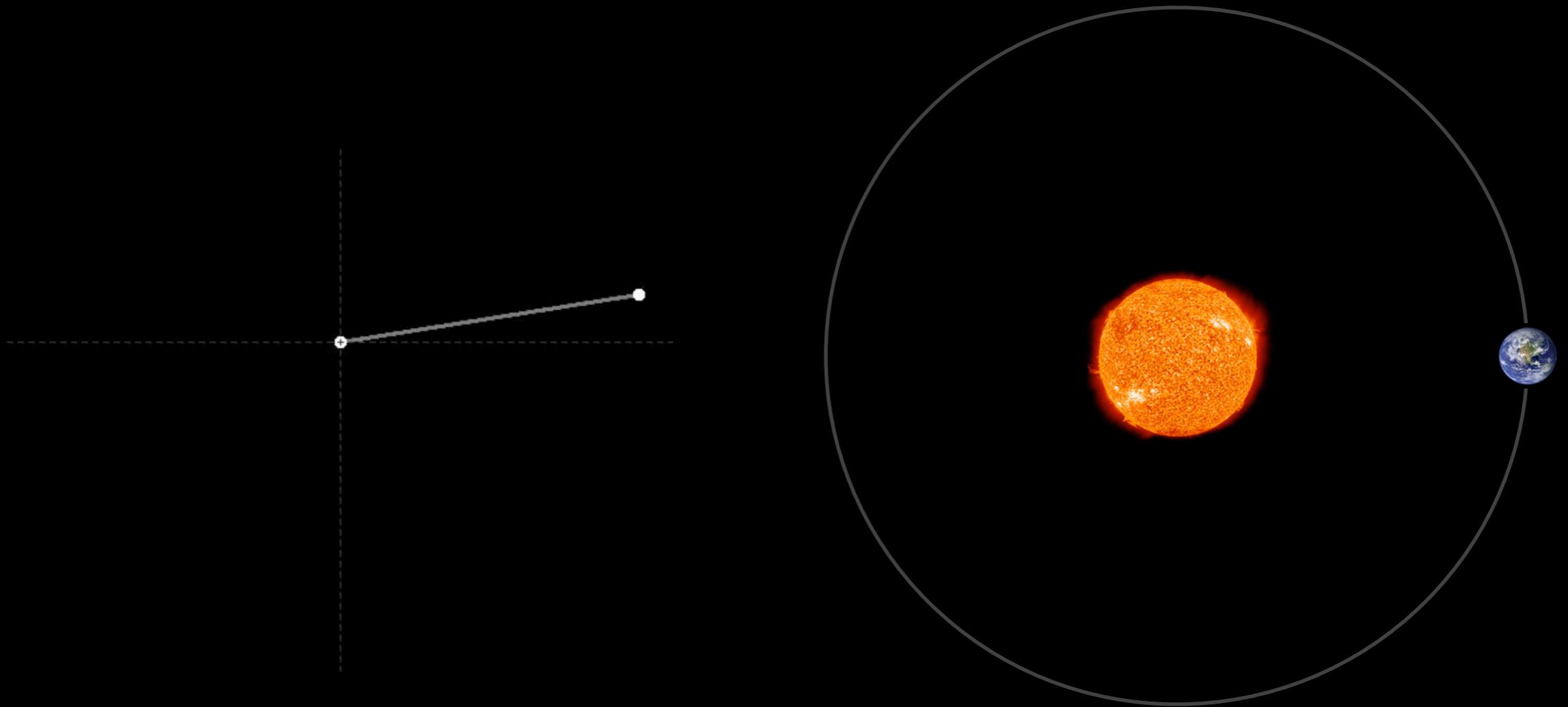
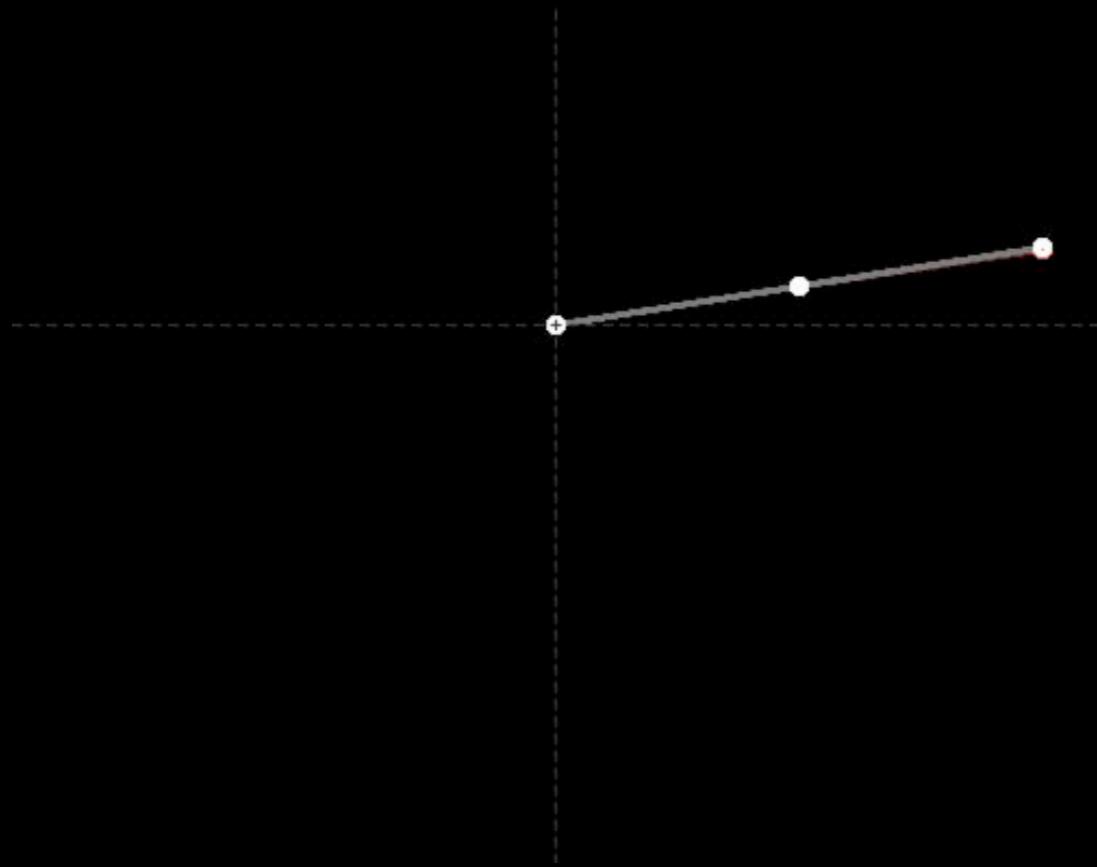


