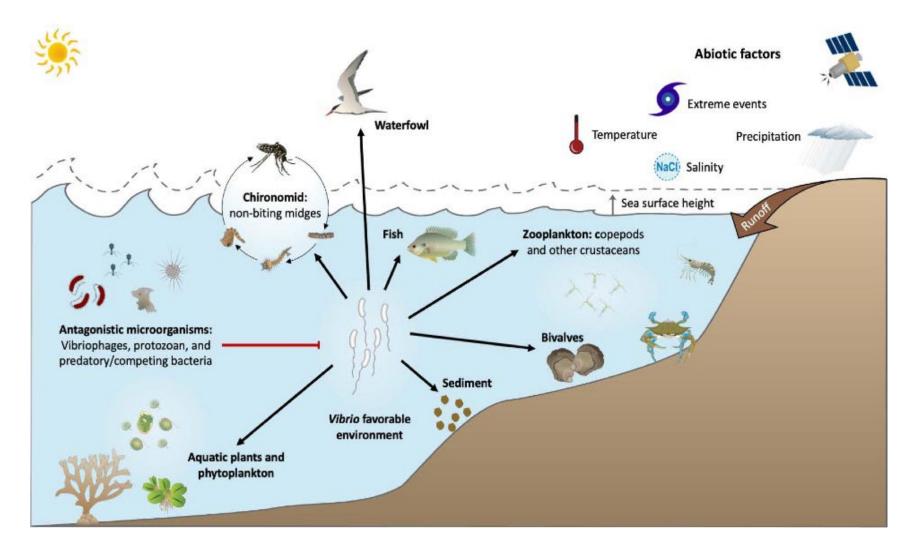




Map of Chesapeake Bay – beginning of the cholera chronicle **1965-1975 An early contribution of marine microbiology to human health: Determination of the** *Vibrio cholerae* **life cycle**



Vibrio and their natural environment



Model for the Transmission of Vibrio Cholerae from the Environment to Humans

PHYSICAL & CHEMICAL CHARACTERISTICS OF WATER

- temperature
- sunlight
- rainfal

- pH
- dissolved oxygen
- salinity & nutrients

FECAL SHEDDING returns V. cholerae to the water. **BIOLOGICAL CHARACTERISTICS**

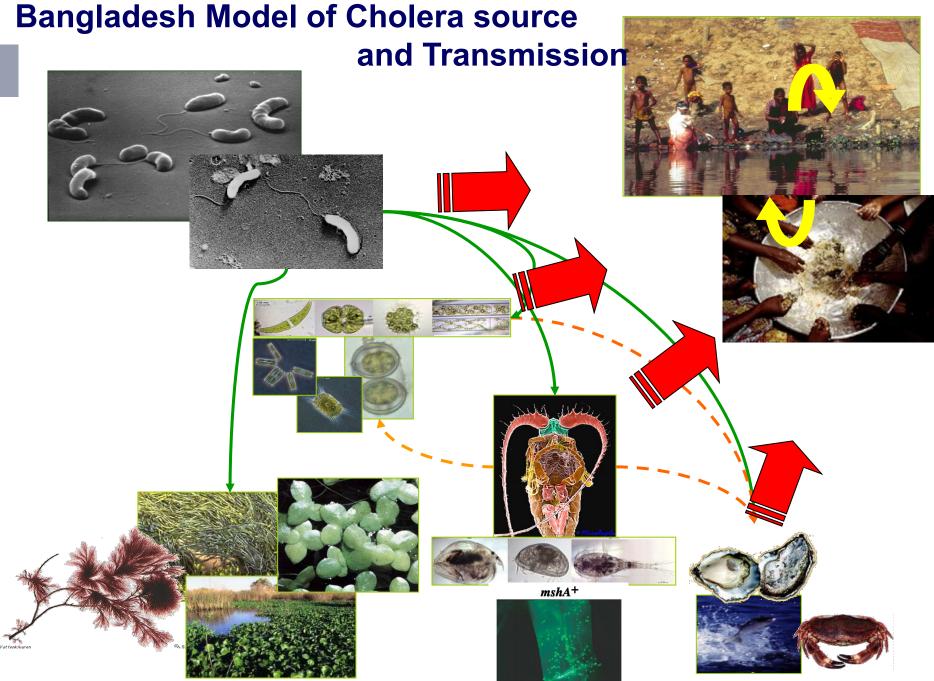
- algae bloom
- phytoplankton bloom

ZOOPLANKTON BLOOM (enters into non-culturable state)

V. CHOLERAE viable but non-culturable state in the water column & attached to particulates. Commensal or symbiotic relationships.

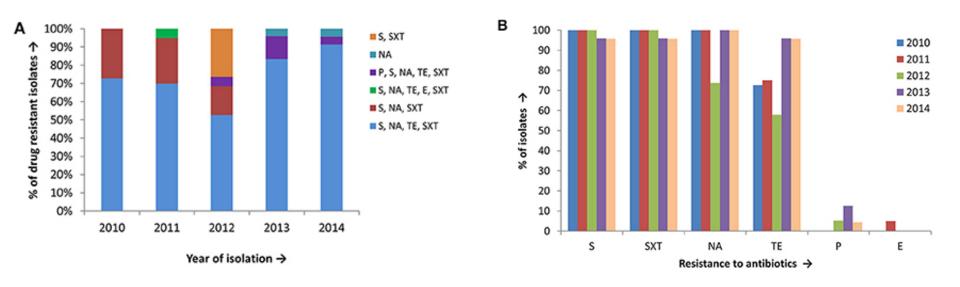
TRANSMISSION OF V. CHOLERAE to humans via ingested water containing colonized copepods or other vectors.



