

CENTER FOR
INFERENCE &
DYNAMICS
OF INFECTIOUS DISEASES



Pandemics in the age of Big Data

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LABORATORY FOR THE MODELING OF BIOLOGICAL
AND SOCIO TECHNICAL SYSTEMS



CENTER FOR
INFERENCE &
DYNAMICS
OF INFECTIOUS DISEASES



EBOLA
CHALLENGE

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M.Litvinova, D.Mistry, A.
Pastore y Piontti, K.Sun,
S.Haque, N. Samay,
Q.Zhang,
(Northeastern University, USA)

M.E. Halloran
(Fred Hutchinson Cancer Research
Center, USA)

N. Dean, D. Rojas,
I.M. Longini
(University of Florida, USA)

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(Greenwich University, UK)

S. Merler, L. Fumanelli, P.
Poletti
FBK, Trento, Italy

C.Gioannini, L.Rossi,
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P. Milano, M.Selim, E.Ubaldi,
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Italy)

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G.Chowell C.Viboud
(Georgia State University) (Fogarty, NIH, USA)

L.Simonsen
(George Washington University, USA)

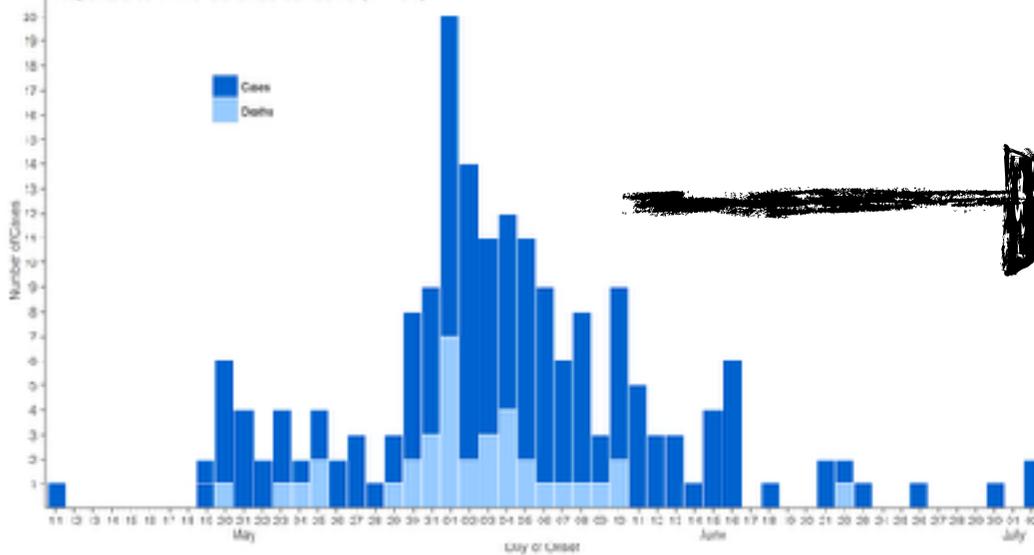
Epidemic modeling

“I simply wish that, in a matter which so closely concerns the wellbeing of the human race, no decision shall be made without all the knowledge which a little analysis and calculation can provide”

Daniel Bernoulli ~1760

Confirmed cases of MERS-CoV in the Republic of Korea and China

Reported to WHO as of 22 Jul 2015 (n=186)

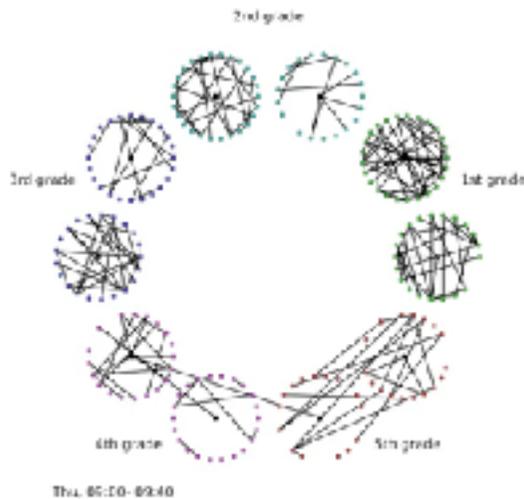


Please note that the underlying data is subject to change as the investigations around cases are ongoing. Case data estimated if not available. Source: WHO

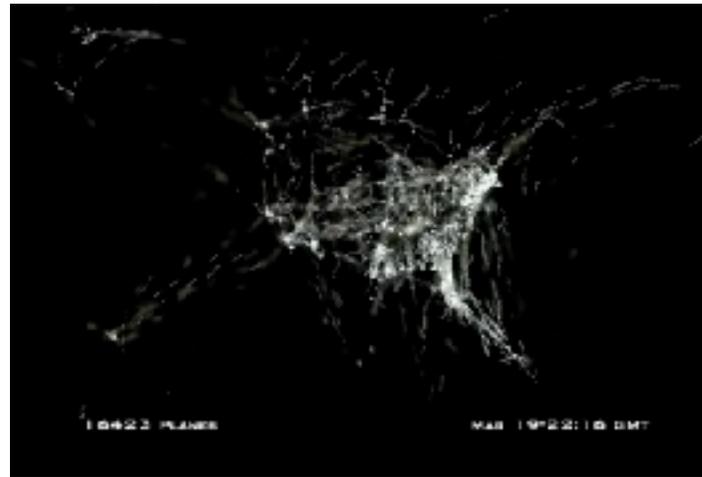


DATA AVAILABILITY HAS EXPLODED

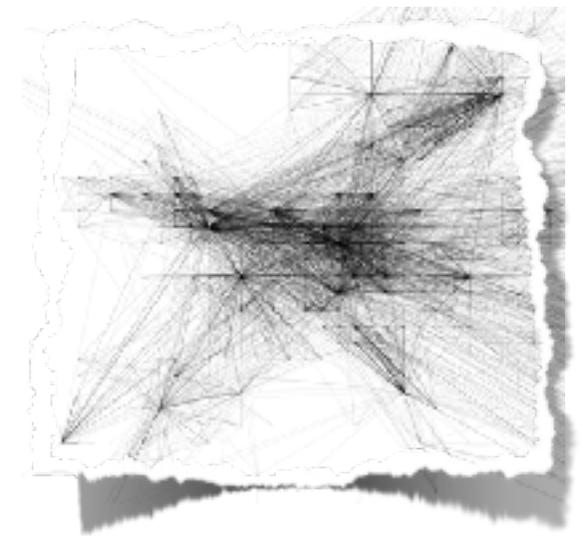
Human interactions/
contact networks



Mobility and
epidemic spreading



Networks
heterogeneity and
complexity



Within school contact
patterns
(@Sociopatterns)

Multiscale integration
of mobility networks in
the analysis of
potentially pandemic
pathogens spread.