predictability

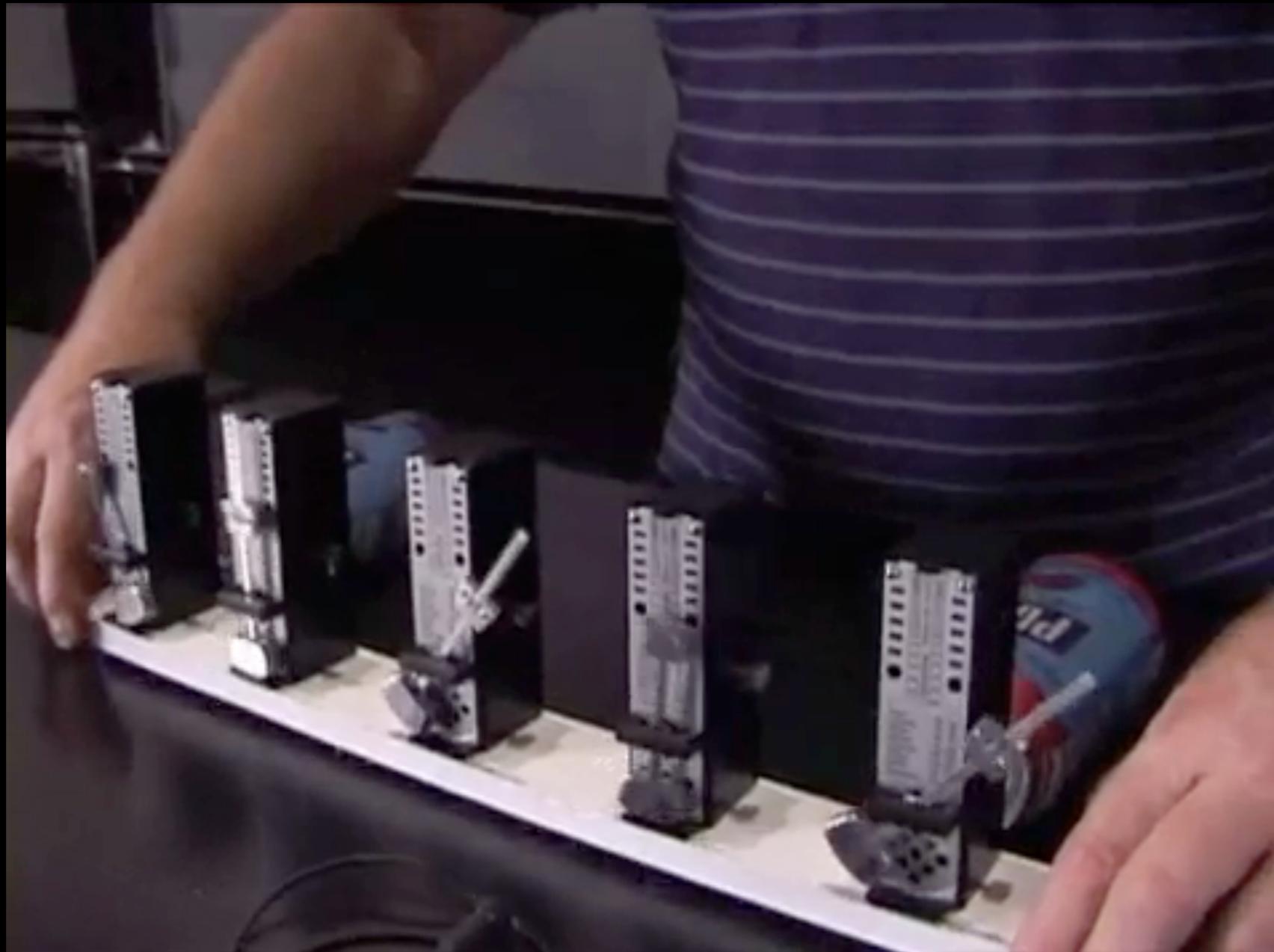


predictability



predictability = knowing initial conditions

predictability

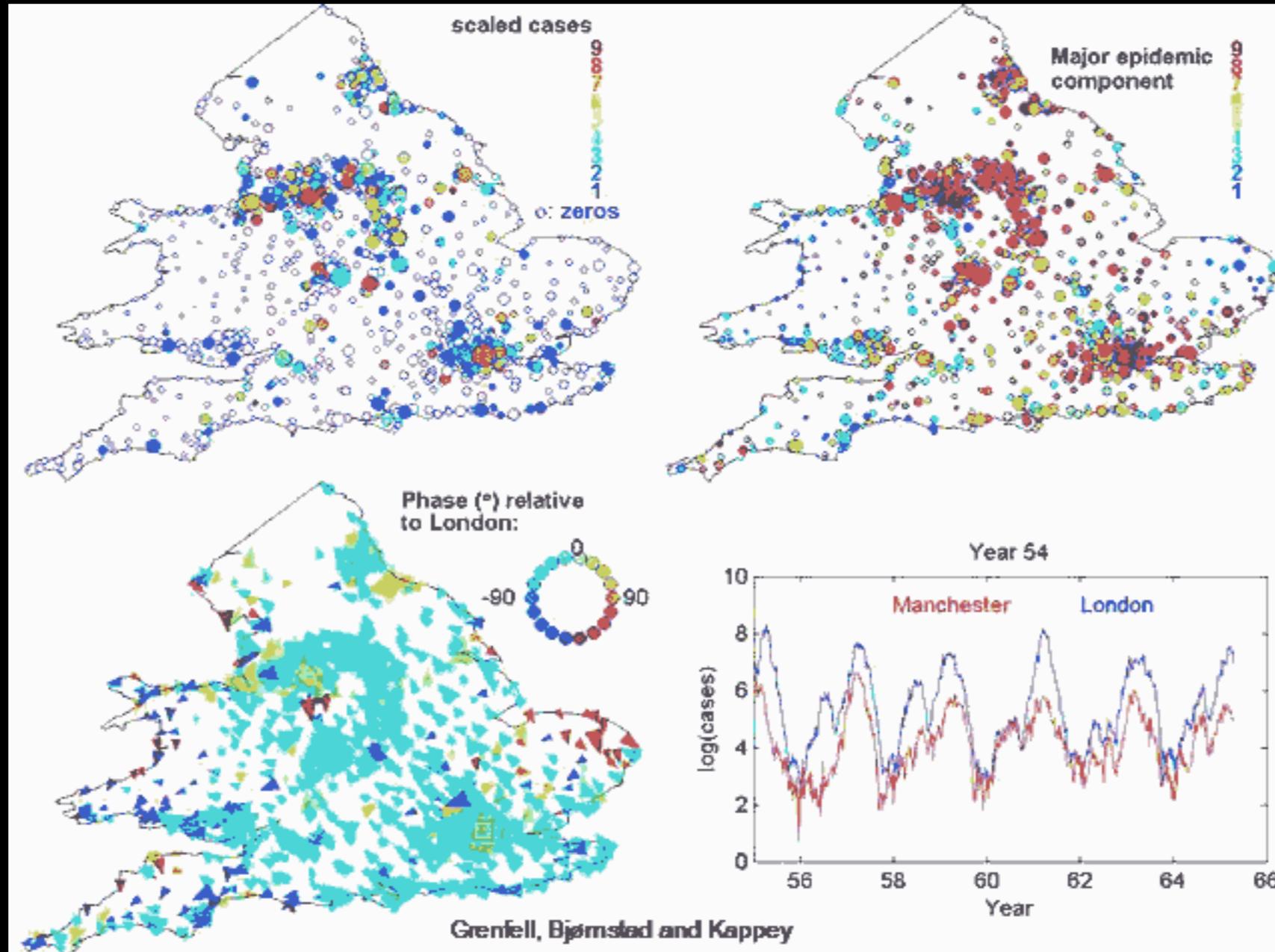


predictability = knowing parametric conditions

the millennium bridge, London



traveling waves in measles epidemic



Grenfell, B., et al., Nature 414, 716–723 (2001).

predictability in global disease dynamics

1760

MATHEMATICS IN EPIDEMIOLOGY



DANIEL BERNOULLI
(1700-1782)

- model for smallpox inoculation
- use of differential equations
- published in 1766
- the use of calculus, in particular differential equations was novel

'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.

Mem Math Phy Acad Roy Sci Paris 1766

inoculation debate

pro



Charles Marie de la Condamine
(1701–1774)

con



Pierre Louis Moreau de
Maupertuis (1698–1759)

D'ALEMBERT WAS A MEMBER OF THE ROYAL SOCIETY

KNEW OF BERNOULLI'S WORK SINCE 1760 PRESENTATION BY CONDAMINE

GENERALIZED BERNOULLI'S MODEL AND PUBLISHED IT BEFORE HIM IN 1765

D'ALEMBERT DID NOT LIKE LEONARD EULER, EULER WAS A GOOD FRIEND OF BERNOULLI HOWEVER.

BERNOULLI WAS VERY ANGRY ABOUT D'ALEMBERT'S BEHAVIOUR

What do you say about the enormous platitudes of the great d'Alembert about the probabilities; as I find myself too frequently unjustly treated in his publications, I have decided already some time ago to read nothing anymore which comes from his pen; I have taken this decision on the occasion of a manuscript about inoculation which I sent to the Academy in Paris eight years ago and which was greatly appreciated because of the novelty of the analysis; it was, I dare say, like incorporating a new province into the body of mathematics; it seems that the success of this new analysis caused him pains of the heart;

he has criticized it in a thousand ways all equally ridiculous, and after having it well criticized, he pretends to be the first author of a theory which he did not only hear mentioned. He, however, knew that my manuscript could only appear after some seven or eight years, and he could only have knowledge about it in his capacity as member of the Academy, and in this respect my manuscript should have stayed sacred until it was made public. *Dolus an virtus quis in hoste requirat!*

2013

global mobility and dense populations



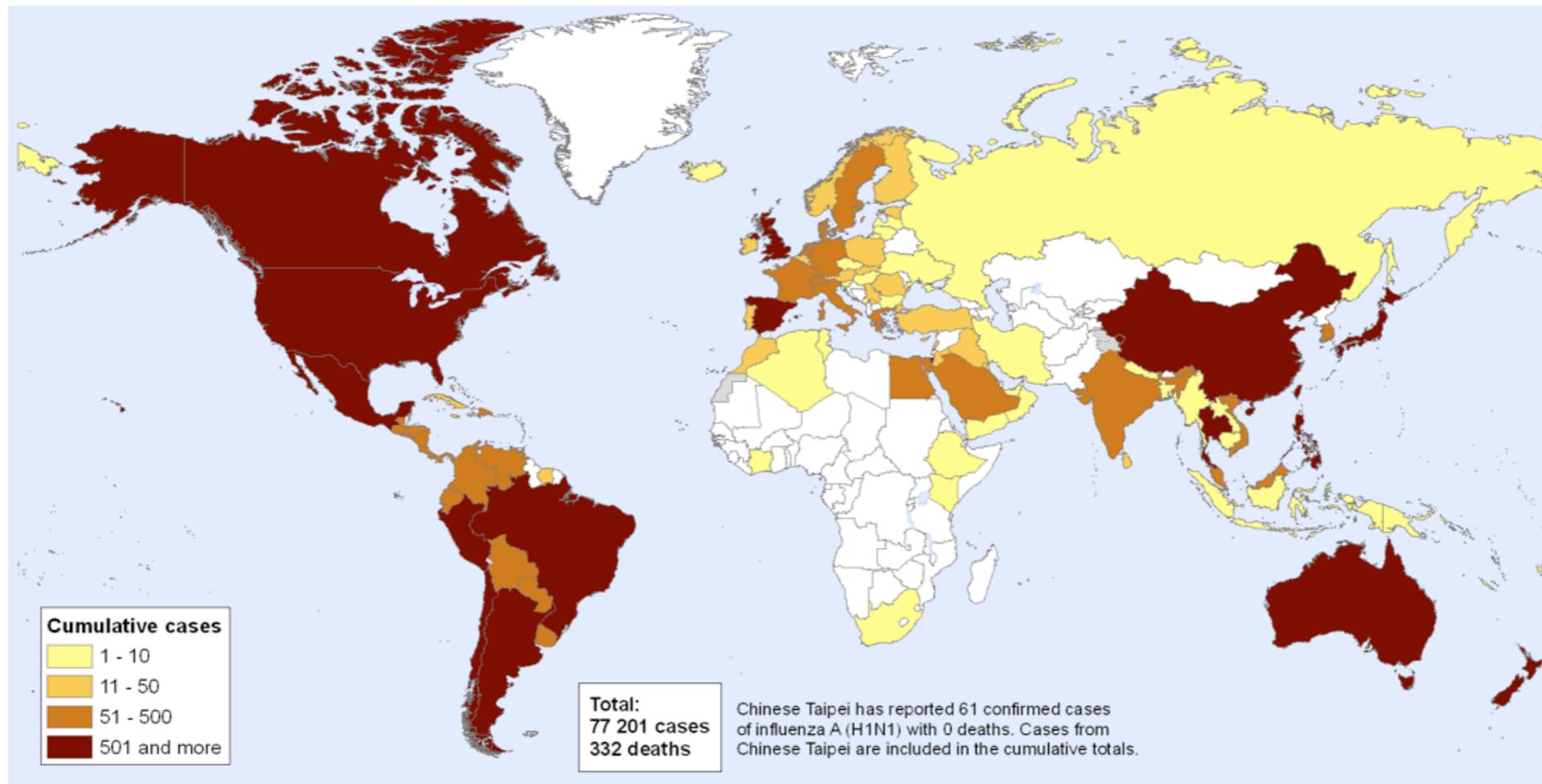
- > 50 % urban population
- population > 7 billion
- 3 billion passengers / yr
- 5 trillion km / yr



H1N1 (swine flue) 2009

Pandemic (H1N1) 2009,
Number of laboratory confirmed cases as reported to WHO

Status as of 01 July 2009
09:00 GMT



The boundaries and names shown and the designations used on this map do not imply the expression of any opinion whatsoever on the part of the World Health Organization concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. Dotted lines on maps represent approximate border lines for which there may not yet be full agreement.

Map produced: 01 July 2009 11:38 GMT

Data Source: World Health Organization
Map Production: Public Health Information
and Geographic Information Systems (GIS)
World Health Organization