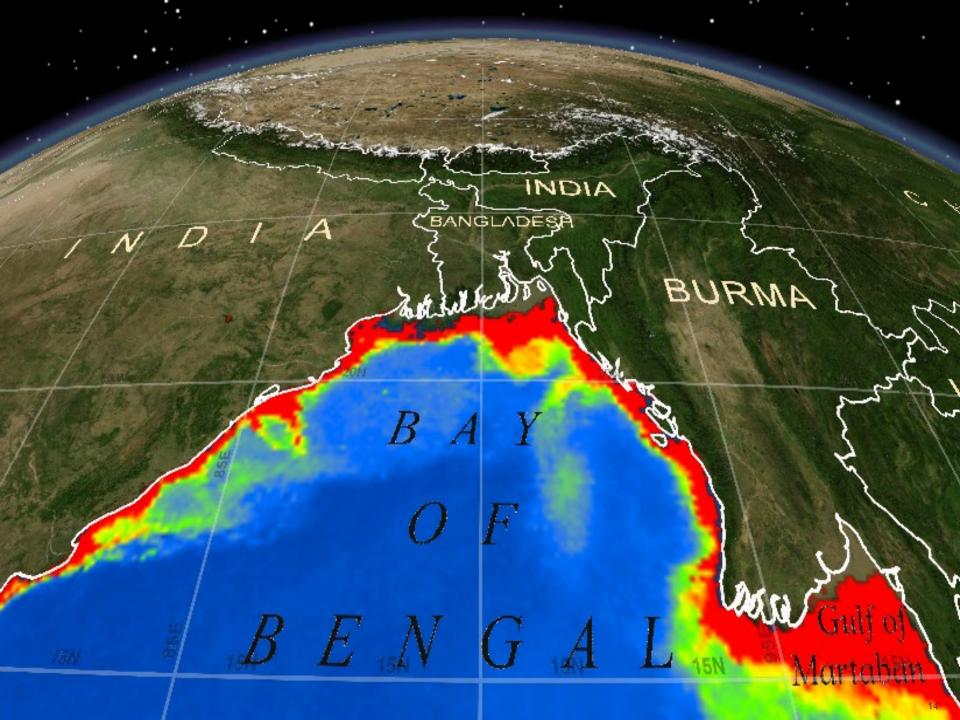


G. Constantin de Magny

Drug resistance profile of *V. cholerae* O1 from environmental sources (Mathbaria 2010-2014)

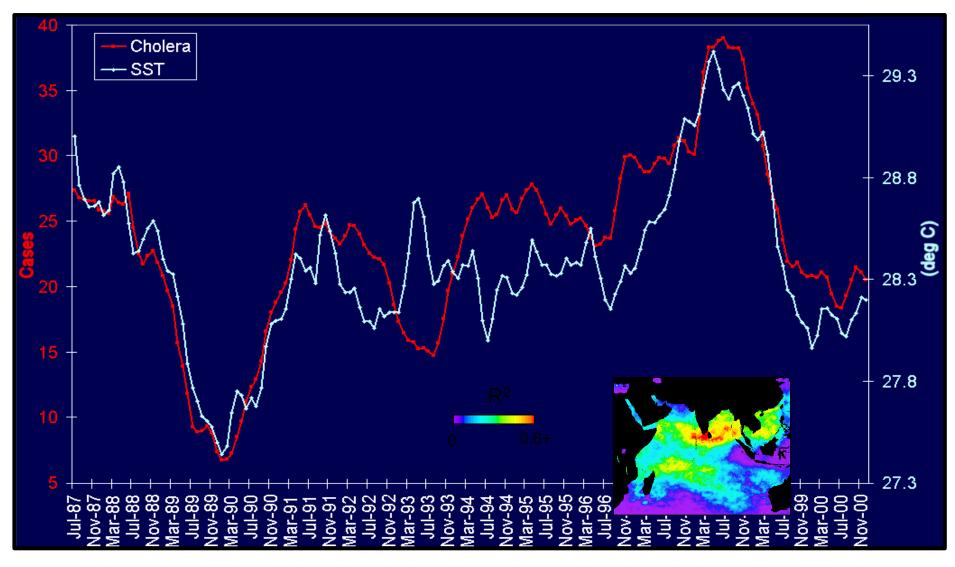


Rashed et al. 2017, Front Microbiol



Cholera and SST in the Indian Ocean 1985 - 2000

Six-month SST lead: R² = 0.72



Lobitz et al., 2000, PNAS Vol. 97, No. 4 pp. 1438-1443

Epidemic Cholera

- Sporadic deadly outbreak
- Usually occurs inland after disasters
- Temperatures may increase growth of bacteria in aquatic bodies.

Typical cholera seasonality

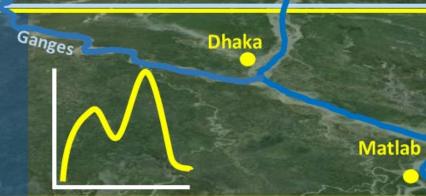


Mixed-mode Cholera

- Usually two seasonal peaks
- One peak related to seawater intrusion; Second peak associated with widespread inundation
- Specific to Bengal Delta region

Endemic Cholera

- Cholera persists throughout year in coastal regions
- Seawater intrusion from coasts to inland