Hubs, community,
clustering, heavy
tails, ...

Big Data

Big data narrative, fourth paradigm etc.

Large number of pitfalls:

- no dynamical understanding
- black box (algorithmic issues)
- See Hosni, Vulpiani, Philosophy & Technology (2017) MUST READ!

~~BIG DATA~~ → NEW DATA



New data (big, or small)

- The focus is on understanding these data sets in a scientific sense and more deeply the real world processes which produced the data (Theory)
- Mechanistic approach
- Effective equations
- Initial conditions
- Model's calibration

A MULTIDISCIPLINARY APPROACH TO EPIDEMIC ANALYSIS

01

MODELING

Elaborate stochastic infectious-disease models to support a wide range of epidemiological studies, covering different types of infections and intervention scenarios.

03

COMPUTATIONAL THINKING

The computer is the laboratory. Models run on high performance computers to create in-silico experiments that would be hardly feasible in real systems, to guide our understanding of typical non-linear behavior and tipping points of epidemic phenomena.

02

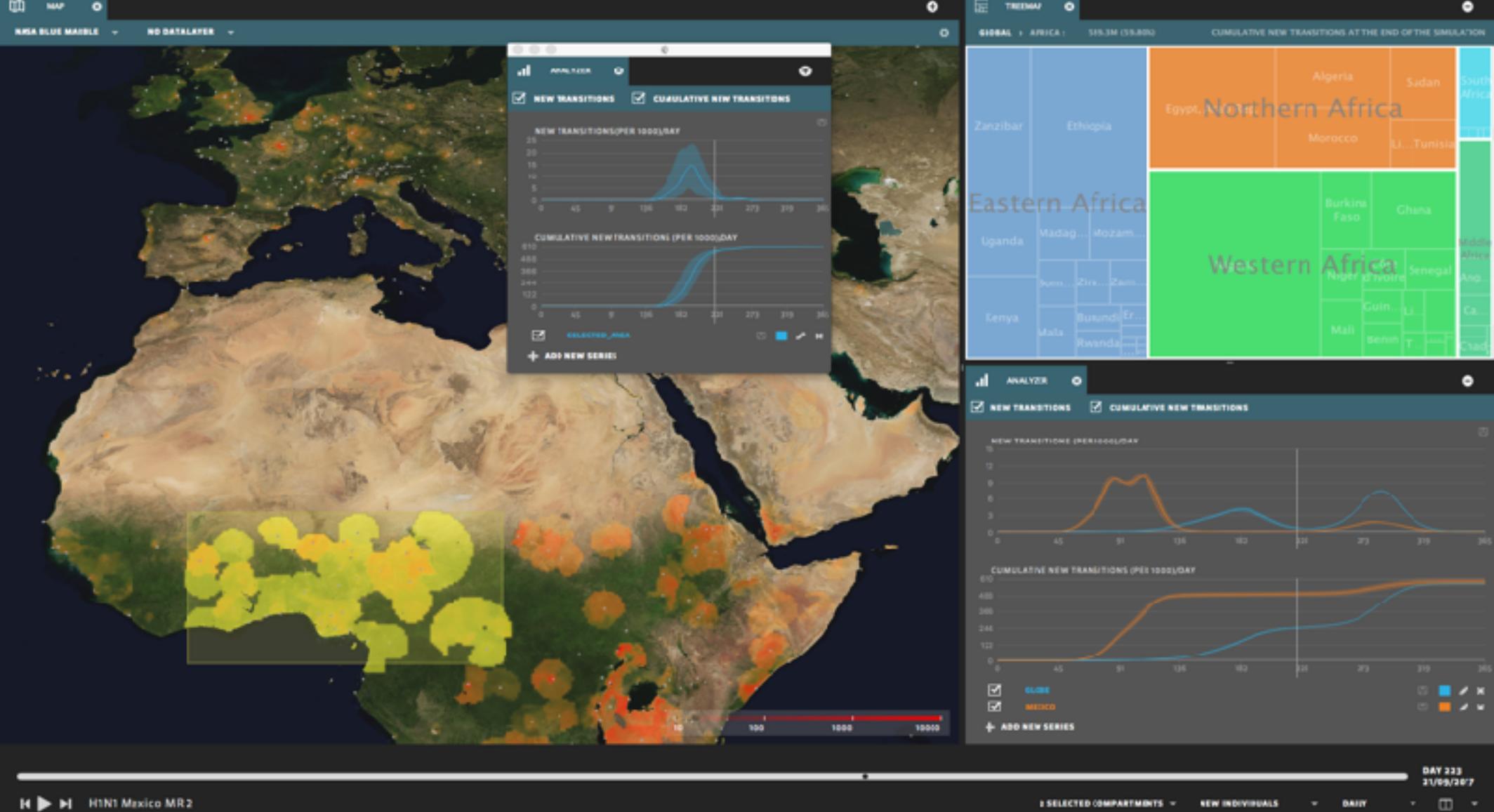
REAL-WORLD DATA

Real-world data on population and mobility networks are integrated in structured spatial epidemic models to generate data-driven simulations of the worldwide spread of infectious diseases.

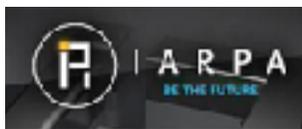
04

TOOLS DEVELOPMENT

Computational tools help in modeling the spread of a disease, understanding observed epidemic patterns, and studying the effectiveness of different intervention strategies. These tools are available to researchers, healthcare professionals, and policy makers.