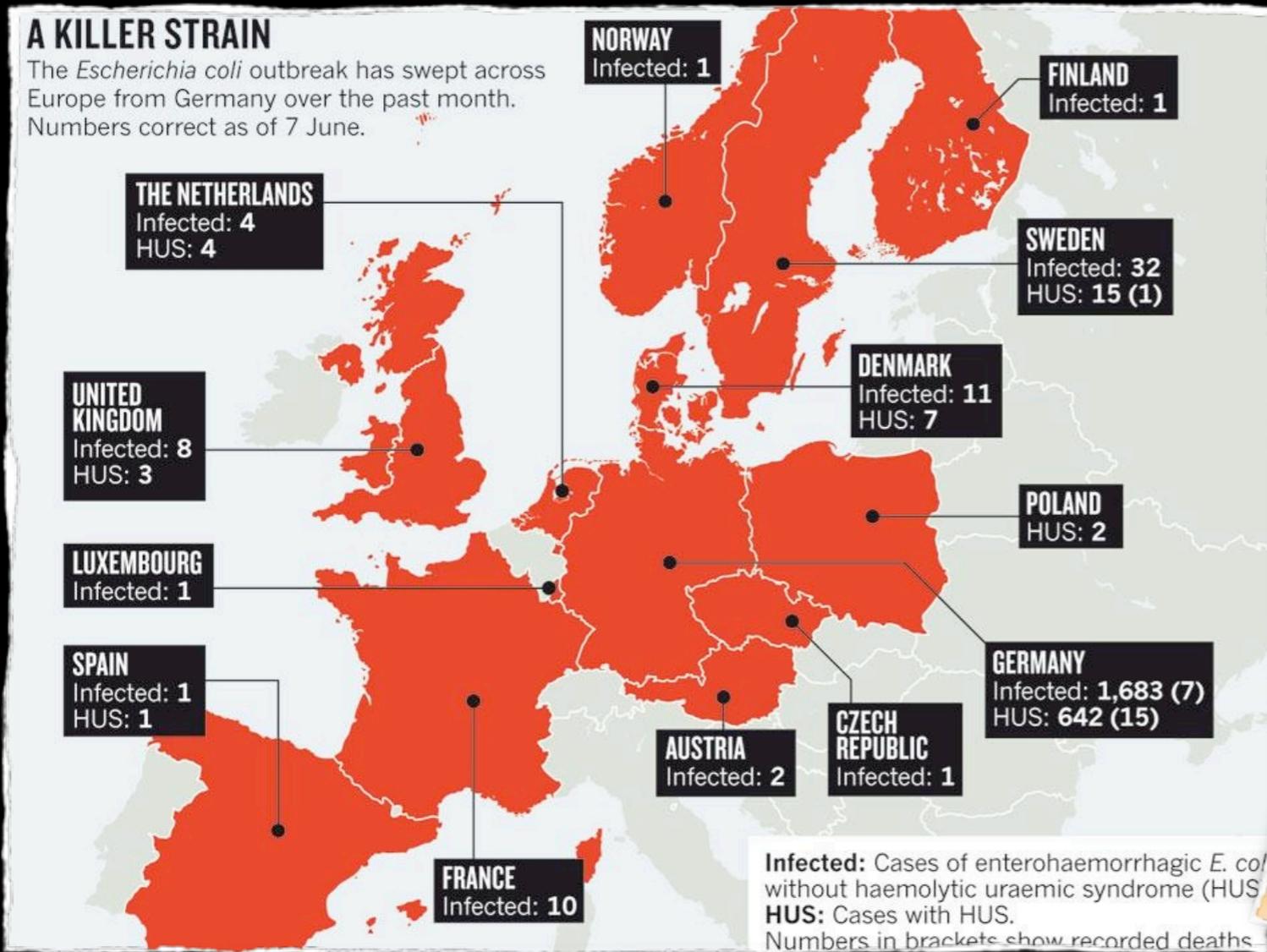


© WHO 2009. All rights reserved



EHEC 2011

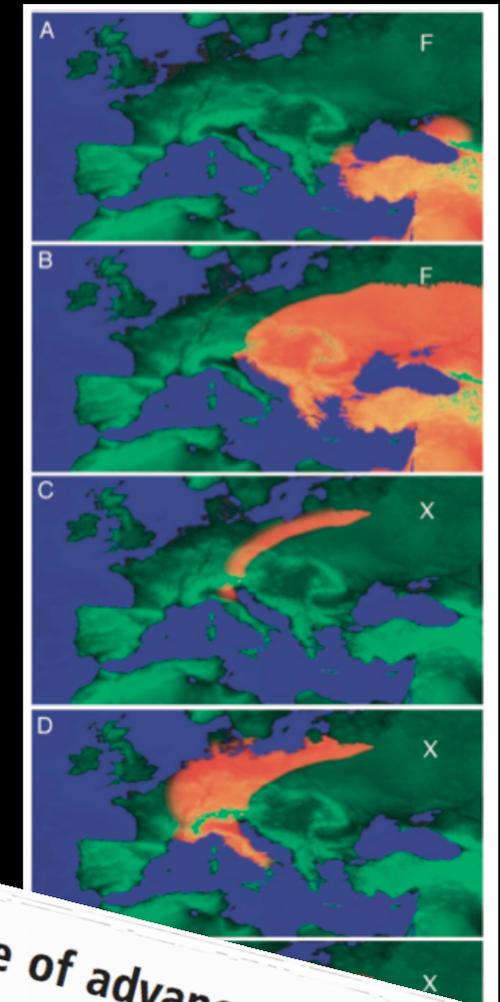
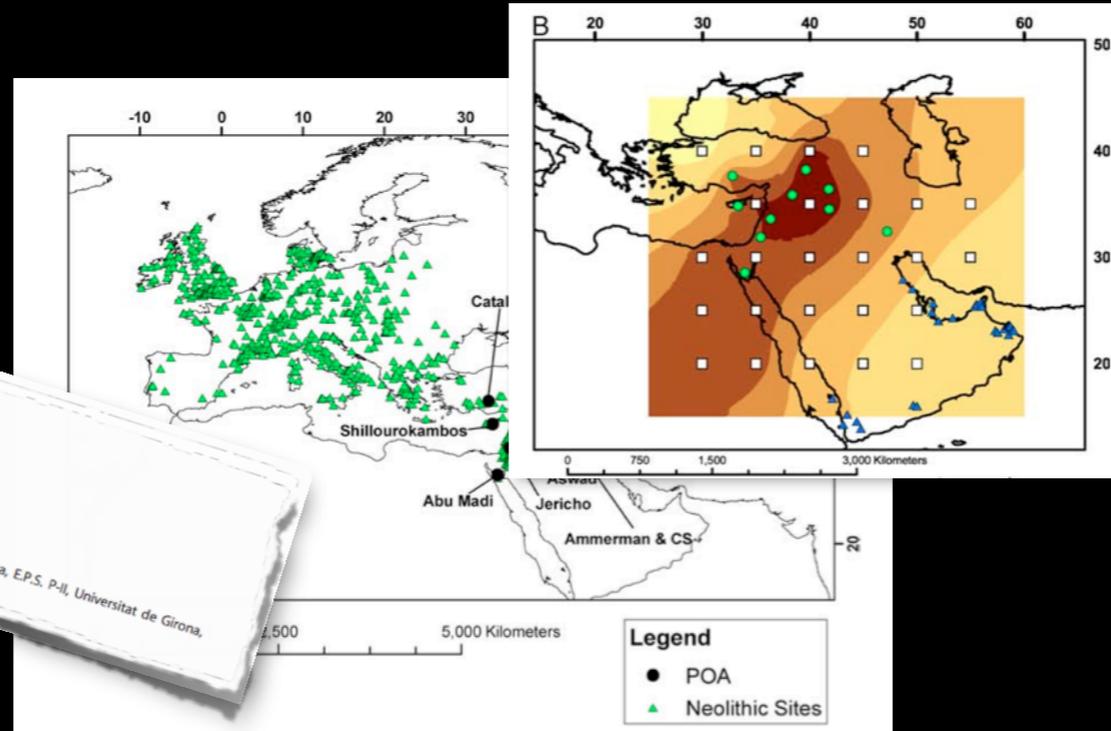
- foodborn disease
- E. Coli
- May - June 2011
- 4000 affected, 60 died



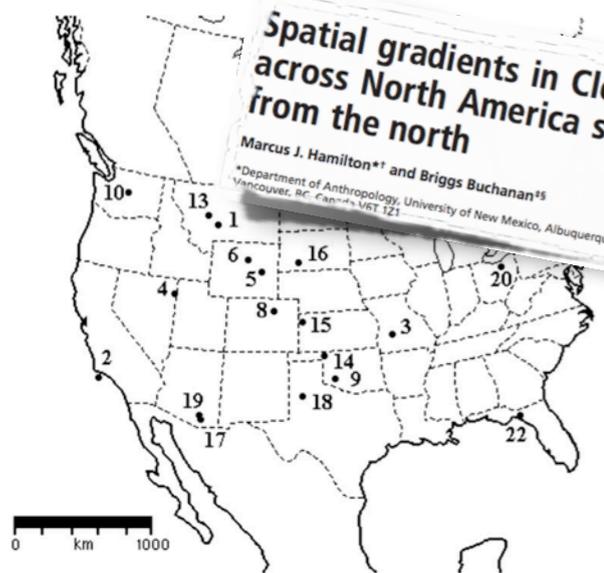
Frank, et al., New England Journal of Medicine (2011).

finding outbreaks

Tracing the Origin and Spread of Agriculture in Europe
 Ron Pinhasi^{1*}, Joaquim Fort², Albert J. Ammerman³
¹ School of Human and Life Sciences, Whitelands College, Roehampton University, London, United Kingdom, ² Departament de Física, E.P.S. P-II, Universitat de Girona, Campus de Montilivi, Catalonia, Spain, ³ Department of Classics, Colgate University, Hamilton, New York, United States of America



Spatial gradients in Clovis-age radiocarbon dates across North America suggest rapid colonization from the north
 Marcus J. Hamilton^{*†} and Briggs Buchanan^{*‡}
^{*}Department of Anthropology, University of New Mexico, Albuquerque, NM 87131; and [†]Department of Anthropology, University of British Columbia, Vancouver, BC, Canada V6T 1Z1



Cultural hitchhiking on the wave of advance of beneficial technologies
 Graeme J. Ackland^{*†}, Markus Signitzer^{*}, Kevin Stratford^{*}, and Morrel H. Cohen^{†‡§}
^{*}School of Physics, University of Edinburgh, Edinburgh EH9 3JZ, United Kingdom; [†]Department of Physics and Astronomy, Rutgers, The State University of New Jersey, Piscataway, NJ 08854; and [‡]Department of Chemistry, Princeton University, Princeton, NJ 08544

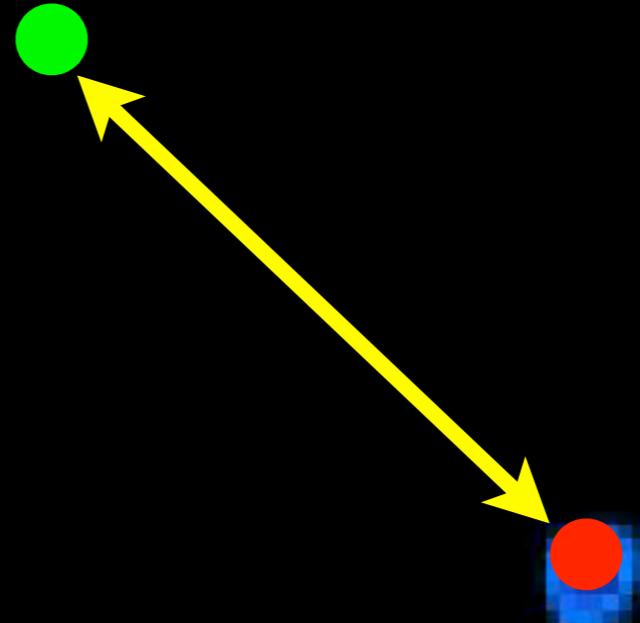
Ackland, G. J., Signitzer, M., Stratford, K. & Cohen, M. H. P PNAS (2007).
 Pinhasi, R., Fort, J. & Ammerman, A., Plos Biol (2005)
 Hamilton, M. J. & Buchanan, B., PNAS, (2007).

questions

Speed?

Arrival?

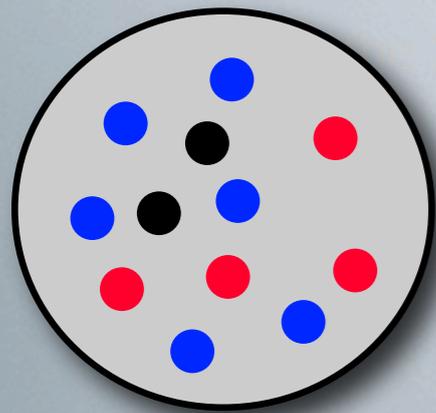
Origin?



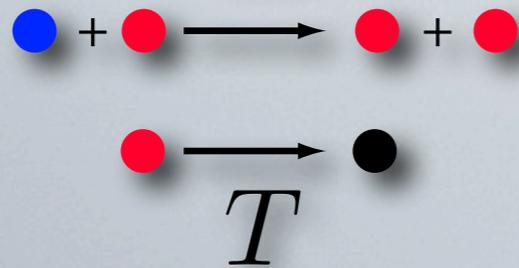
$$T = \frac{D}{V}$$

Noble, Nature (1974) vol. 250 (5469) pp. 726-728
Fort, J. & Méndez, V., Phys Rev Lett (1999).