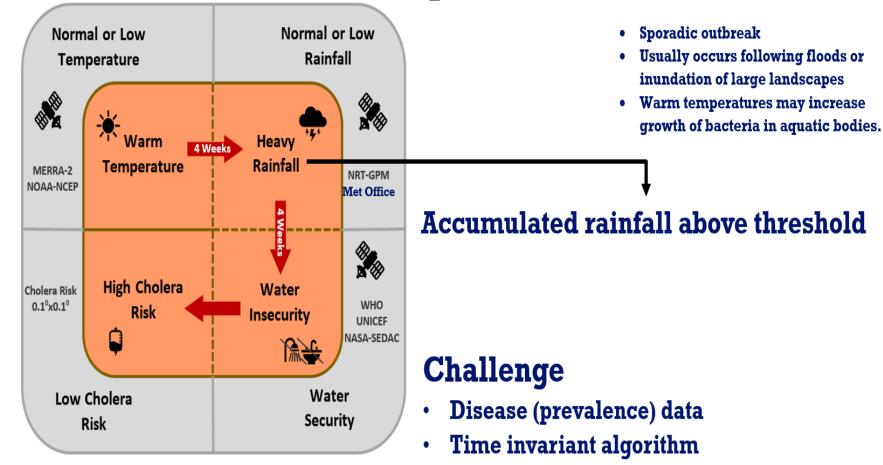
Mathbaria





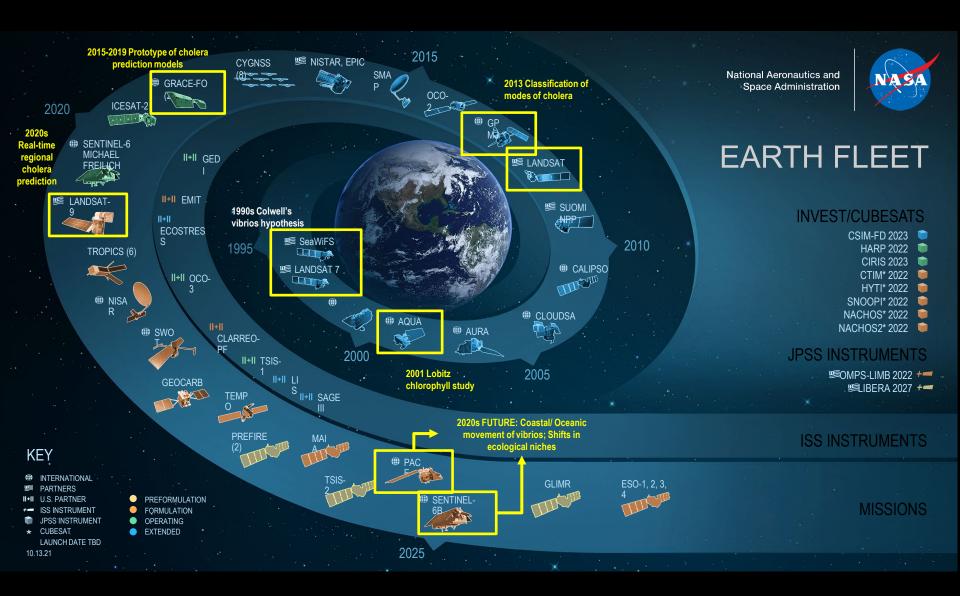
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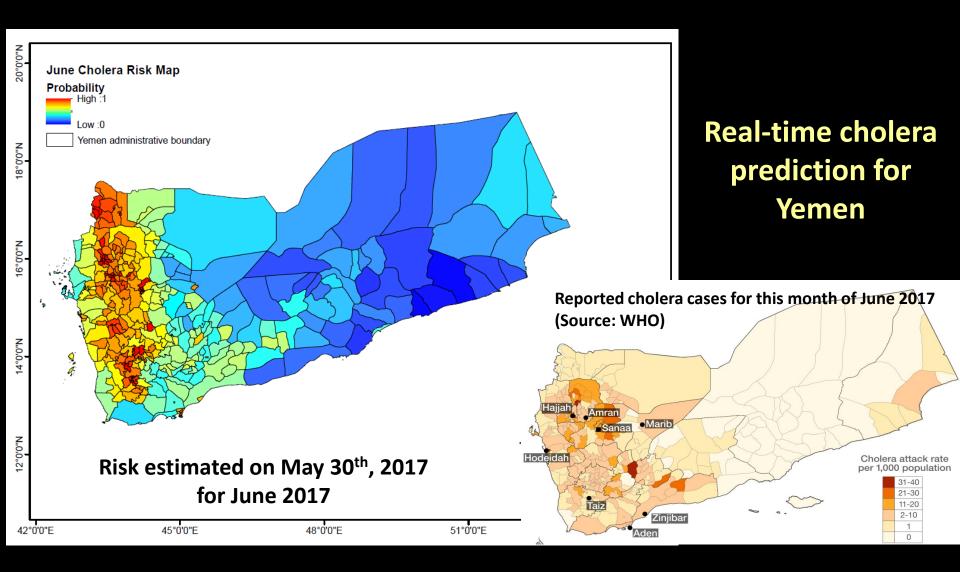
Epidemic Mode of Cholera



Warm temperature= above climatological average temperature Heavy rainfall= above climatological average precipitation Water insecurity=lack of access to water and sanitation access High cholera risk=probability of cholera greater than 50%

• Search for self-adaptive algorithm





Shotgun whole (meta)genome sequencing

Biological specimen

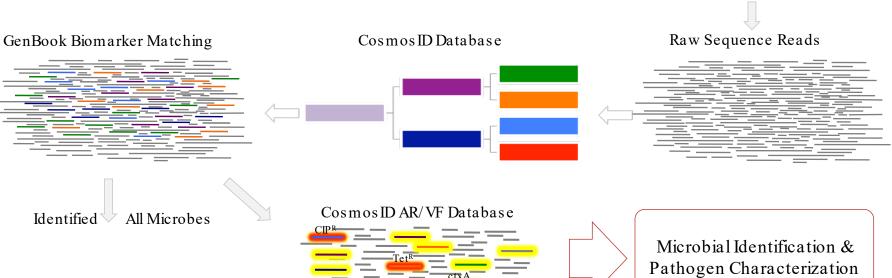


Community DNA



DNA Sequencing

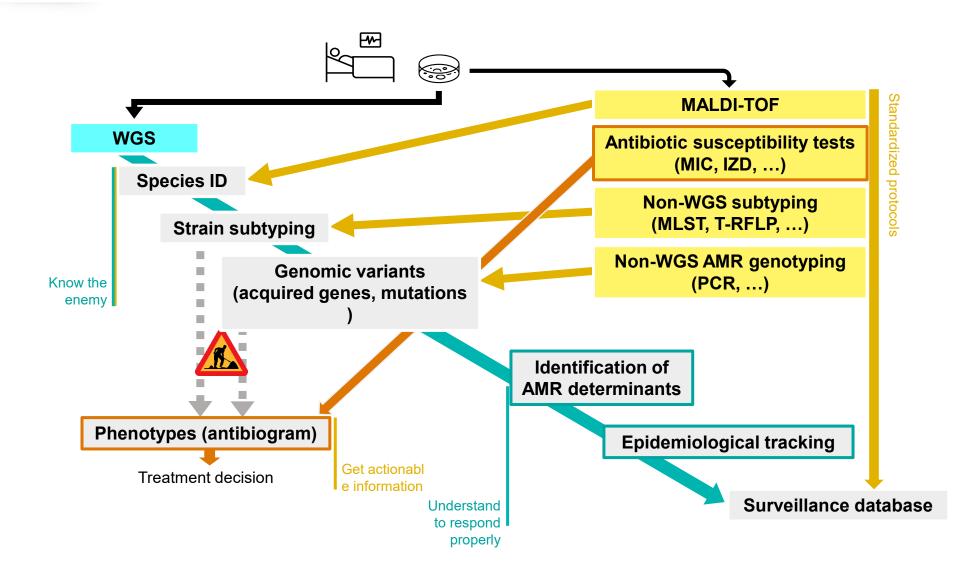
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COSMOSID® Unlocking The Microbiome



What WGS provides



Microbiome Analysis of Acute Diarrheal Patients Compared with Healthy Individuals

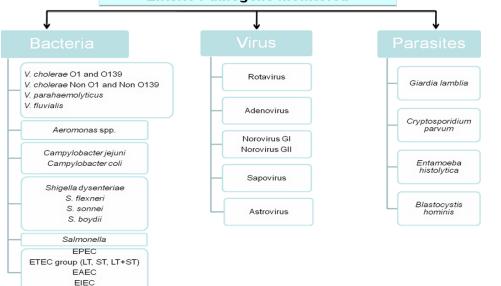


pre-publication results

Study Cohort

@ 2% Surveillance (every 50th patient) at the National Institute of Cholera and Enteric Diseases (NICED), Calcutta, India