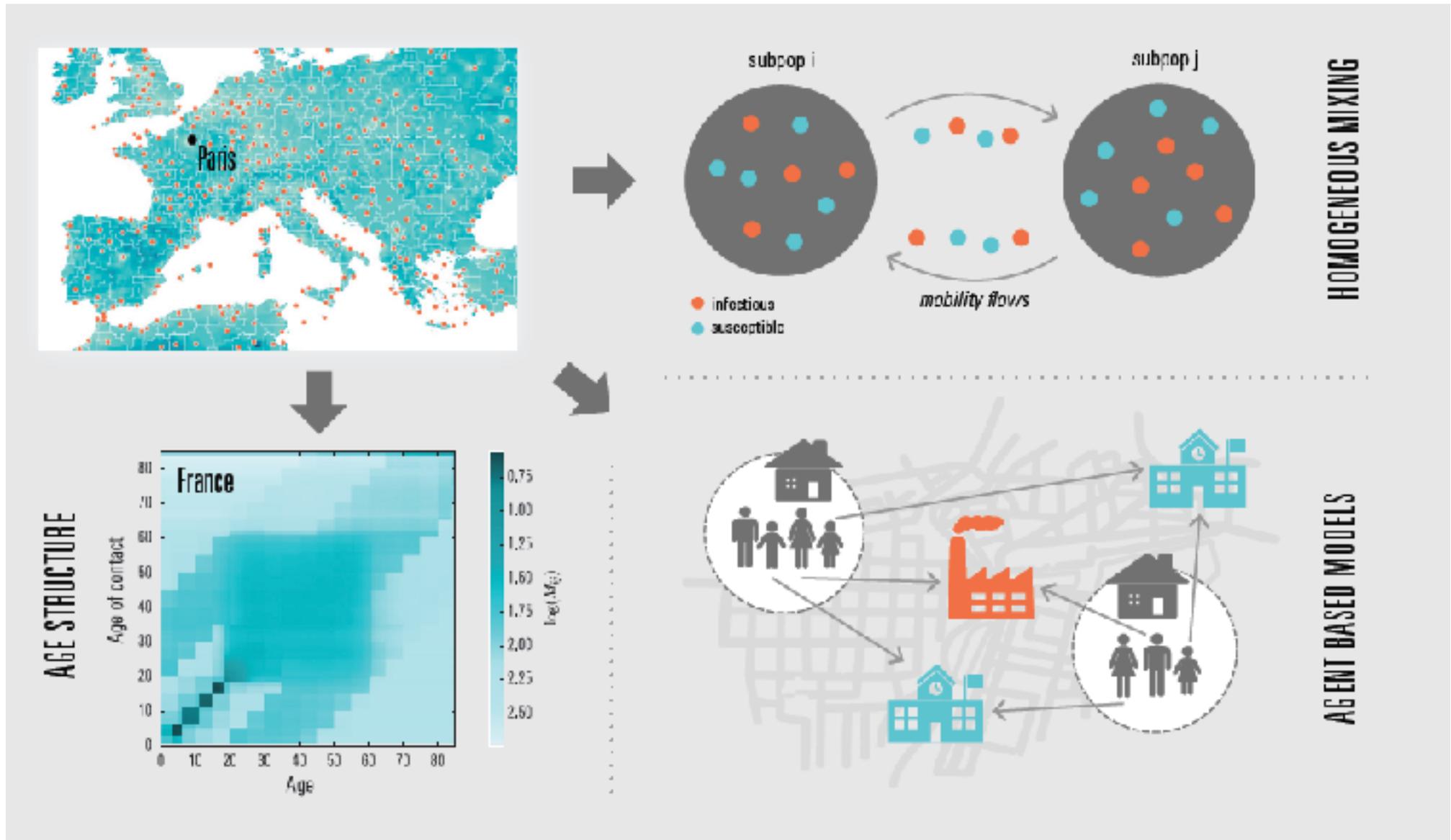
WWW.GLEAMVIZ.ORG



Stochastic Inter population dynamics



Multiple schemes for the stochastic intra-population contagion dynamic



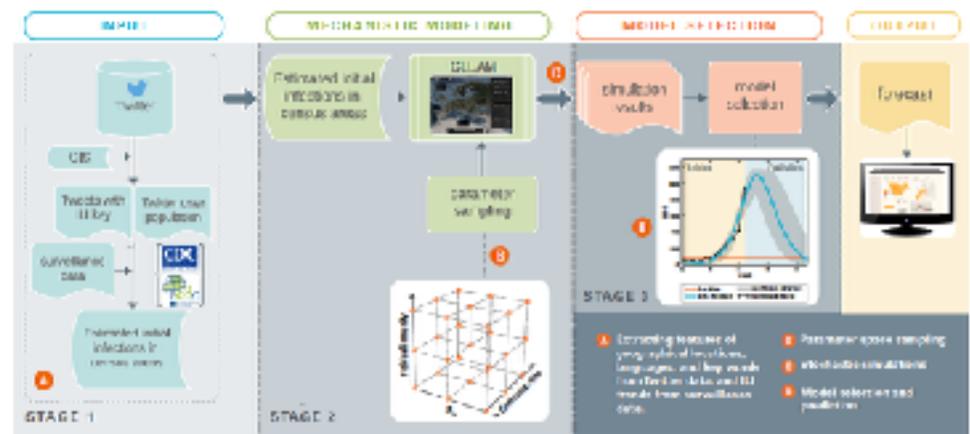
Applications

- SARS Epidemic (retrospective)
- 2009 H1N1pdm
- 2014 West Africa Ebola Epidemic
- Zika epidemic in Latina America
- CDC yearly challenge since 2012

This work has been in collaboration with the CIDID, MIDAS network and a number of national and International Institution.

Research & Development

- Predictability limits/frameworks (Ebola synthetic challenge)
- Synthetic contact matrices
- Forecasting Mechanistic approach with the help of novel data streams.



H1N1 ppm invasion tree

Feb 18 2009



GLEaMviz.org

Chicago
New York
Los Angeles
Houston
Toronto
Vancouver
Calgary
Indianapolis

La Gloria

Sao Paulo
Mexico City
Rio De Janeiro
San Juan
Bogota

Johannesburg
Cairo
Cape Town
Nairobi

Paris
Frankfurt
Amsterdam
Rome
Milan
Moscow
Dublin

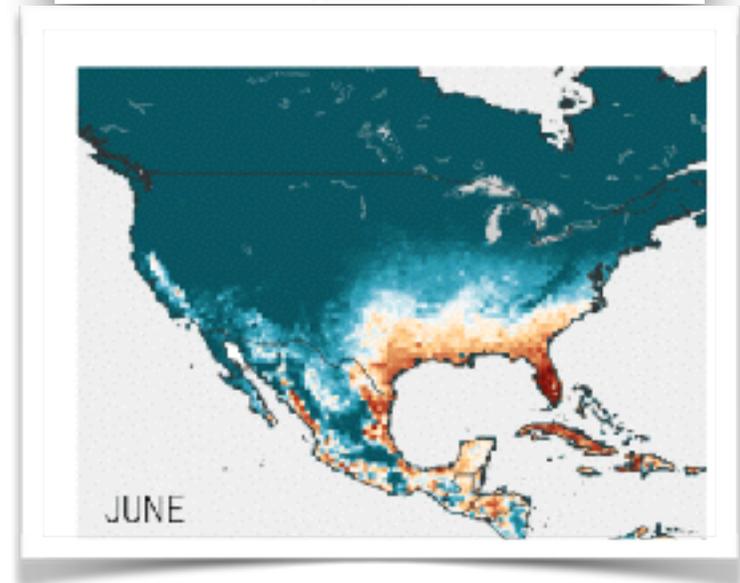
Hong Kong
Tokyo Narita
Bangkok
Singapore
Beijing
Manila

Sydney
Brisbane
Auckland
Perth

Spatial stochastic individual based model

Zhang et al. PNAS 2017 ; doi:10.1073/pnas.1620161114

- Introduce explicitly the coupling of **traveling patterns** (case importation and colonization) on the **disease progression**
- Introduce **seasonal drivers of Mosquito** transmission in the epidemic dynamic.
- Introduce effect of **socio-economic drivers**
- Interplay of traveling pattern, outbreak initial conditions, disease dynamic and seasonal driving in defining the epidemic progression at the regional level.