SIR DYNAMICS



SIR model: α



- infected
- susceptible
- recovered/removed

mean field dynamics:

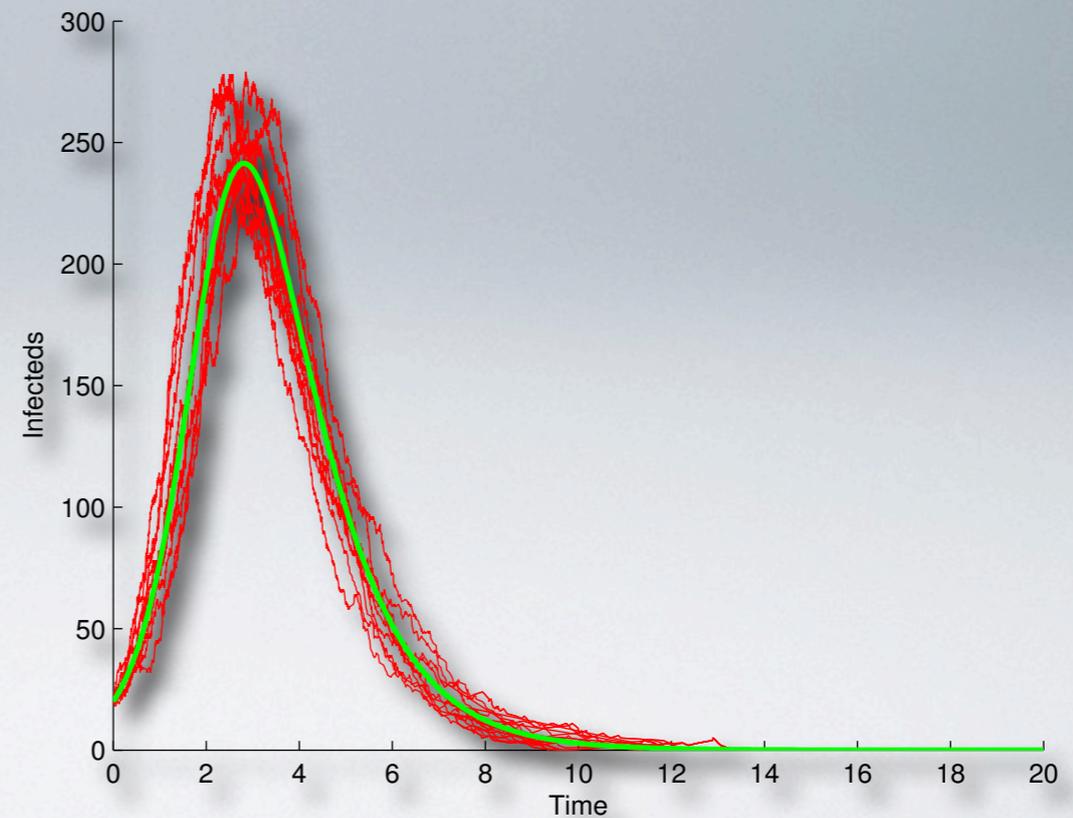
$$\partial_t S = -R_0 I S$$

$$\partial_t I = R_0 I S - I$$

$$R = 1 - S - I$$

The basic reproduction number

$$\alpha \times T = R_0 > 1$$

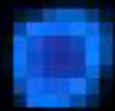


reaction diffusion models

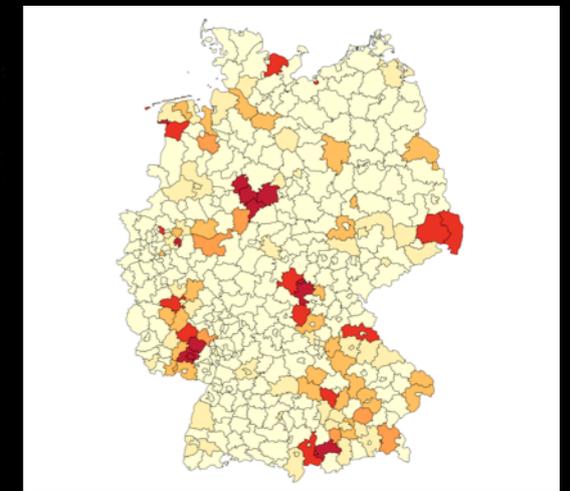
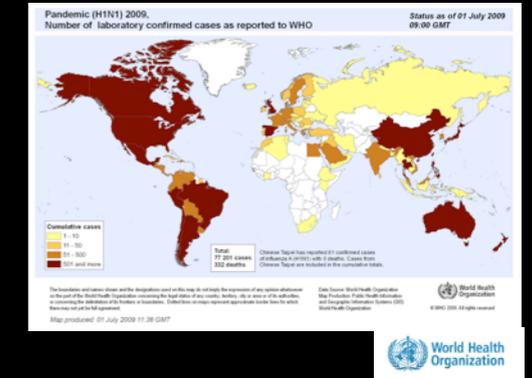
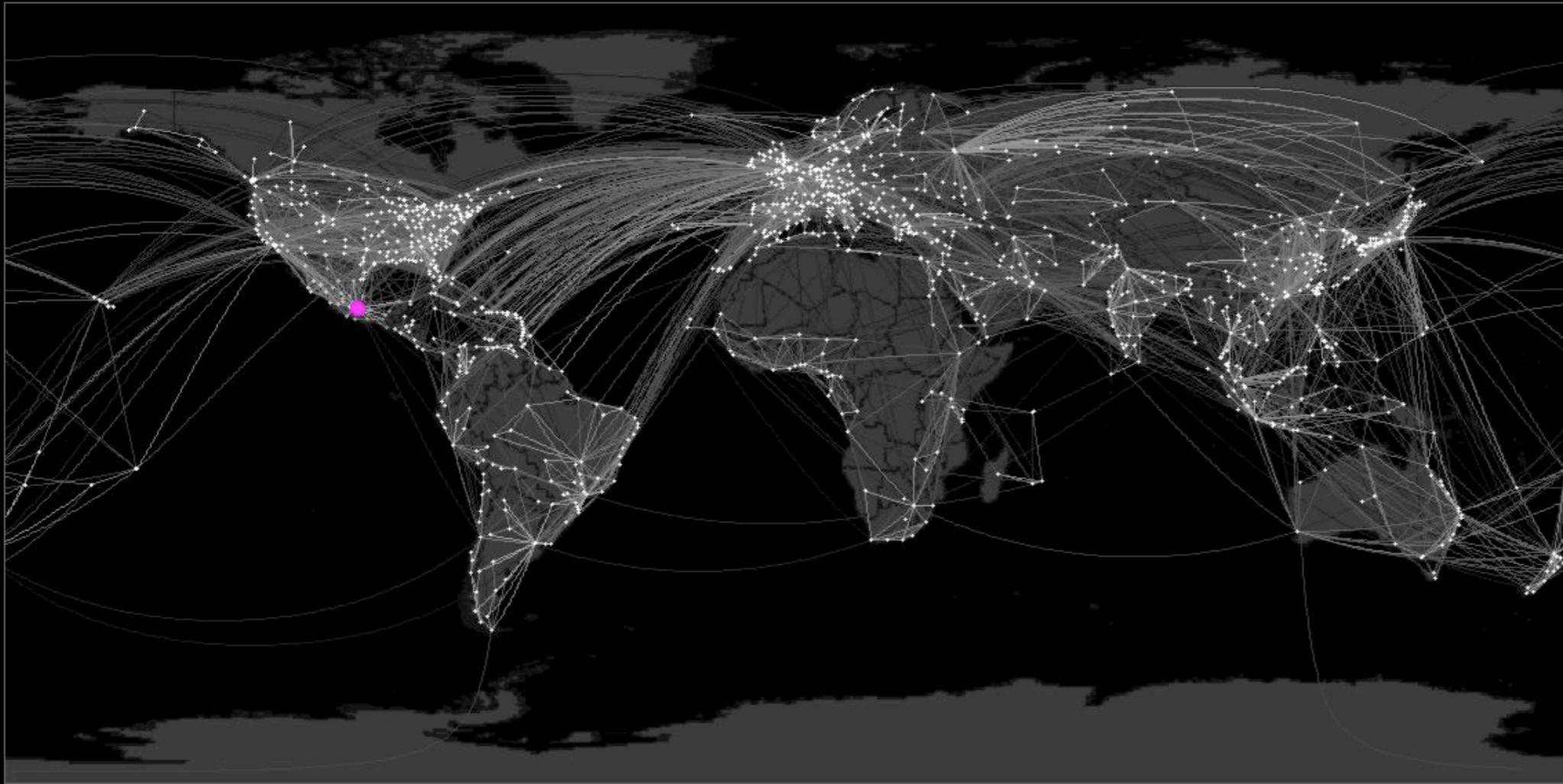
$$\partial_t I = \alpha IS/N - \beta I + D\partial_x^2 I$$