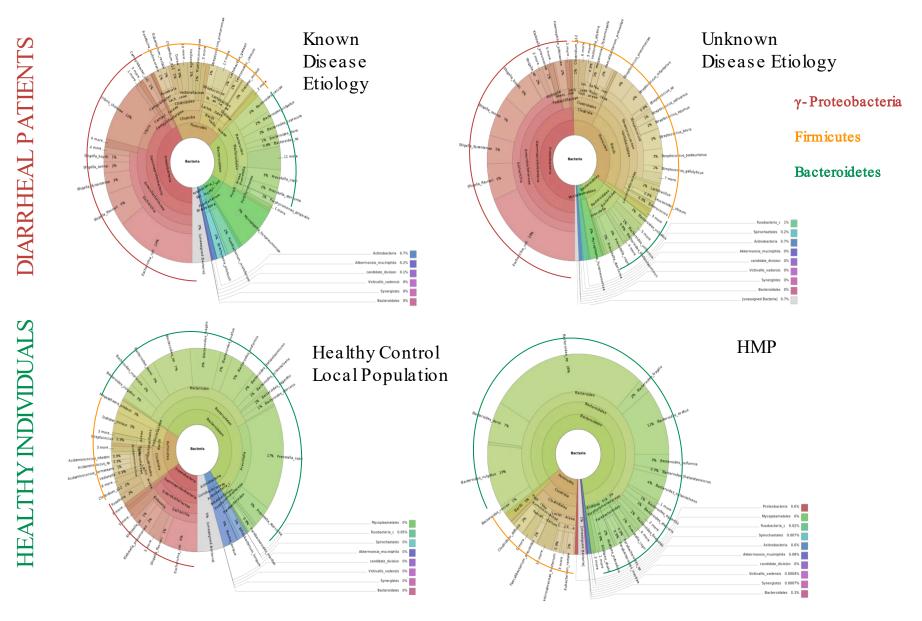
Study Phases	Total # of Samples	Known Etiology	Unknown Etiology	Healthy Control					
PHASE I	9	9	0	0					
PHASE II	28	0	18	10					
PHASE III	37	17	10	10					
	Ent	ric Pathogens monitored							
T		Ļ	\rightarrow						
Ba		Virus	Par						
V. chole	erae O1 and O139 erae Non O1 and Non O139 haemolyticus alis	Rotavirus	Gi	Giardia lamblia					



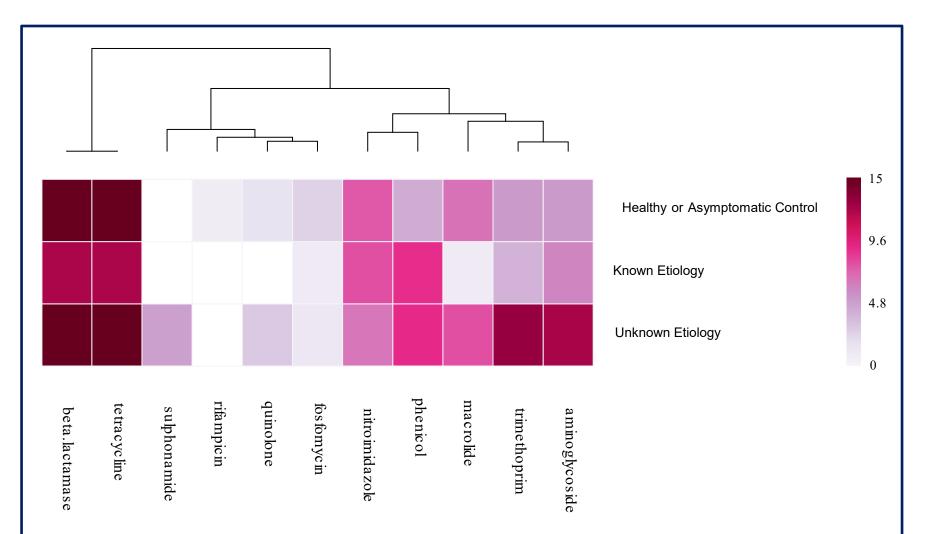
STEC



Microbial Community in Healthy vs Diarrheal Patients

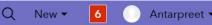


Antimicrobial resistance prevalent in Indian population



 \circ Genes which match at > 50% coverage

• HMP samples had no genes present which matched at this level of coverage







A decision-making initiative for protecting human health and enhancing the resilience of coastal communities under current and changing environments

https://vibrio-prediction-ufl.hub.arcgis.com/

Is COVID-19 polymicrobial and systemic?

How does coronavirus kill? Clinicians trace a ferocious rampage through the body, from brain to toes

Meredith Wadman, Jennifer Couzin-Frankel, Jocelyn Kaiser, Catherine Matacic. Science, Apr. 17, 2020, 6:45 PM

https://www.sciencemag.org/news/2020/04/how-doescoronavirus-kill-clinicians-trace-ferocious-rampagethrough-body-brain-toes



1 Lungs

A cross section shows immune cells crowding an inflamed alveolus, whose walls break down during attack by the virus, diminishing oxygen uptake. Patients cough, fevers rise, and it takes more and more effort to breathe.

2 Liver

Up to half of hospitalized patients have enzyme levels that signal a struggling liver An immune system in overdrive and drugs given to fight the virus may be causing the damage.

3 Kidneys

Kidney damage is common in severe cases and makes death more likely. The virus may attack the kidneys directly, or kidney failure may be part of whole-body events like plummeting blood pressure.

4 Intestines

Patient reports and biopsy data suggest the virus can infect the lower gastrointestinal tract, which is rich in ACE2 receptors. Some 20% or more of patients have diarrhea.

5 Brain

Some COVID-19 patients have strokes, seizures, mental confusion, and brain inflammation. Doctors are trying to understand which are directly caused by the virus.

6 Eyes

Windpipe

Bronchii

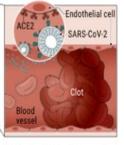
Bile duct

a

Conjunctivitis, inflammation of the membrane that lines the front of the eye and inner eyelid, is more common in the sickest patients.

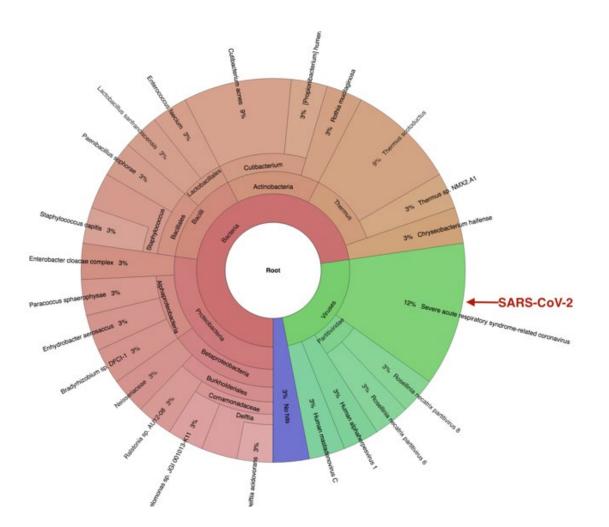
7 Nose

Some patients lose their sense of smell. Scientists speculate that the virus may move up the nose's nerve endings and damage cells.