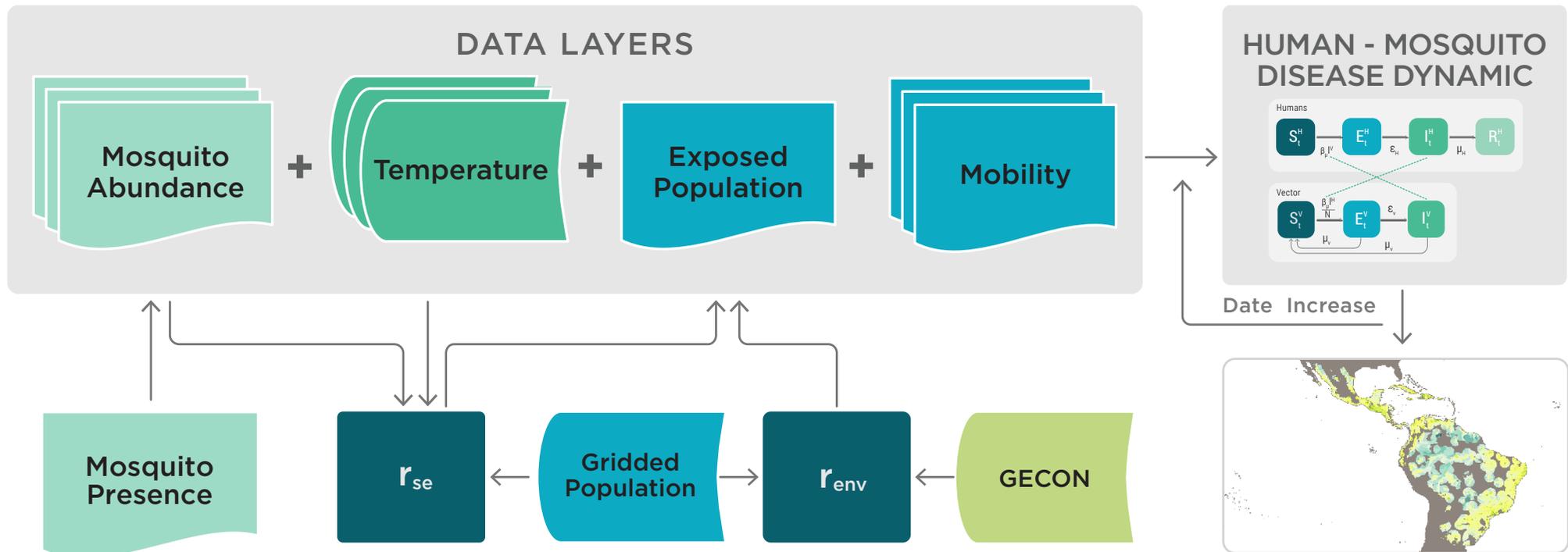


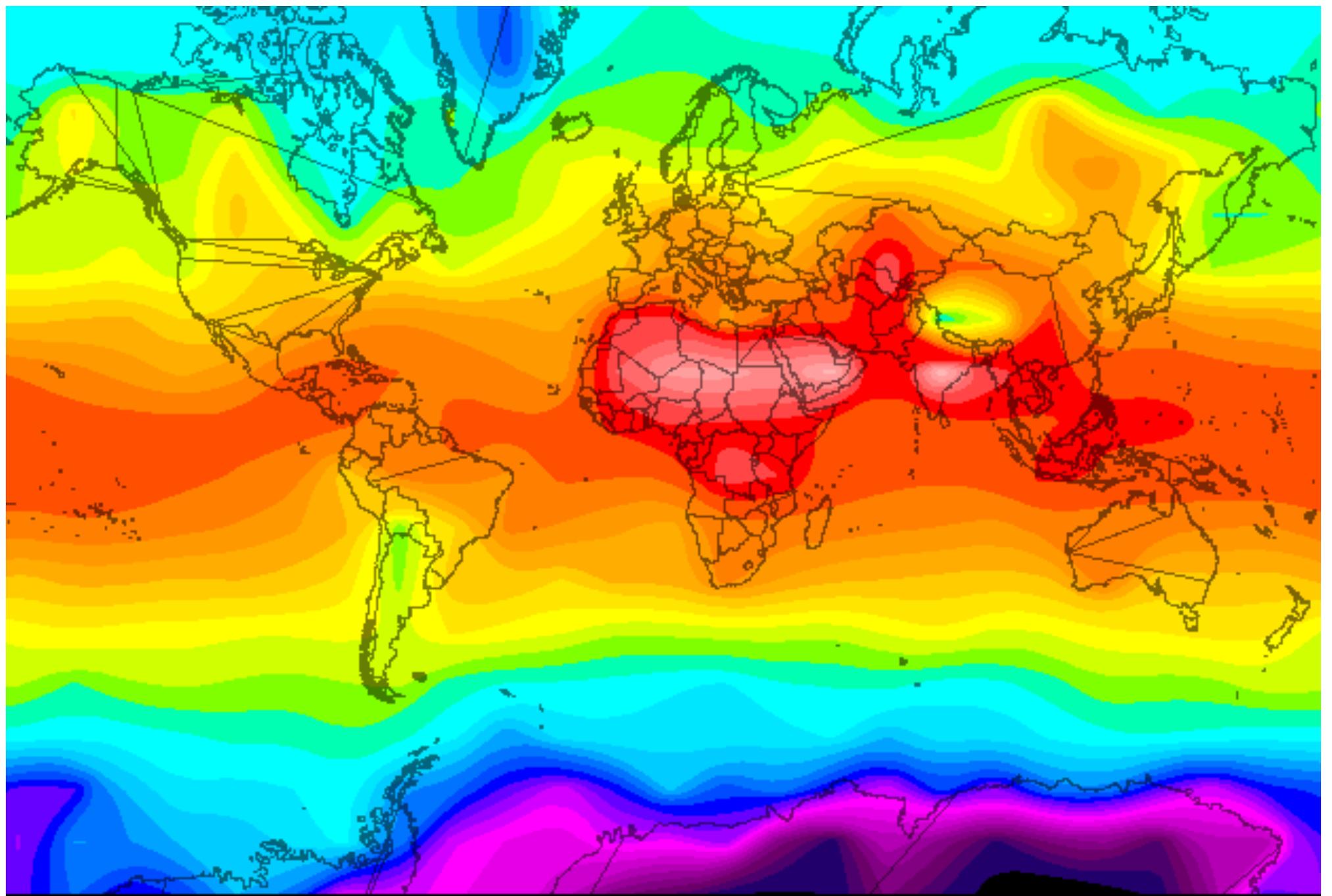
Model structure

Explicit modeling of
airline traffic national/
international +
commuting patterns
and local mobility