$$\partial_t S = -\alpha IS/N + D\partial_x^2 S$$

$$v \propto \sqrt{(R_0 - 1)D}$$



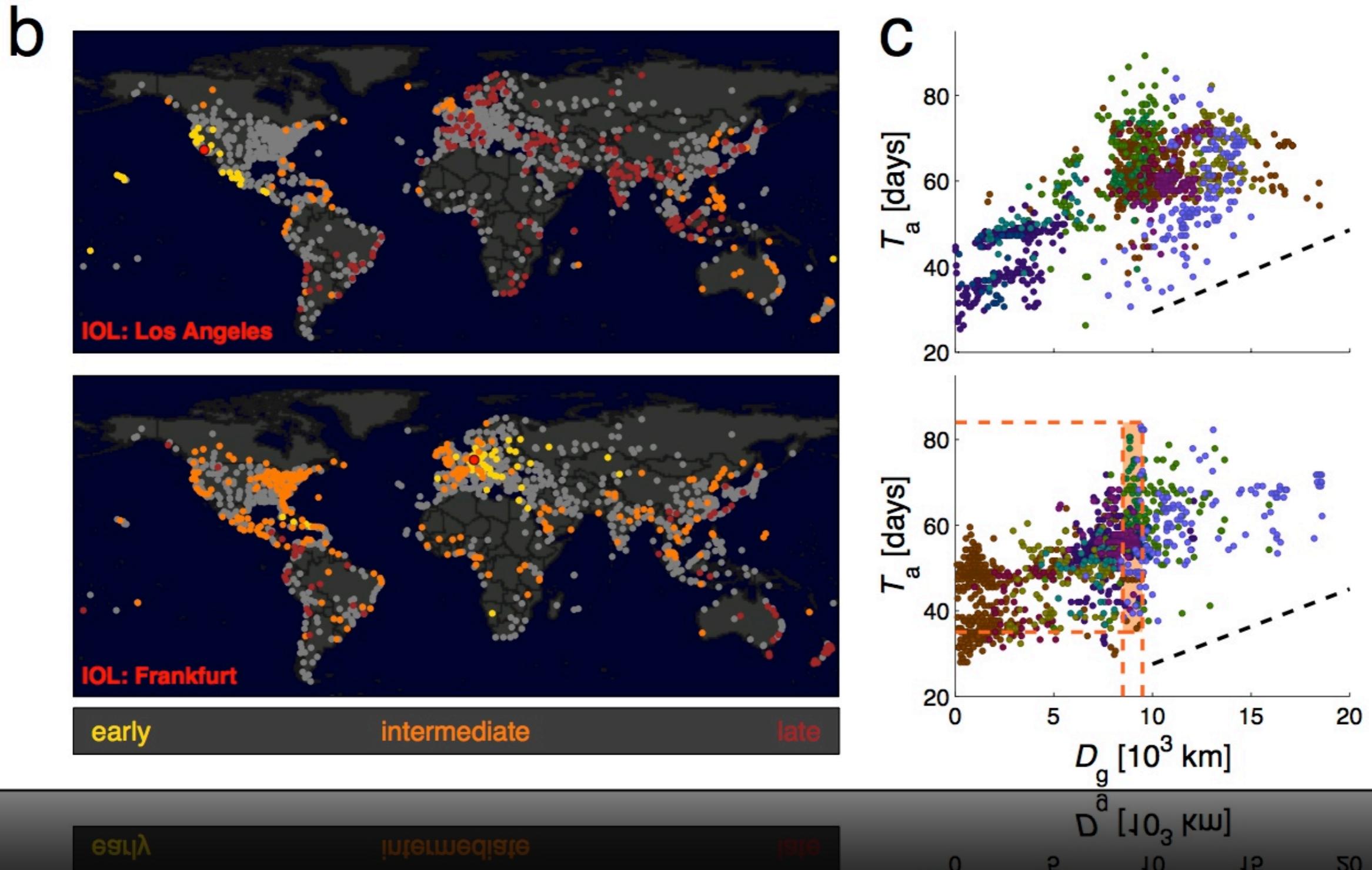


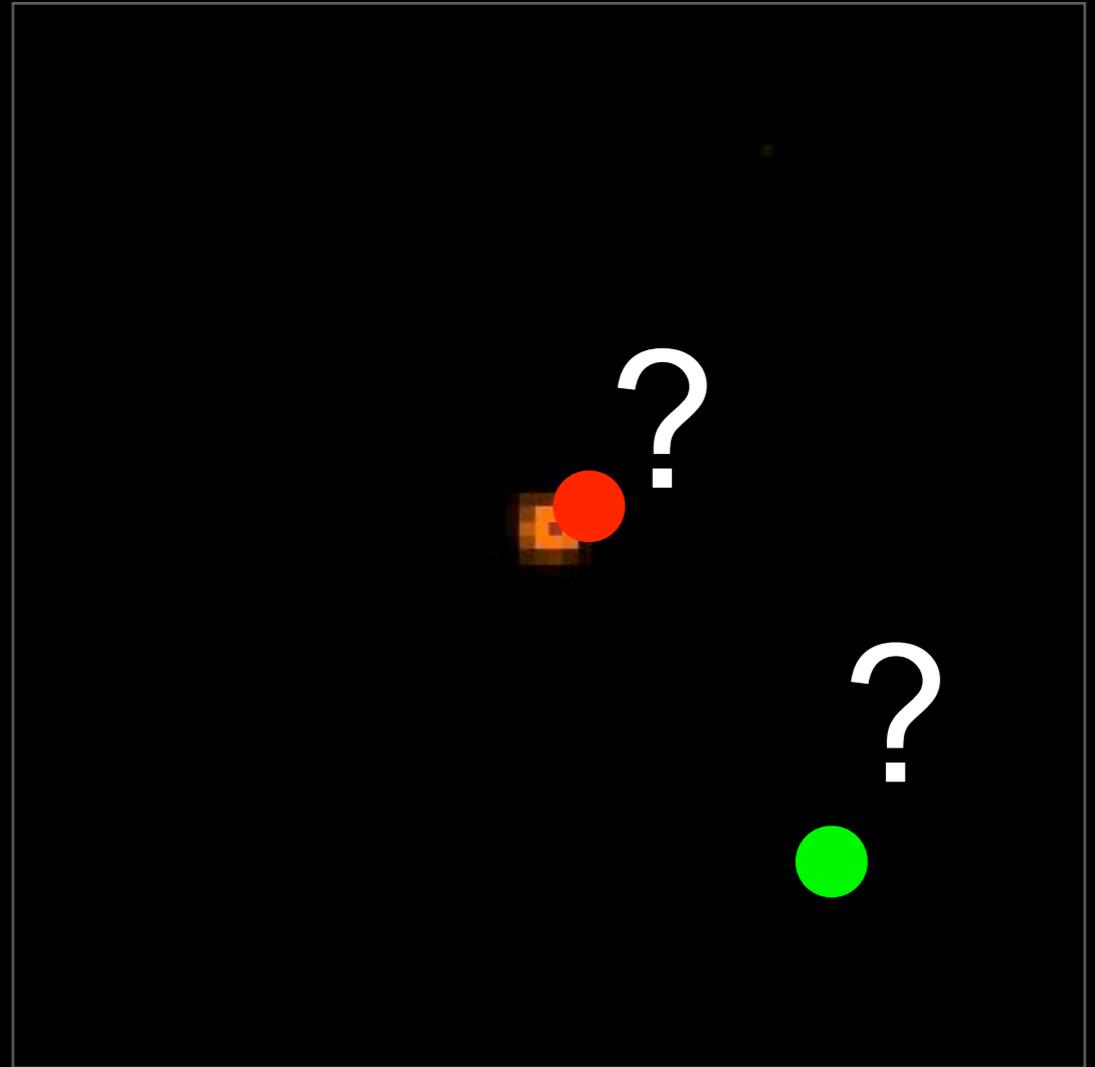
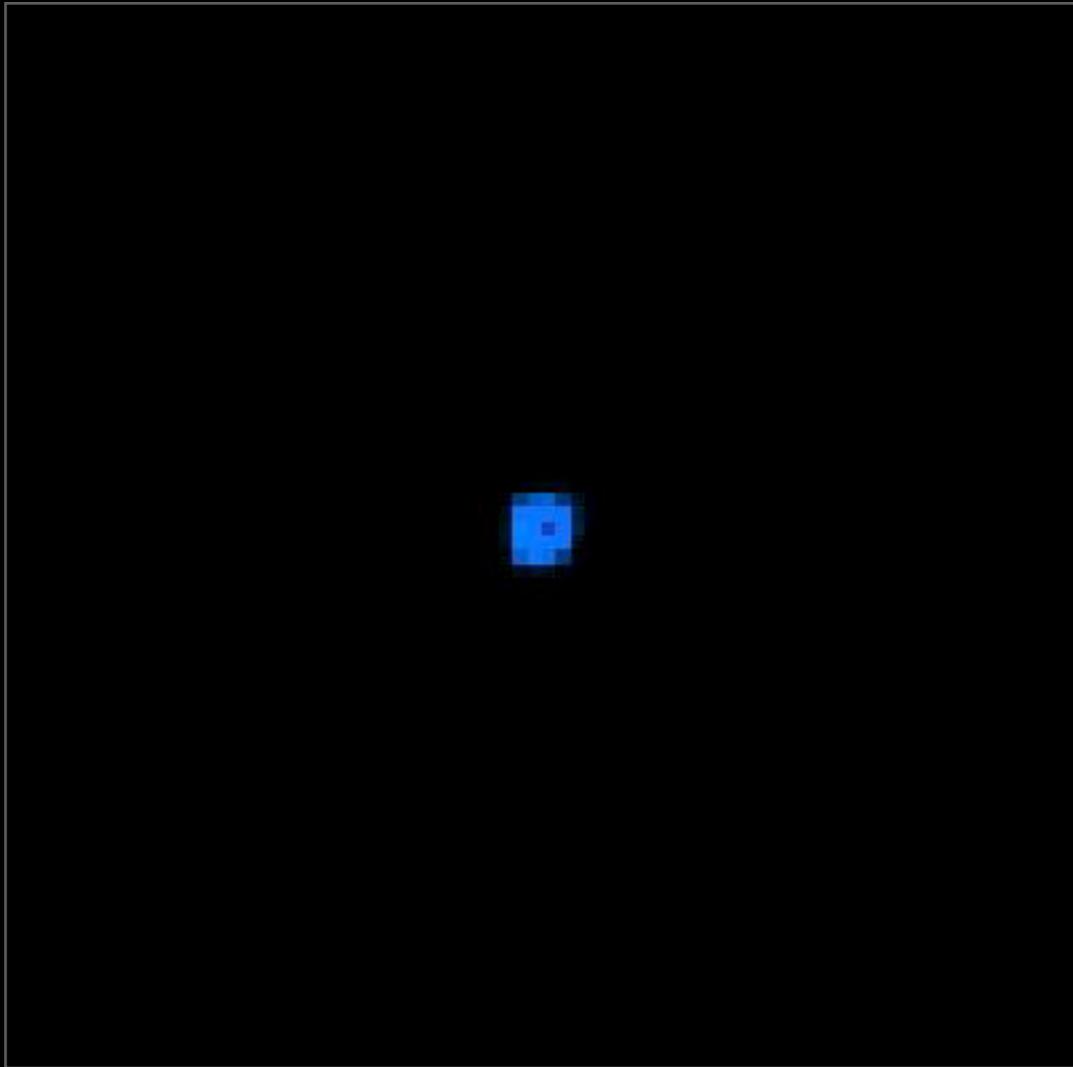


$$\partial_t j_n = \alpha s_n j_n - \beta j_n + \gamma \sum_{m \neq n} P_{nm} (j_m - j_n).$$

$$\partial_t s_n = -\alpha s_n j_n + \gamma \sum_{m \neq n} P_{nm} (s_m - s_n).$$

geographic distance and arrival times in a hypothetical pandemic





quantitative computational models

GLEaMviz



Hufnagel et al. Forecast and control of epidemics in a **globalized** world. PNAS (2004)

Ferguson et al. Strategies for containing an emerging influenza **pandemic** in Southeast Asia. Nature (2005)

Hollingsworth et al. Will **travel restrictions** control the **international spread** of pandemic influenza?. Nat Med (2006)

Colizza et al. The role of the **airline transportation** network in the prediction and predictability of global epidemics. PNAS (2006)

Ferguson et al. Strategies for mitigating an influenza **pandemic**. Nature (2006)

Colizza et al. Modeling the **worldwide spread** of pandemic influenza: Baseline case and containment interventions. Plos Med (2007)

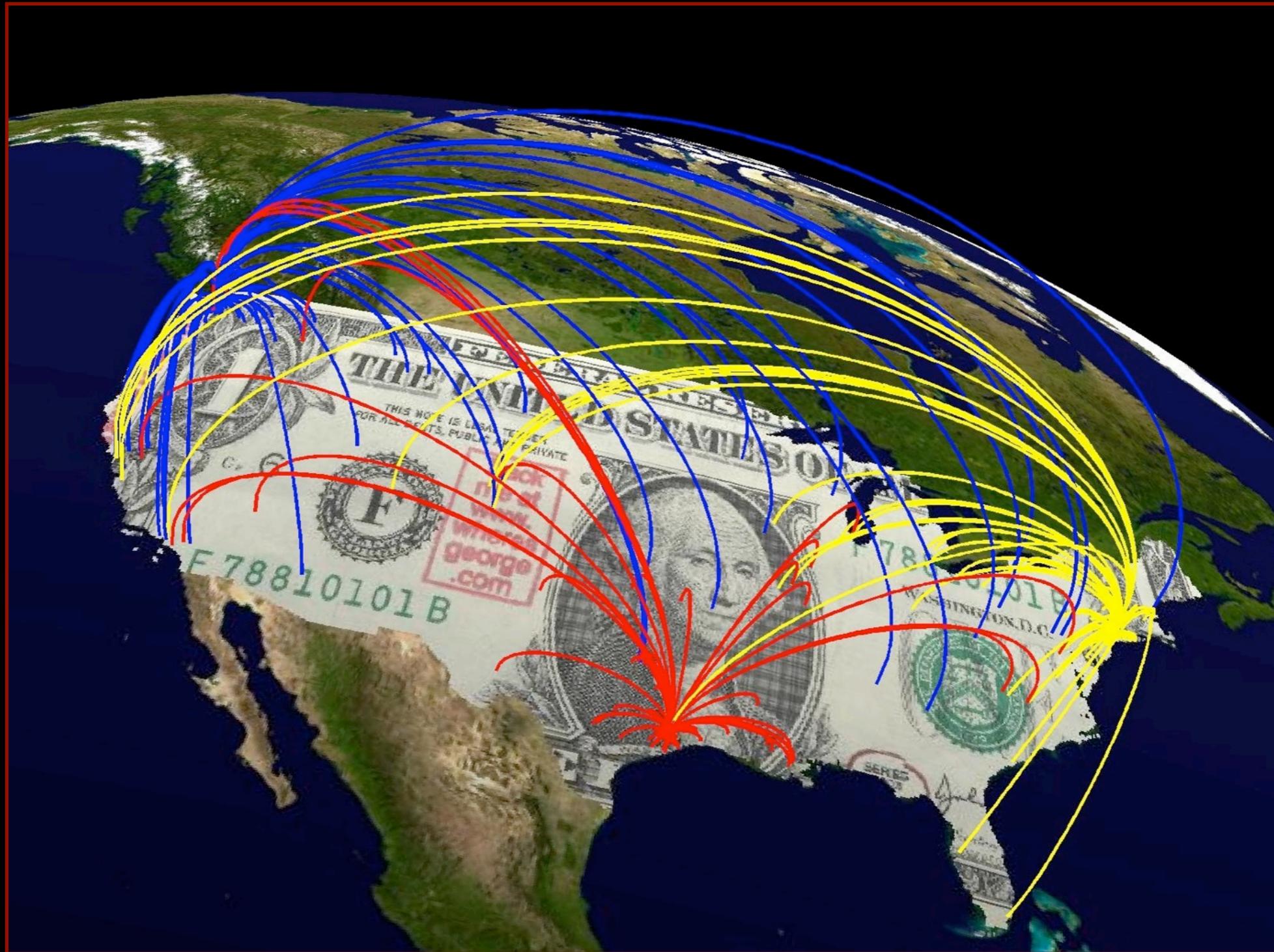
Colizza et al. Reaction-diffusion processes and metapopulation models in heterogeneous networks. Nat Phys (2007)