8 Heart and blood vessels The virus (green) enters cells, likely including those lining blood vessels, by binding to ACE2

vessels, by binding to ACE2 receptors on the cell surface. Infection can also promote blood clots, heart attacks, and cardiac inflammation. Identification of Bacteria and Viruses Present in Respiratory Samples in which SARS-CoV-2 has been Detected



SARS Cov-2 viral RNA has been detected in 48.1% of stool samples

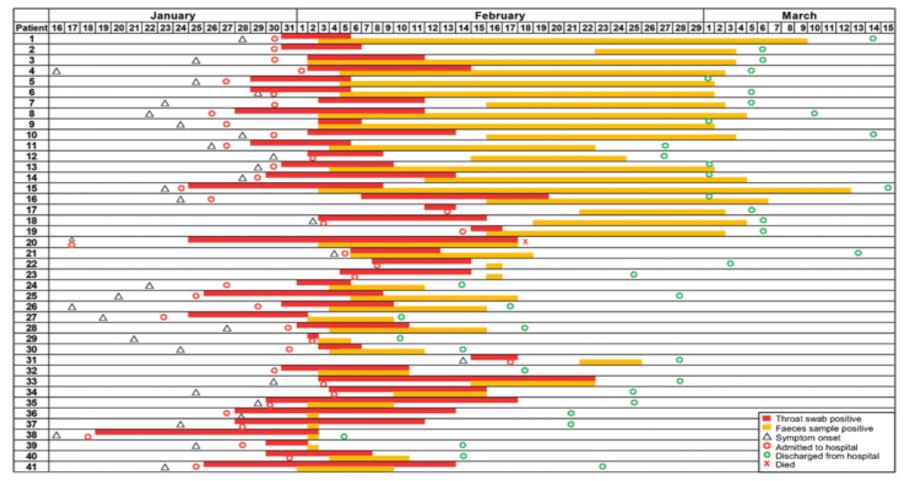
Stool viral RNA positivity rate

							P	revalen	ce		Weight	Weight
Study	Events	Total						(%)		95%-CI	(fixed)	(random)
Xiao F	39	73		- 2				53.42	[41.37;	65.20]	55.8%	45.2%
Zhang W	4	15	- *	4	-			26.67	[7.79]	55.10]	9.0%	10.9%
Zhang J	5	14		* 1				35.71	[12.76]	64.86]	9.9%	11.9%
Wang W	5	13		* 1				38.46	[13.86]	68.42]	9.4%	11.4%
Young BE	4	8						50.00	[15.70]	84.30]	6.1%	7.7%
Kim JY	0	2 ⊷						0.00	[0.00;	84.19]	1.3%	1.7%
Yang Z	3	3		4			-	100.00	[29.24;	100.00]	1.3%	1.8%
Cheng SC	0	1 ⊷					_	0.00	[0.00;	97.50]	1.2%	1.5%
Holshue	1	1 -					-	100.00	[2.50;	100.00]	1.2%	1.5%
Cai J	5	6					_	83.33	[35.88]	99.58]	2.6%	3.3%
Zeng L	1	1 -					-	100.00	[2.50;	100.00]	1.2%	1.5%
Zhang Y	1	1 -		-			-	100.00	[2.50;	100.00]	1.2%	1.5%
Fixed effect model		138		-	>			48.88	[40.41;	57.41]	100.0%	-
Random effects model				<	>		_			57.94]		100.0%
Heterogeneity: $I^2 = 7\%$, τ^2	= 0.0361,	p = 0.38	1	1		1	1					
		0	20	40	60	80	100	D				

Gastroenterology

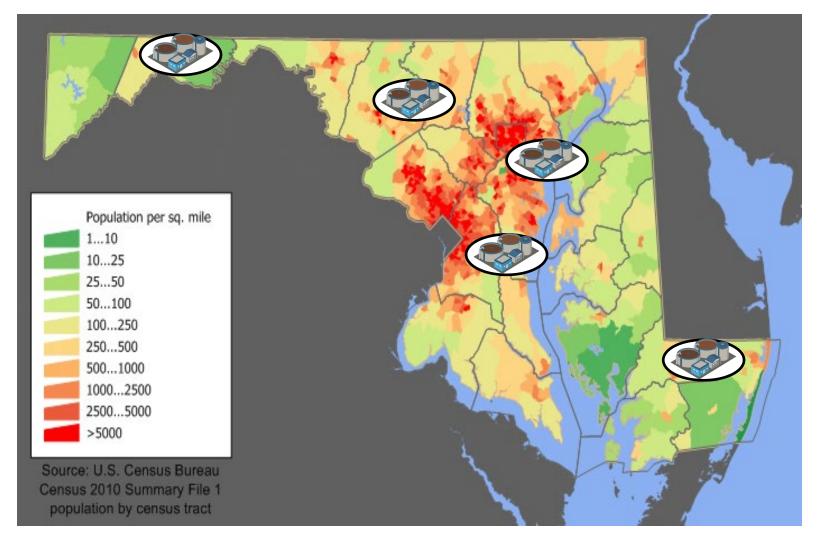
Cheung et al., (2020). Gastrointestinal Manifestations of SARS-CoV-2 Infection and Virus Load in Fecal Samples from the Hong Kong Cohort and Systematic Review and Meta-analysis. Gastroenterology. Pre-Proof

Positive Stool Samples Detected After Respiratory Sample Tested Negative During Recovery

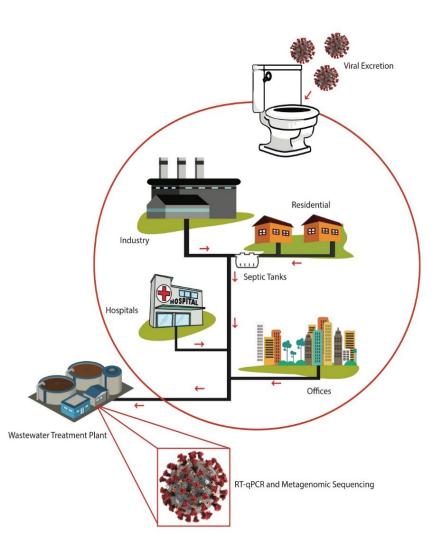


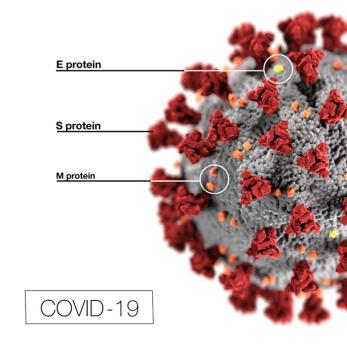
Wu Y, Guo C, Tang L, et al. Prolonged presence of SARS-CoV-2 viral RNA in faecal samples. 2020 The lancet Gastroenterology – hepatology. Volume 5, Issue 5, 434 - 435