Mosquitoes abundance
+ local climate drivers +
socio-economic
indicators

Dynamic stochastic
model providing time
evolution of the
epidemic





Today's High Temperatures

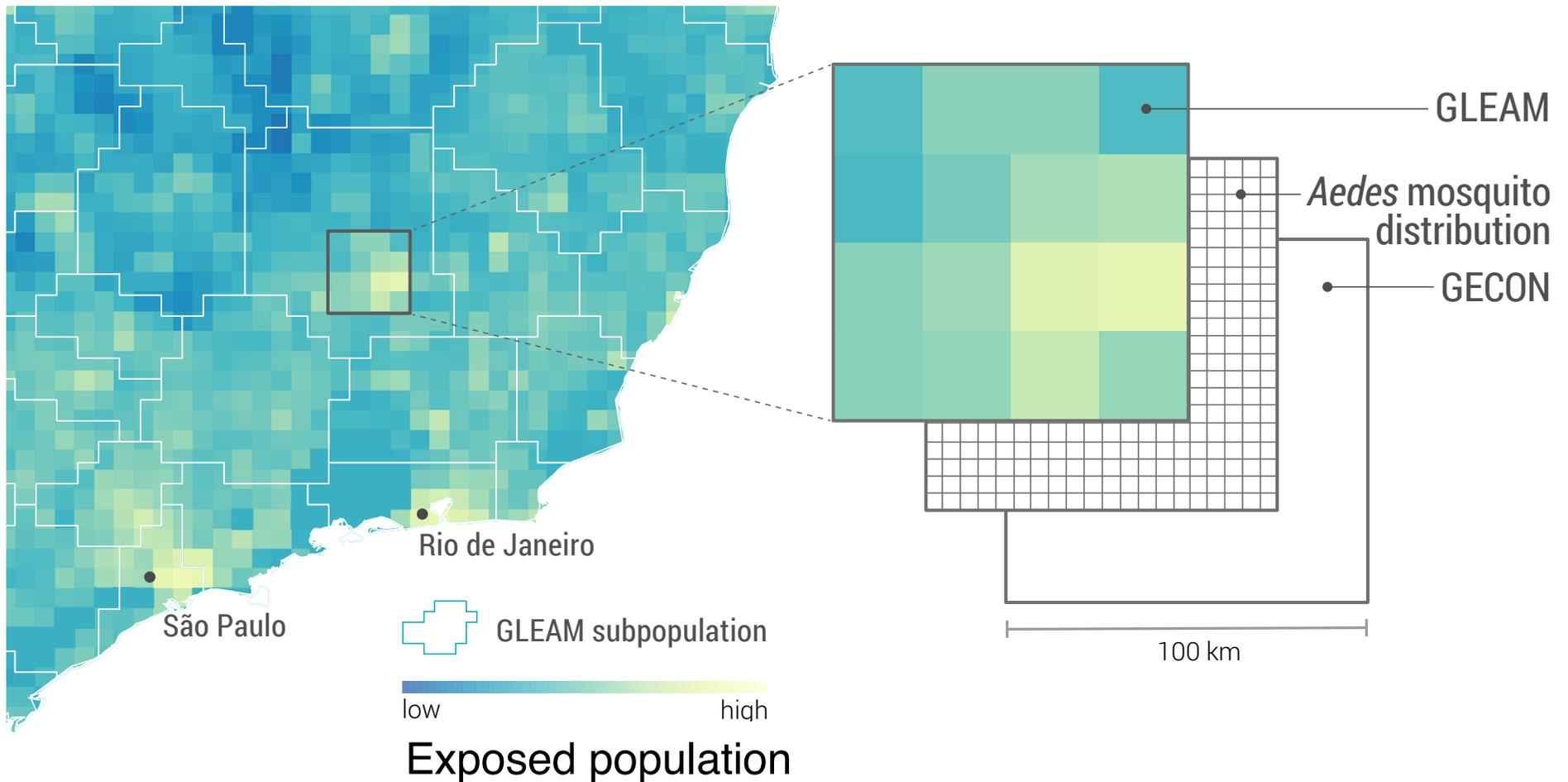
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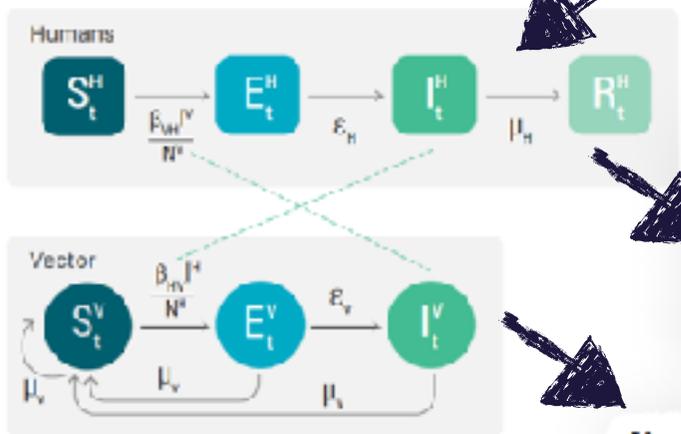
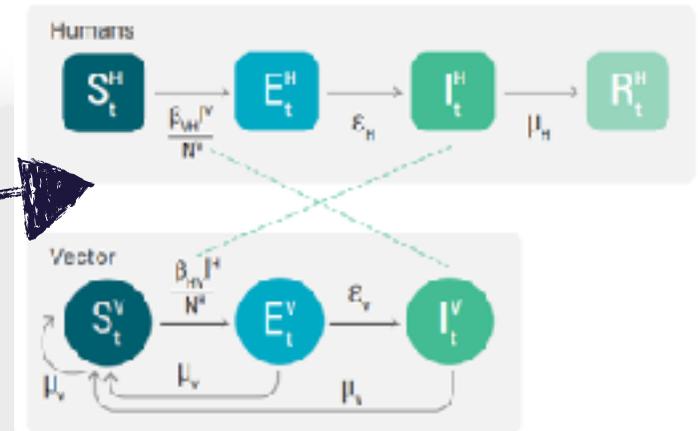
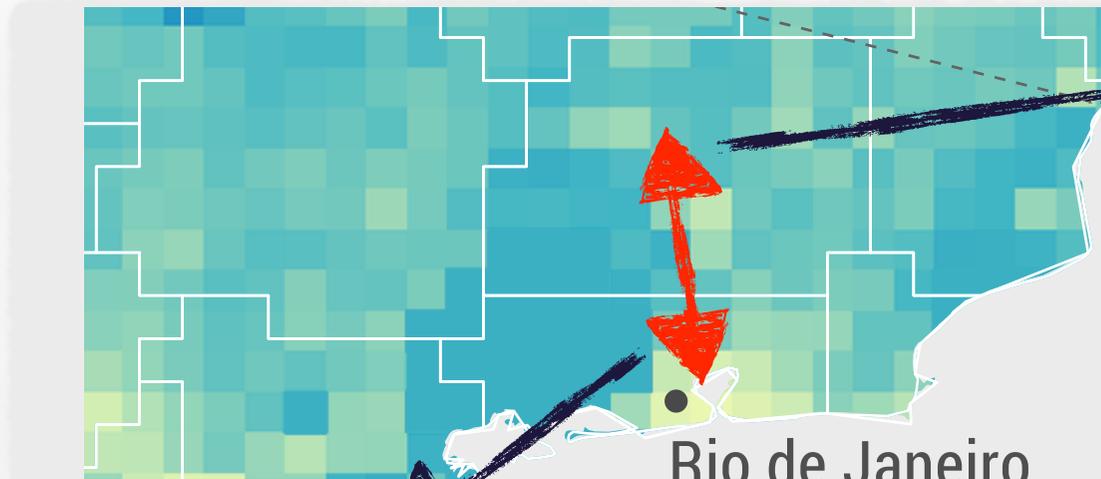
Model resolution