Colizza et al. Invasion threshold in heterogeneous metapopulation networks. Phys Rev Lett (2007) vol. 99 (14)

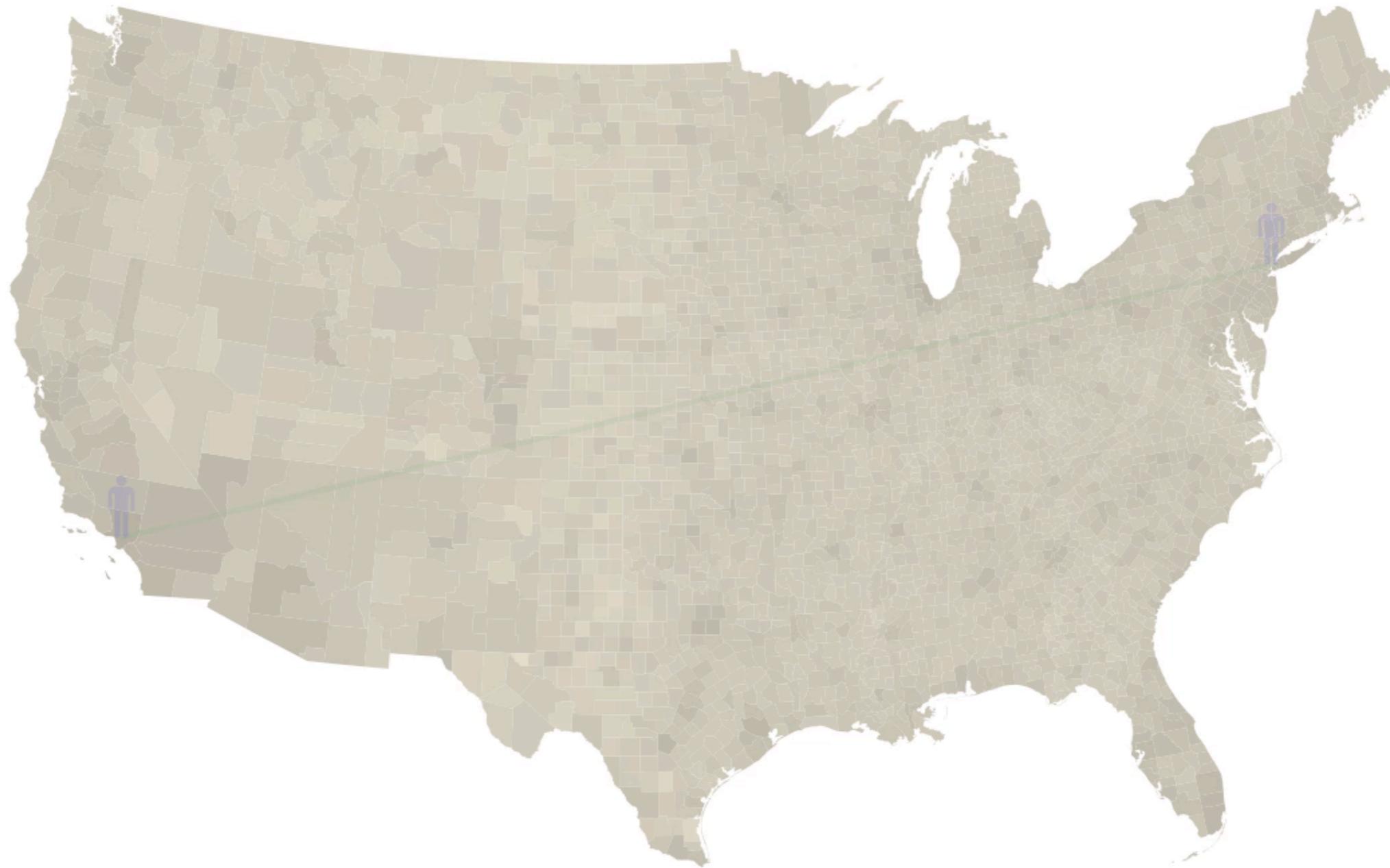
Riley. **Large-scale spatial-transmission** models of infectious disease. Science (2007)

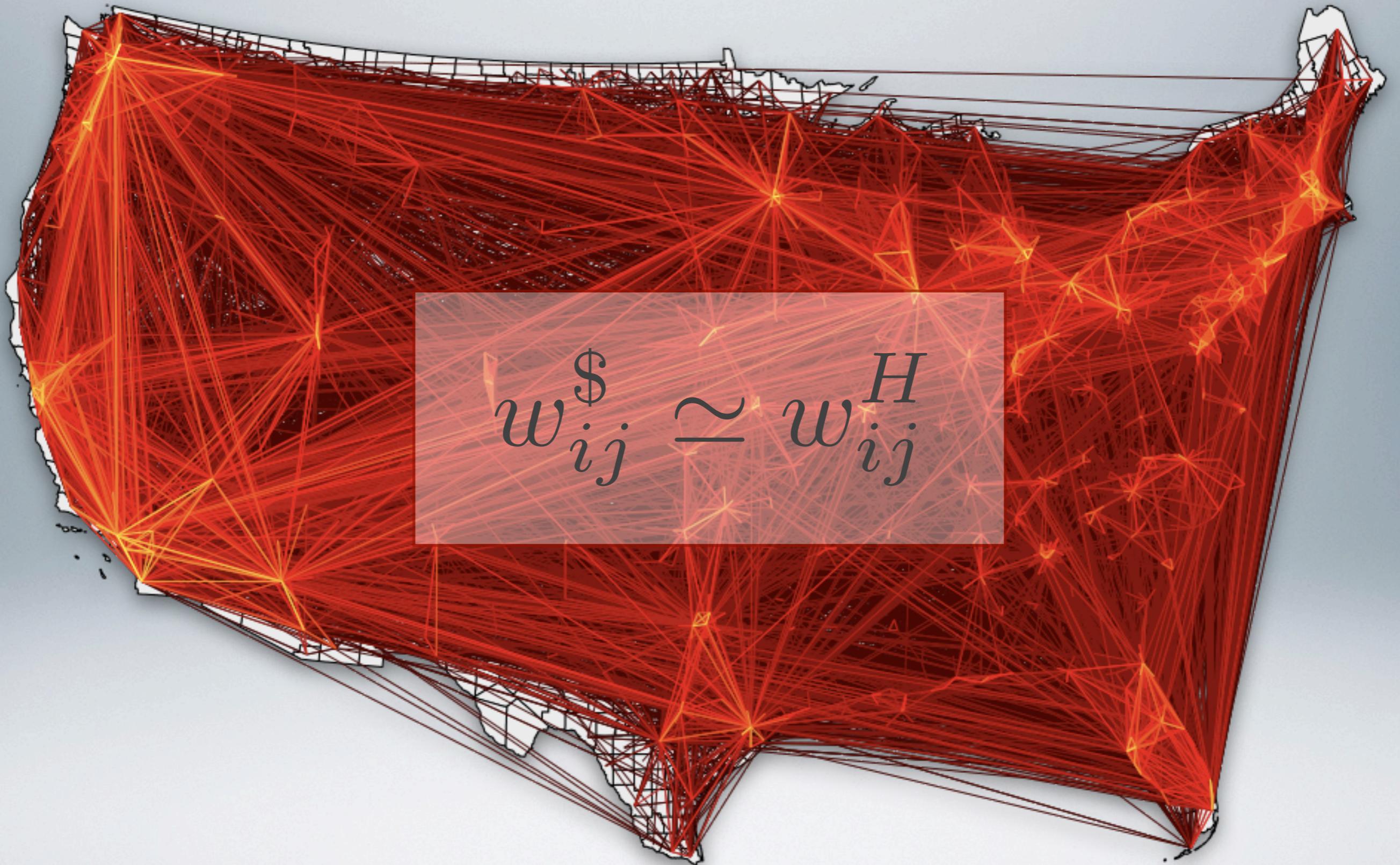
Colizza et al. Epidemic modeling in metapopulation systems with heterogeneous coupling pattern: Theory and simulations. J Theor Biol (2008)

measuring human mobility



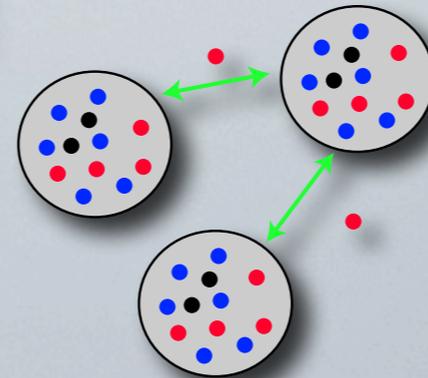
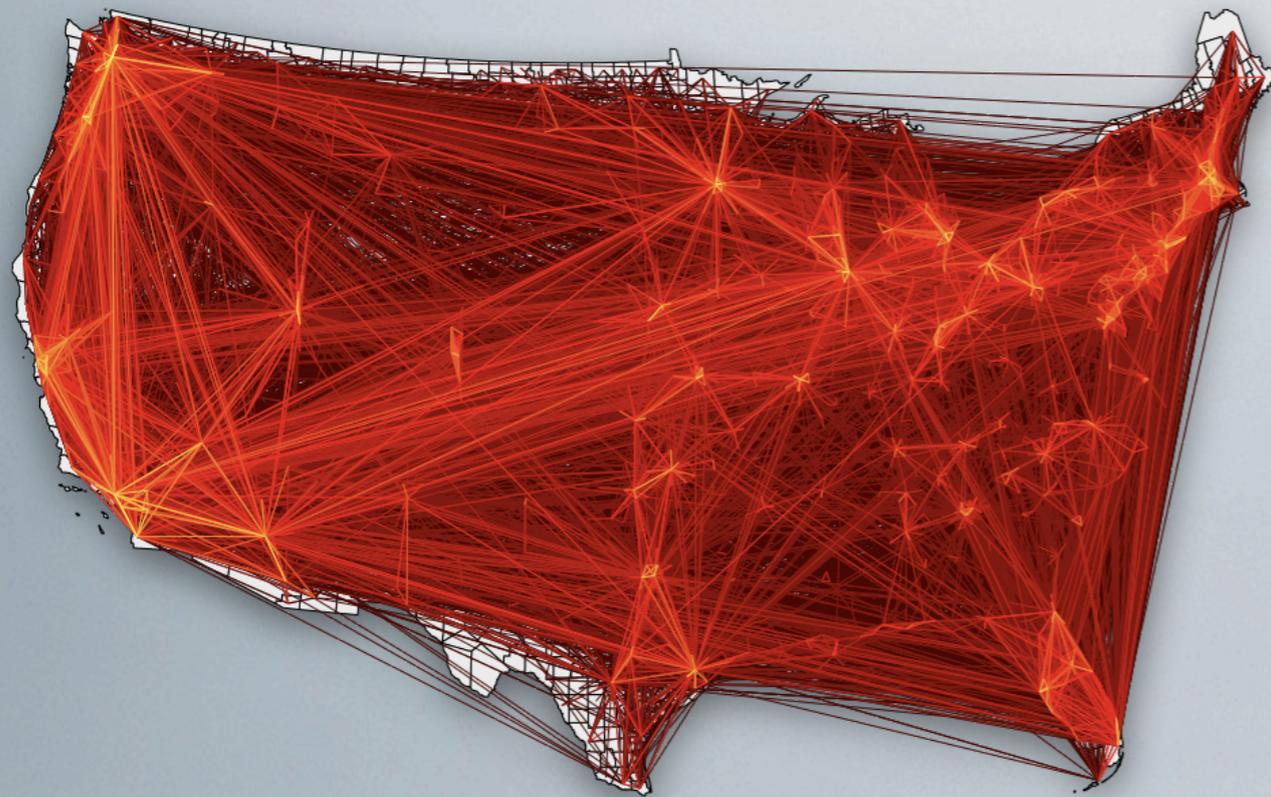
THE FLUX OF DOLLARS