COVID-19 tracking in wastewater in Maryland, USA, 2020-2022

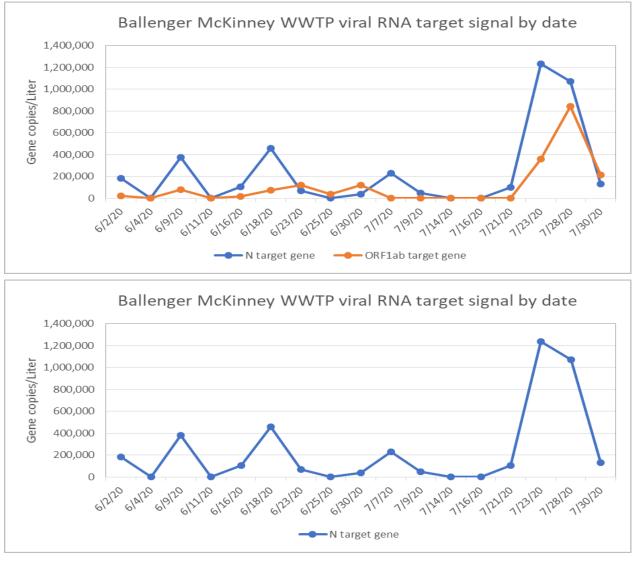


Wastewater surveillance





Results from Frederick, Maryland sites, 2020





Case Study: Mount St. Mary's University

) Home

() Share

> P 27

- Twice weekly sampling of dormitory effluent
- Covid spike triggered testing of individual students
- 221 students tested
 - 10 positive
 - 9 asymptomatic
- "It could have become quite a spreading event," said Donna Klinger, a spokeswoman for the university
- Coronavirus positive students isolated and wastewater tests done twice weekly

Sections E Other Washington Abst Get one year for \$29 Sign in 1 Descring Dir in Carbons

Maryland Politics

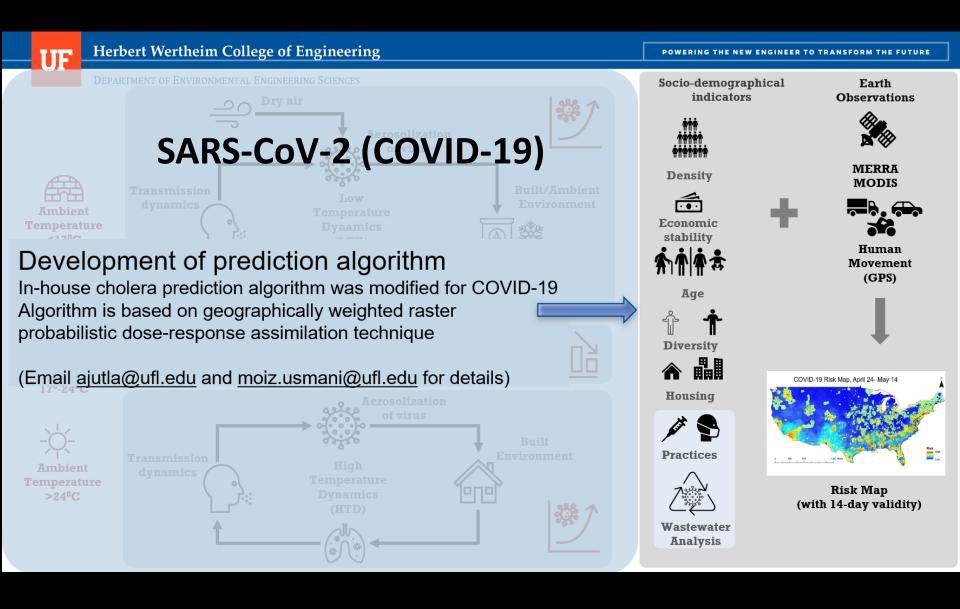
Flushing out the coronavirus: Universities, cities and states are testing wastewater for the virus



Alan Matow-Bariges collects untrested wastewater at the Ballenger-VcKinney treatment facility in Frederick, Md, on Oec. 18. Twice a week, samples are collected and sent to a lab in Rockville to be screened for the coronavirus. (Katherine Frey/The Washington Posc)

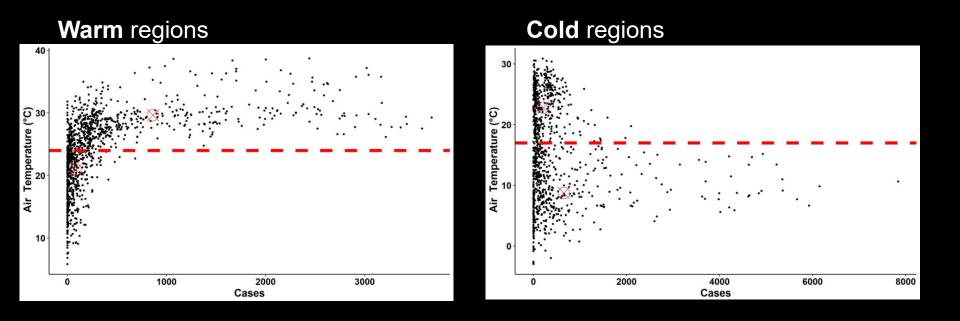






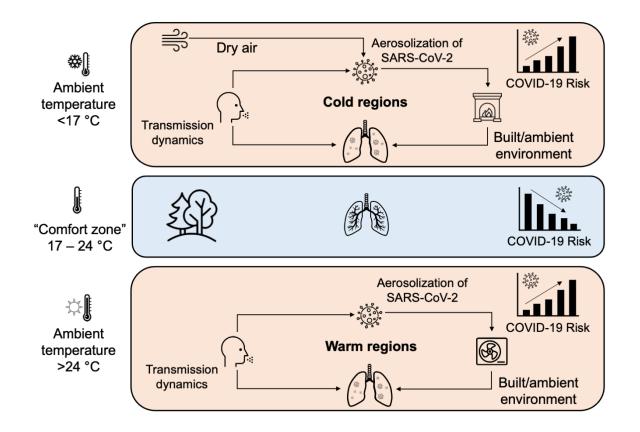
https://covid-ufl.hub.arcgis.com/apps/covid-19-risk-map/explore

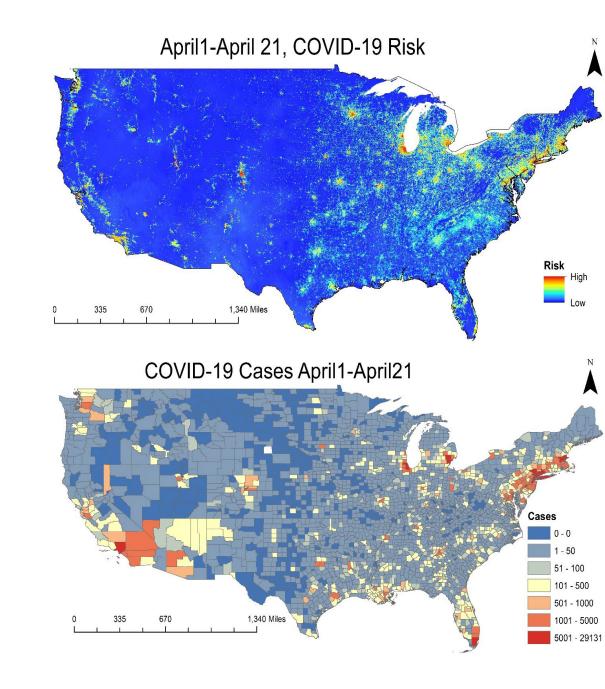
Clustering of disease prevalence vs ambient air temperature