25 km x 25 km within census areas in all the Americas.

A few quantities can be projected up to 1km x 1 km



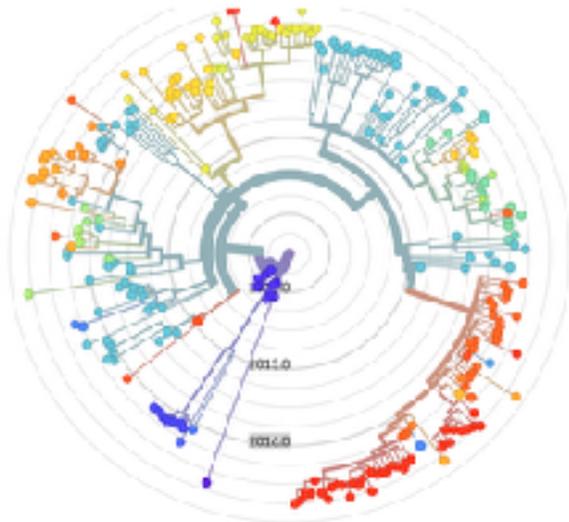
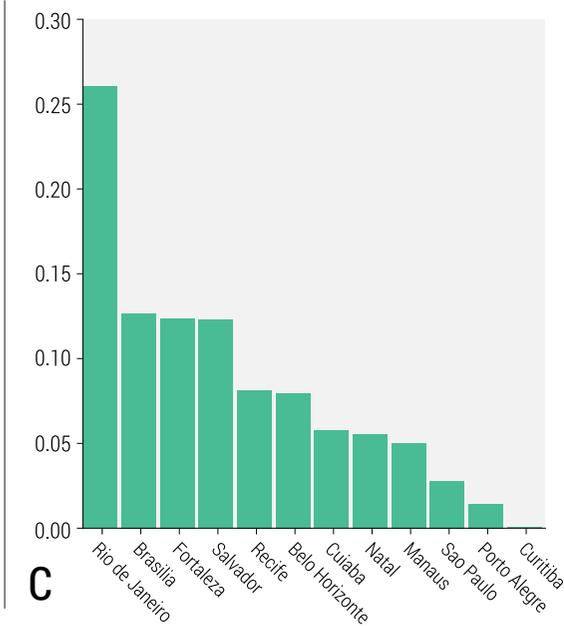
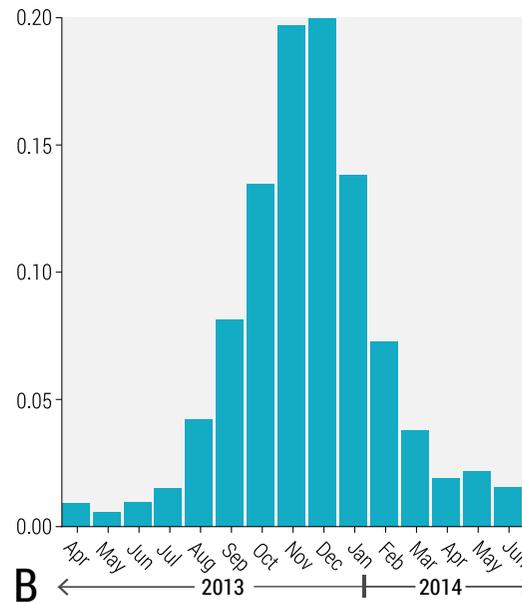
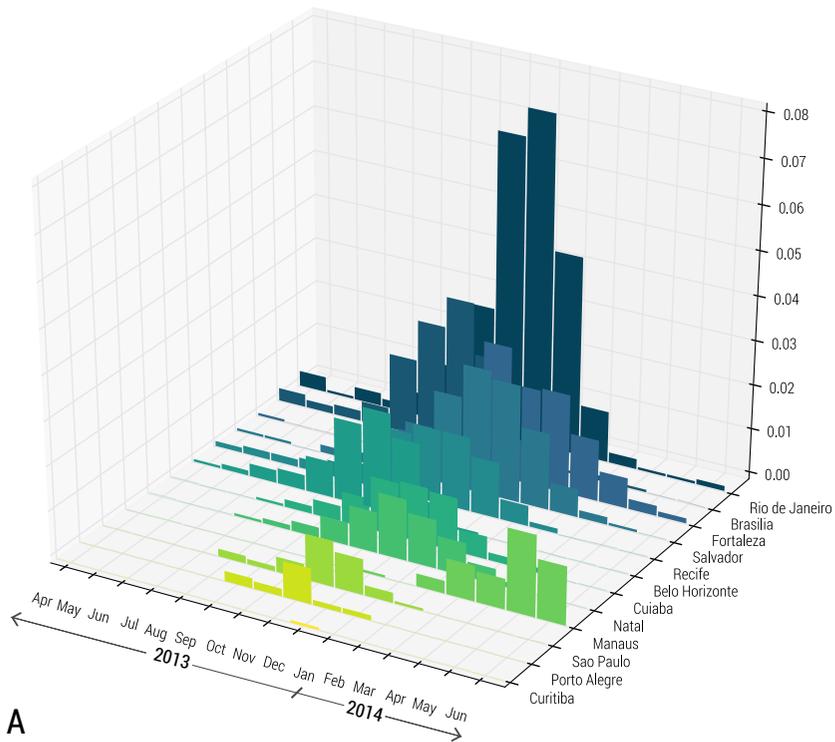
EPIDEMIC DYNAMICS