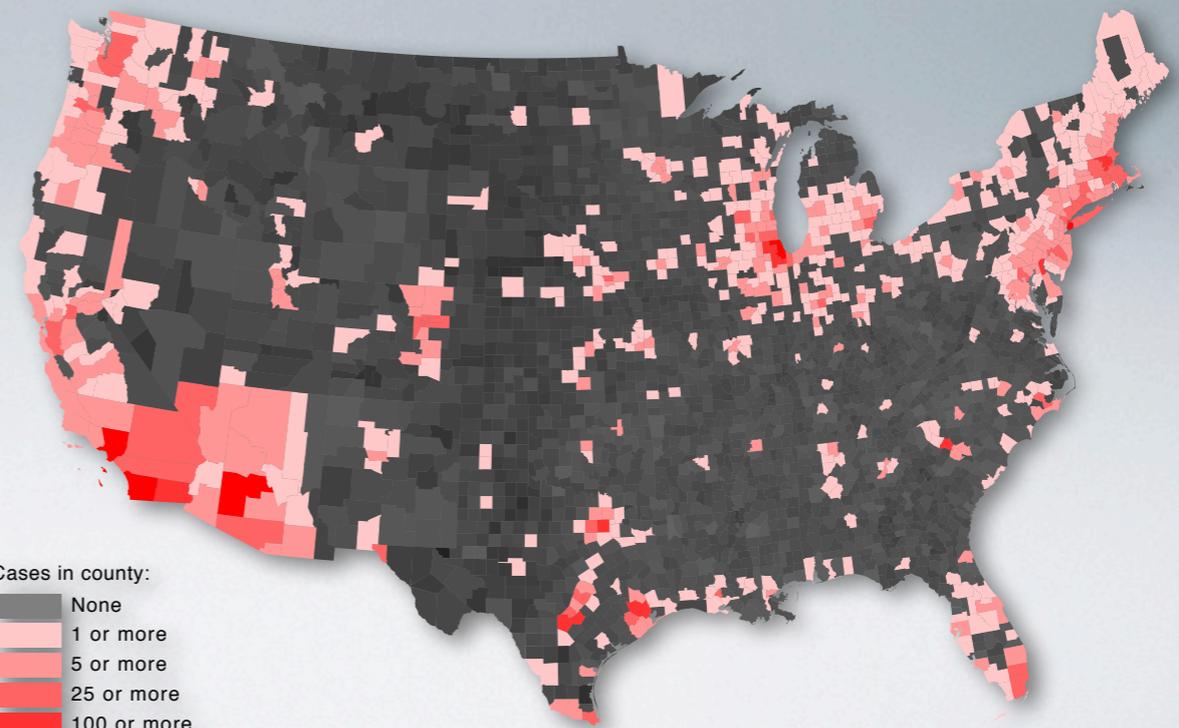
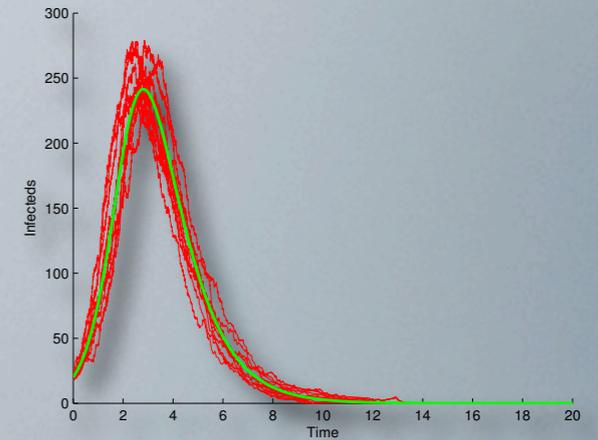