Usmani et al. 2022, AM J Trop Med Hyg

Hypothesis for environmental COVID-19 risk prediction

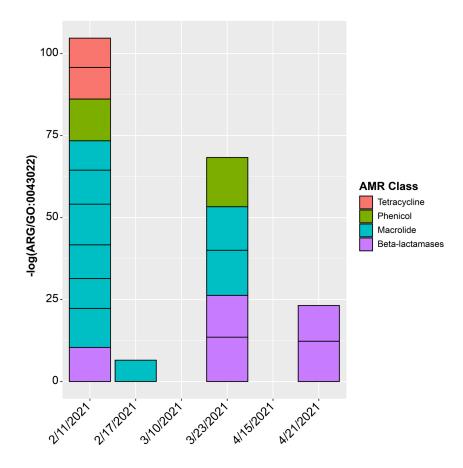




Environmental COVID-19 risk prediction

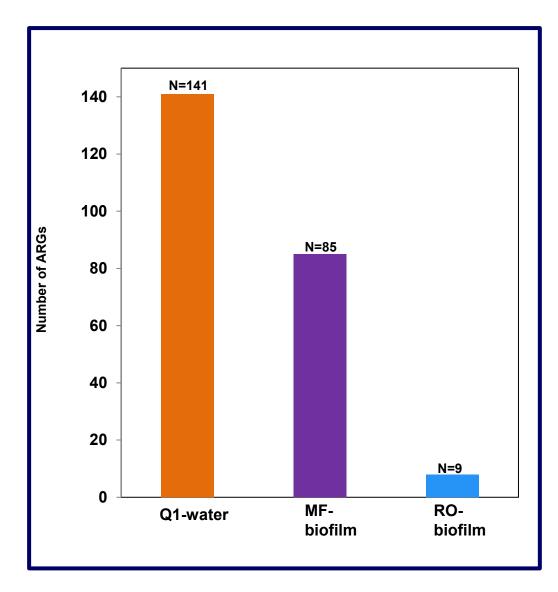
Usmani, Brumfield, Jutla, & Colwell; *in preparation*

Expression of AMR in wastewater (RNA-seq)

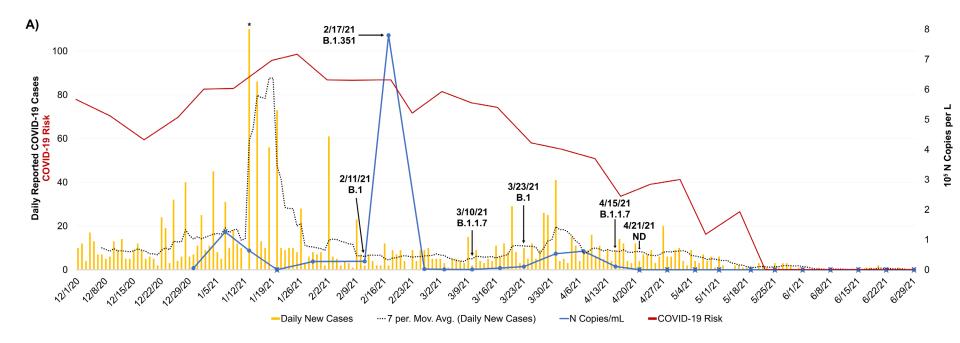


Brumfield et al. 2022, mBio Vol. 13, No. 4, Microbiome Analysis for Wastewater Surveillance during COVID-19

Distribution of antibiotic resistance genes (ARGs) and stepwise reduction of ARG's in MF and RO-biofilms

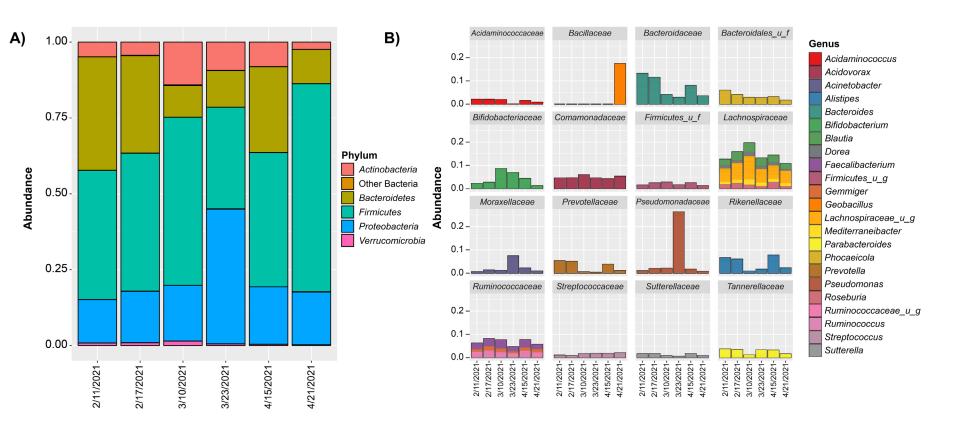


Detection of SARS-CoV-2 in wastewater and predicted risk

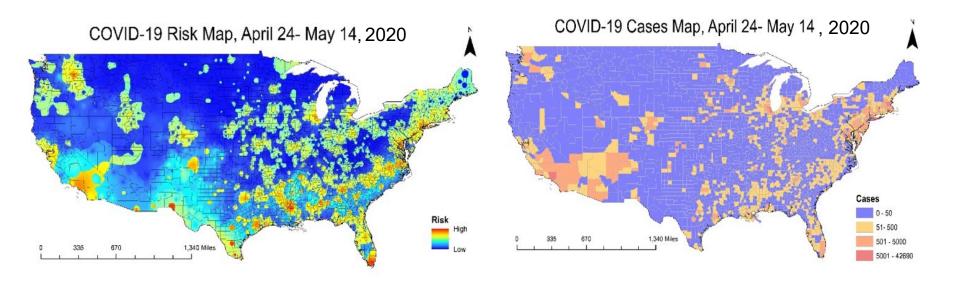


Brumfield et al. 2022, under review

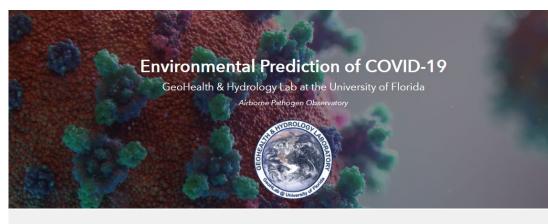
Microbiome profiles of wastewater (DNA metagenomics)