$$\begin{aligned}
 S_{t+1}^H &= S_t^H - Bin(S_t^H, \lambda_t^H) \\
 E_{t+1}^H &= E_t^H + Bin(S_t^H, \lambda_t^H) - Bin(E_t^H, c_H) \\
 I_{t+1}^H &= I_t^H + Bin(F_{t,t}^H, \epsilon_H) - Bin(I_t^H, \mu_H) \\
 R_{t+1}^H &= R_t^H + Bin(I_t^H, \mu_H)
 \end{aligned}$$

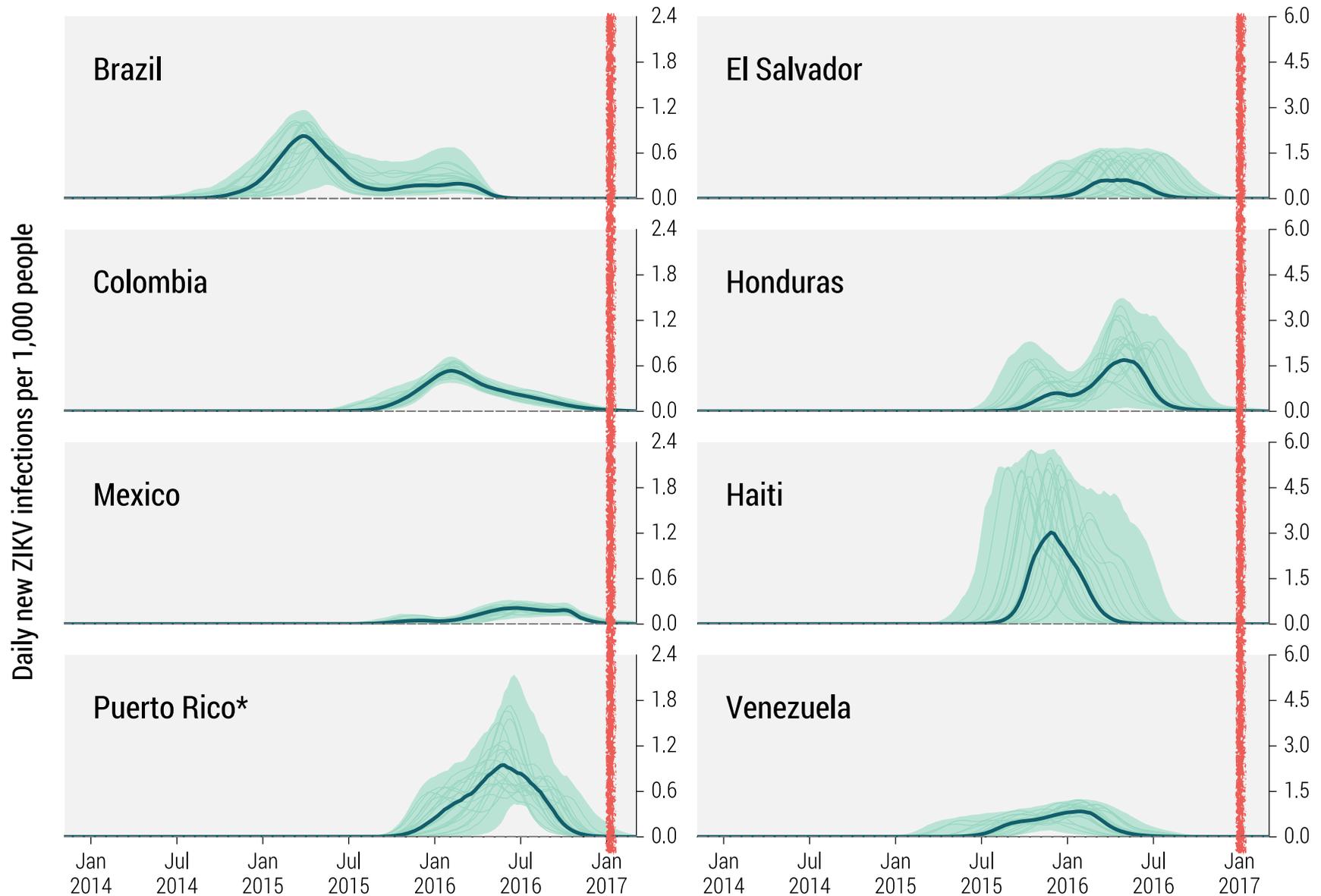
$$\begin{aligned}
 S_{t+1}^V &= S_t^V - Bin(S_t^V, \lambda_t^V) + Bin(I_t^V, \mu_V) + Bin(E_t^V, \mu_V) \\
 E_{t+1}^V &= E_t^V - Bin(E_t^V, \mu_V) - Bin(E_t^V, \epsilon_V) + Bin(S_t^V, \lambda_t^V) \\
 I_{t+1}^V &= I_t^V + Bin(E_t^V, \epsilon_V) - Bin(I_t^V, \mu_V)
 \end{aligned}$$

Epidemiological explanations: Monte-Carlo analysis of ZIKAV introduction in the Americas



Model provides posterior distributions for the place and date of introduction in Brazil in good agreement with Phylogenetic analysis

Epidemic declining in 2017



Number of infections

Attack Rate and Microcephaly Cases, Model Projection (Median with 95%CI)

Date of Projection	Attack Rate %	Date of Projection	Microcephaly Cases* (0.95 % first trimester risk)	Microcephaly Cases* (2.19 % first trimester risk)	Microcephaly Cases* (4.52 % first trimester risk)
By Feb. 01, 2016	4.33 2.73 - 6.83	By Feb. 01, 2016	0 0 - 4	0 0 - 10	1 0 - 20
By Feb. 28, 2017	11.9 10.5 - 13.5	By Dec. 10, 2017	219 194 - 248	504 447 - 572	1041 922 - 1180

*Estimates for microcephaly cases are based on risk calculation provided in Michael A. Johansson et al. New England Journal of Medicine (2016); Cauchemez et al. The Lancet 387, 2125-2132 (2016). Estimates may change as new and/or more precise risk multipliers will be available.