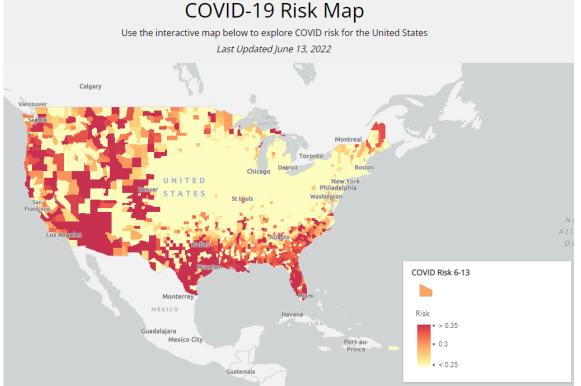
Prediction of coronavirus risk



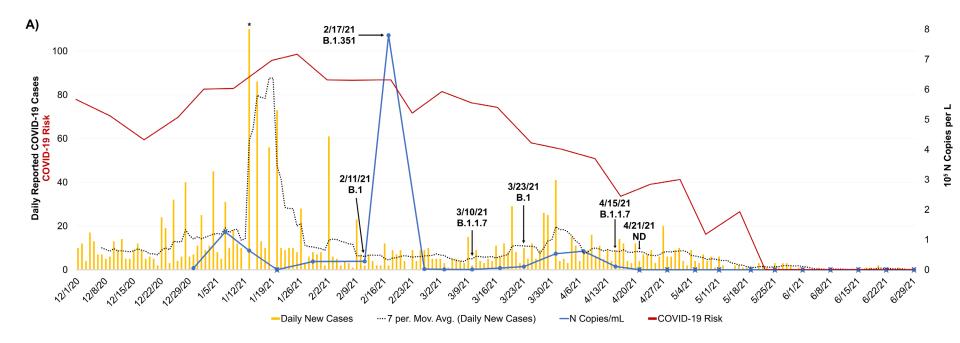
Left panel: **Prediction** made on April 24th 2020 and valid until May 14th, 2020. Right panel: **Actual number** of COVID19 cases during those three weeks: a heuristic prediction model developed in GeoHLab DEPARTMENT OF ENVIRONMENTAL ENGINEERING SCIENCES



https://covidufl.hub.arcgis.com/

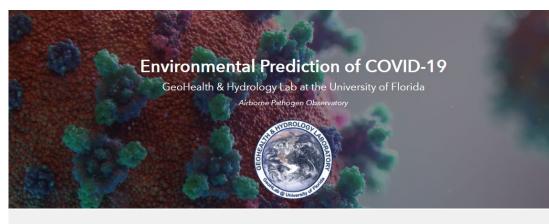


Detection of SARS-CoV-2 in wastewater and predicted risk

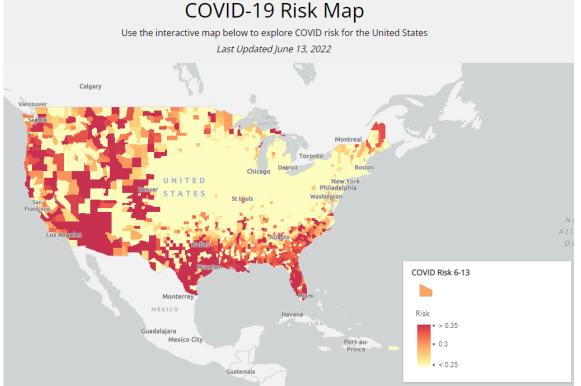


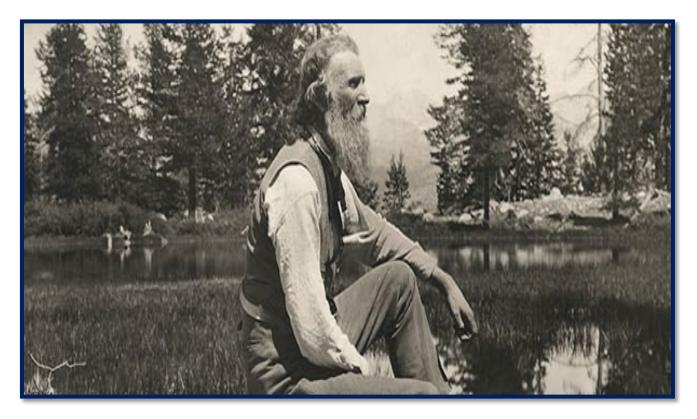
Brumfield et al. 2022, under review

DEPARTMENT OF ENVIRONMENTAL ENGINEERING SCIENCES



https://covidufl.hub.arcgis.com/





"When one tugs at a single thing in nature, he (and she) find it hitched to the rest of the universe."

> John Muir (1838-1914)

Collaborators and Colleagues

ICDDR,B - Dhaka

- Dr. Tahmeed Ahmed
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- A.K. Ashraful Aziz
- Dr. A.S.G. Faruque
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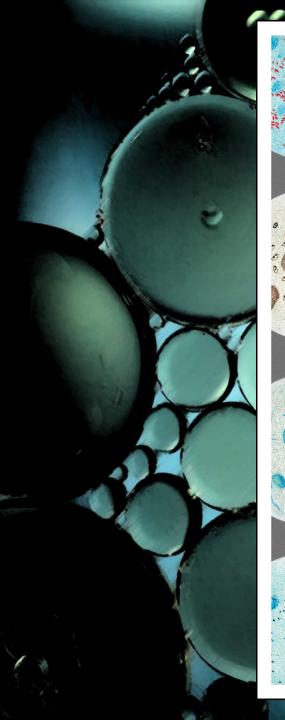
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- Patrick Monfort, Univ of Montpellier, FR

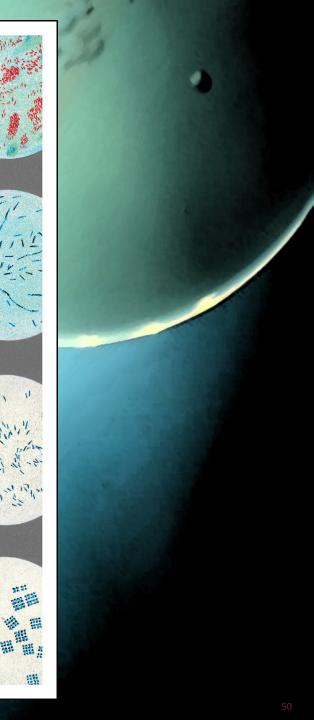
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- Moiz Usmani, University of Florida
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- Miguel Talledo, Lima, Peru
- Ron Taylor, Darmouth, College, NH
- Byron Wood, NASA Ames
- Huai-shu Xu, China
- Linda Zall, Office of the Chief Scientist
- Fred Zimmerman, ISciences, Burlington, VT



A Lab of One's Own

One Woman's Personal Journey Through Sexism in Science

Former Director of the National Science Foundation
RITA COLWELL, PHD
and
SHARON BERTSCH McGRAYNE







QUESTIONS AND DISCUSSION



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