**The incidence profile is the median incidence calculated each week from the stochastic ensemble output of the model and it may not be representative of specific epidemic realizations.

Colombia

Attack Rate and Microcephaly Cases, Model Projection (Median with 95%CI)

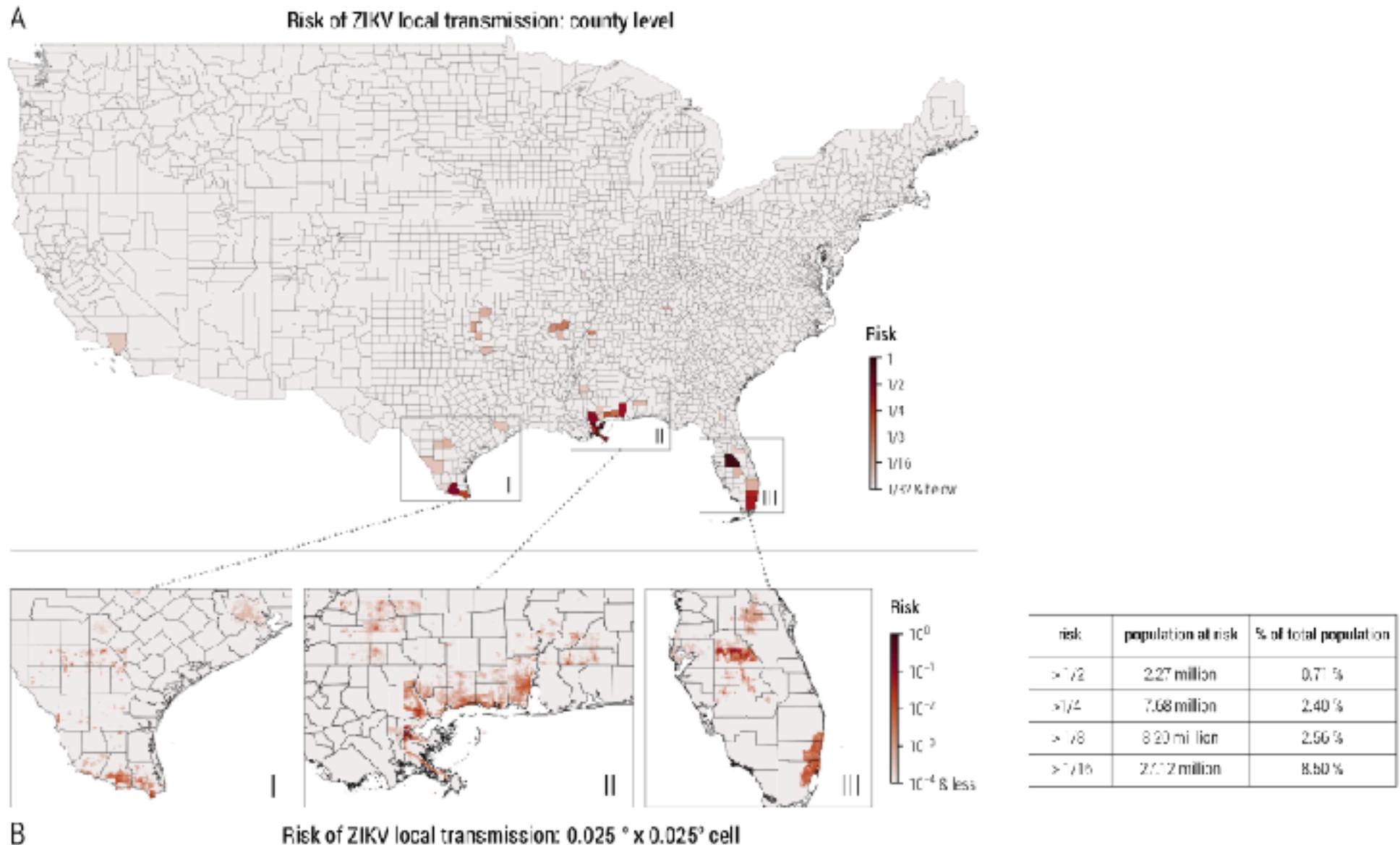
Date of Projection	Attack Rate %	Date of Projection	Microcephaly Cases* (0.95 % first trimester risk)	Microcephaly Cases* (2.19 % first trimester risk)	Microcephaly Cases* (4.52 % first trimester risk)
By Feb. 01, 2016	1.93 0.02 - 6.83	By Feb. 01, 2016	0 0 - 0	0 0 - 0	0 0 - 0
By Feb. 28, 2017	19.6 13.3 - 27.6	By Dec. 10, 2017	19 13 - 26	43 29 - 60	88 60 - 124

*Estimates for microcephaly cases are based on risk calculation provided in Michael A. Johansson et al. New England Journal of Medicine (2016); Cauchemez et al. The Lancet 387, 2125-2132 (2016). Estimates may change as new and/or more precise risk multipliers will be available.

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Puerto Rico

Risk of local transmission in the US



COMPUTATIONAL IMPLEMENTATION

HPC

Large-scale numerical simulations

- About 15 million runs (overall)
- More than 1M instances deployed
- About 125 years of computational time

HPDA

From raw data to actionable summary tables

- Hundred of TB of data analyzed
- Each scenario up to more than 10 TB of raw data

STILL A MAJOR HURDLE FOR REAL-TIME
GLOBAL MODELING

WHERE IS THE COMPUTATIONAL PROBLEM

Statistical ensemble

Each large-scale numerical simulation is a single instance of a stochastic process.

- Thousands (or more) of runs to extract statistical patterns
- The finer the scale, the larger the required number of realizations
- Real time requirement

Model calibration

Calibration on real data is executed via MCL or MCMC scanning high dimensional parameters space

- Millions (or more) of single realizations exploring the level of match with real data.
- Real time requirement

Real-time analysis of large-scale/highly detailed models requires increasing amount of computational resources.

Supercomputing culture must be more and more a key component of the pandemic modeling and analysis research.

Interdisciplinary endeavor not possible without the contribution of the HPC