$$w_{ij}^{\$} \approx w_{ij}^H$$

A MODEL FOR THE SPREAD OF H1N1 IN THE US



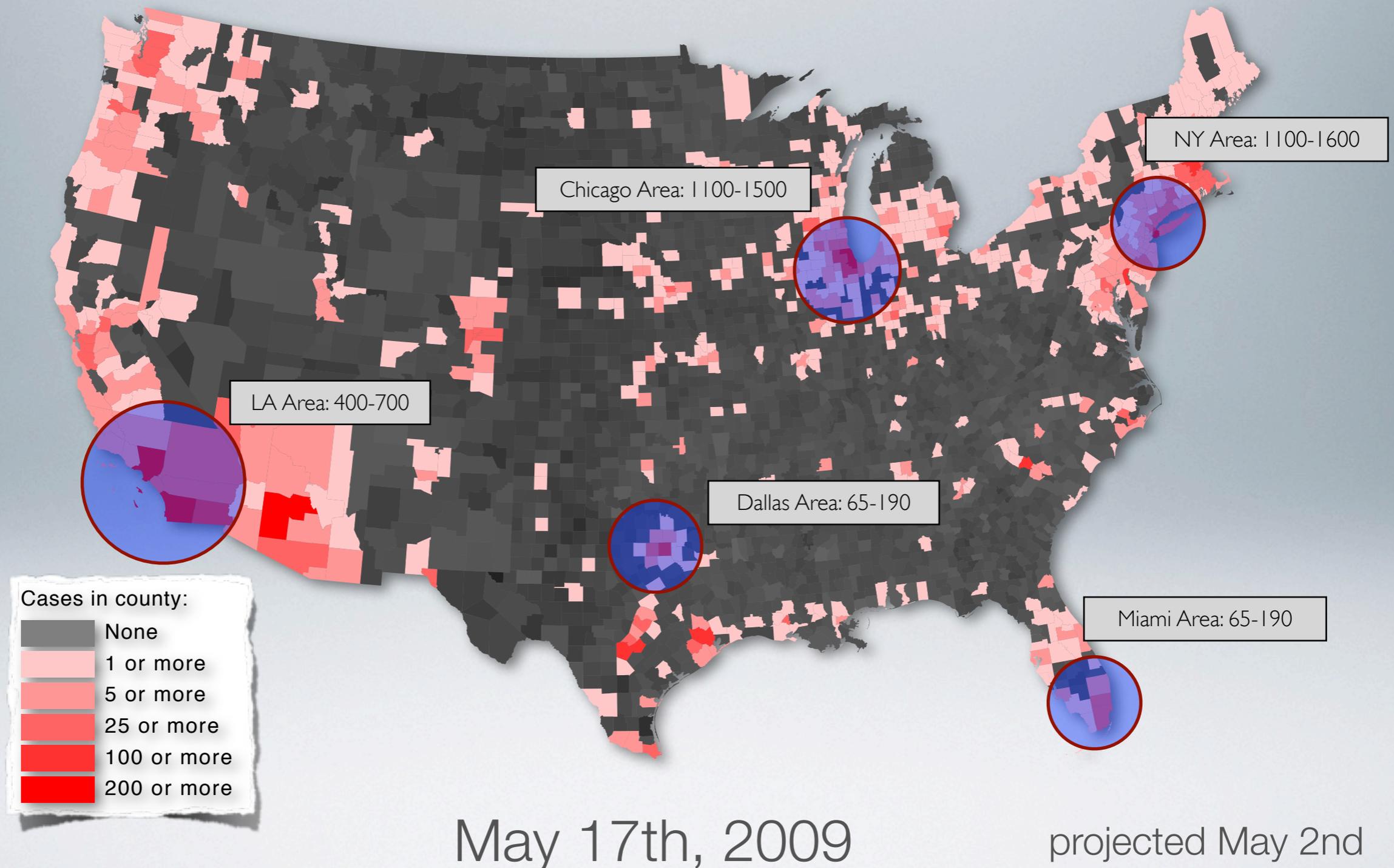
+



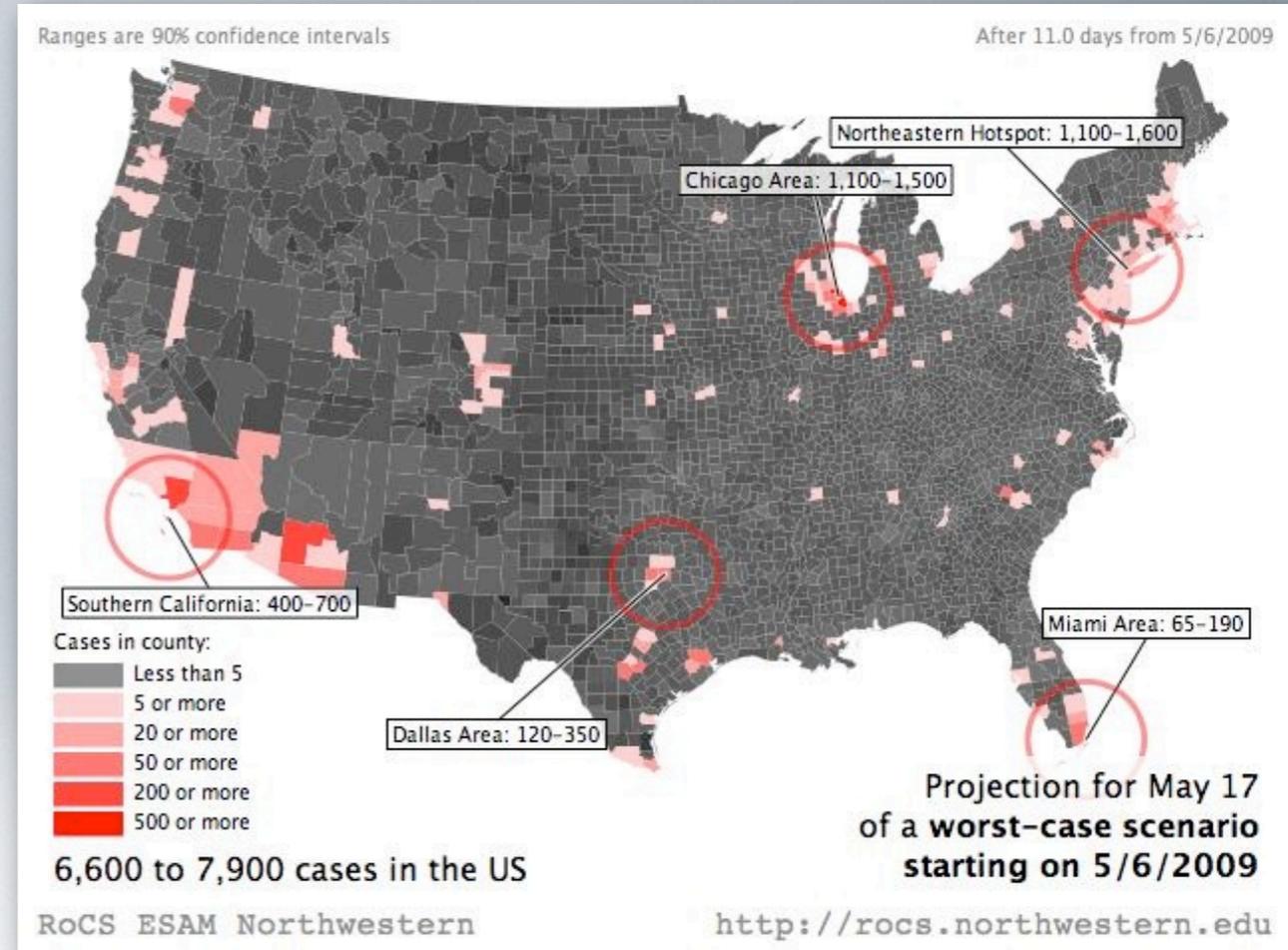
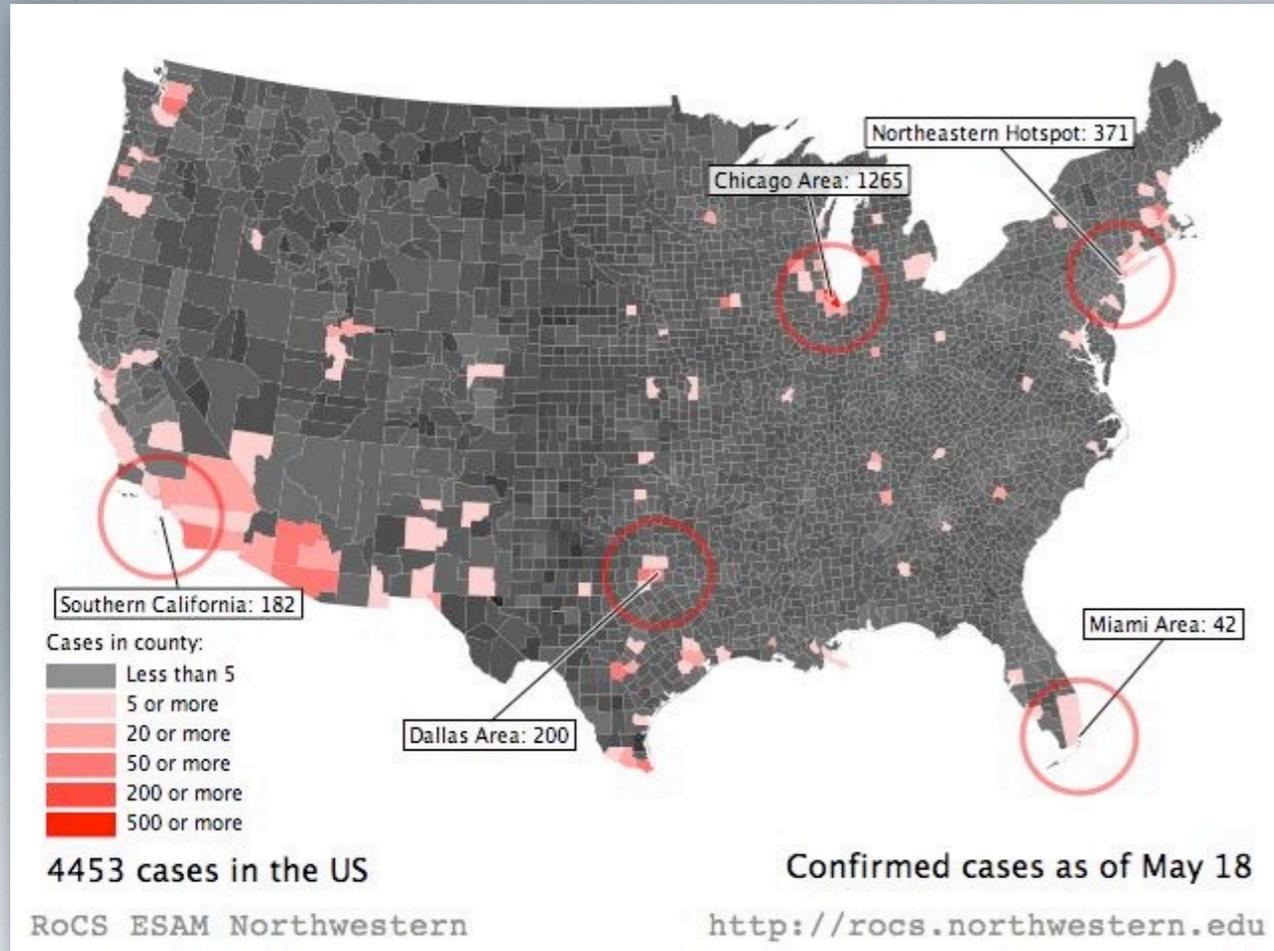
Cases in county:

- None
- 1 or more
- 5 or more
- 25 or more
- 100 or more
- 200 or more

H1N1 PROJECTIONS US

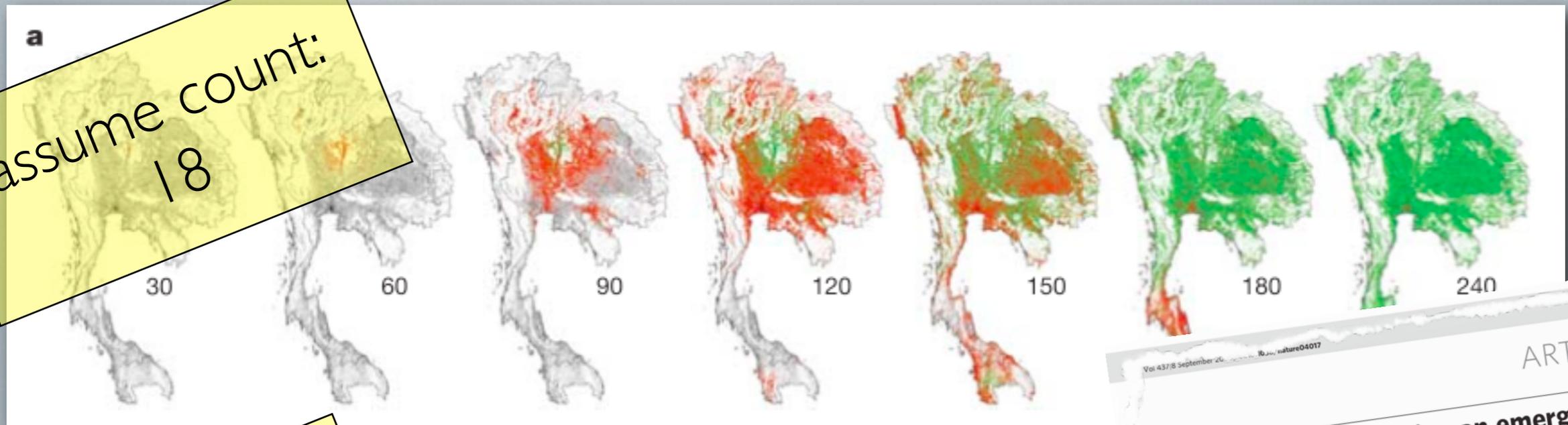


H1N1 PANDEMIC 2009



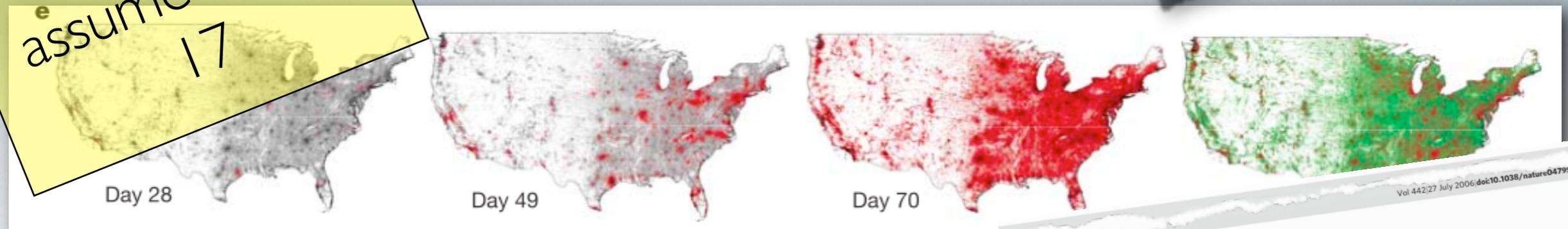
QUANTITATIVE MODELS

assume count:
18



ARTICLES
Vol 437/8 September 2006
Strategies for containing an emerging influenza pandemic in Southeast Asia
Neil M. Ferguson^{1,2}, Derek A.T. Cummings³, Simon Cauchemez⁴, Christophe Fraser¹, Donald S. Burke²

assume count:
17



LETTERS
Vol 442/27 July 2006; doi:10.1038/nature04795
Strategies for mitigating an influenza pandemic
Neil M. Ferguson¹, Derek A. T. Cummings², Christophe Fraser¹, James C. Cajka³, Philip C. Cooley³ & Donald S. Burke²

ROBERT MAY

VIEWPOINT

Uses and Abuses of Mathematics in Biology

Robert M. May

In the physical sciences, mathematical theory and experimental investigation have always marched together. Mathematics has been less intrusive in the life sciences, possibly because they have until recently been largely descriptive, lacking the invariance principles and fundamental natural constants of physics. Increasingly in recent decades, however, mathematics has become pervasive in biology, taking many different forms: statistics in experimental design; pattern seeking in bioinformatics; models in evolution, ecology, and epidemiology; and much else. I offer an opinionated overview of such uses—and abuses.

Darwin once wrote “I have deeply regretted that I did not proceed far enough at least to understand something of the great leading principles of mathematics; for men thus endowed seem to have an extra sense.” With the benefit of hindsight, we can see how much an “extra sense” could indeed

have solved one of Darwin’s major problems. In his day, it was thought that inheritance “blended” maternal and paternal characteristics. However, as pointed out to Darwin by the engineer Fleeming Jenkin and others, with blending inheritance it is virtually impossible to preserve the natural variation within populations that is both observed and essential to his theory of how evolution works. Mendel’s observations on

the particulate nature of inheritance were contemporary with Darwin, and his published work accessible to Darwin. Fisher and others have suggested that Fleeming Jenkin’s fundamental and intractable objections to *The Origin of Species* could have been resolved by Darwin or one of his colleagues, if only they had grasped the mathematical significance of Mendel’s results (1). But half a century elapsed before Hardy and Weinberg (H-W) resolved the difficulties by proving that particulate inheritance preserved variation within populations (2).

Today, the H-W Law stands as a kind of Newton’s First Law (bodies remain in their state of rest or uniform motion in a straight line, except insofar as acted upon by external forces) for evolution: Gene frequencies in a

Zoology Department, Oxford University, Oxford OX1 3PS, UK.

6 FEBRUARY 2004 VOL 303 SCIENCE www.sciencemag.org

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PS, UK.

The increasing speed and sophistication and ease of use of computers enables an increasingly large number of life scientists who have no substantial background in mathematics to explore "mathematical models" and draw conclusions about them.

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Such activity usually consists of representing sensible and evidence-based assumptions as the starting point for a complicated and usually nonlinear dynamical system, assigning particular parameters (often in an arbitrary way), and then letting this complicated system rip.

ROBERT MAY

Uses and Abuses of Mathematics in Biology

Robert M. May

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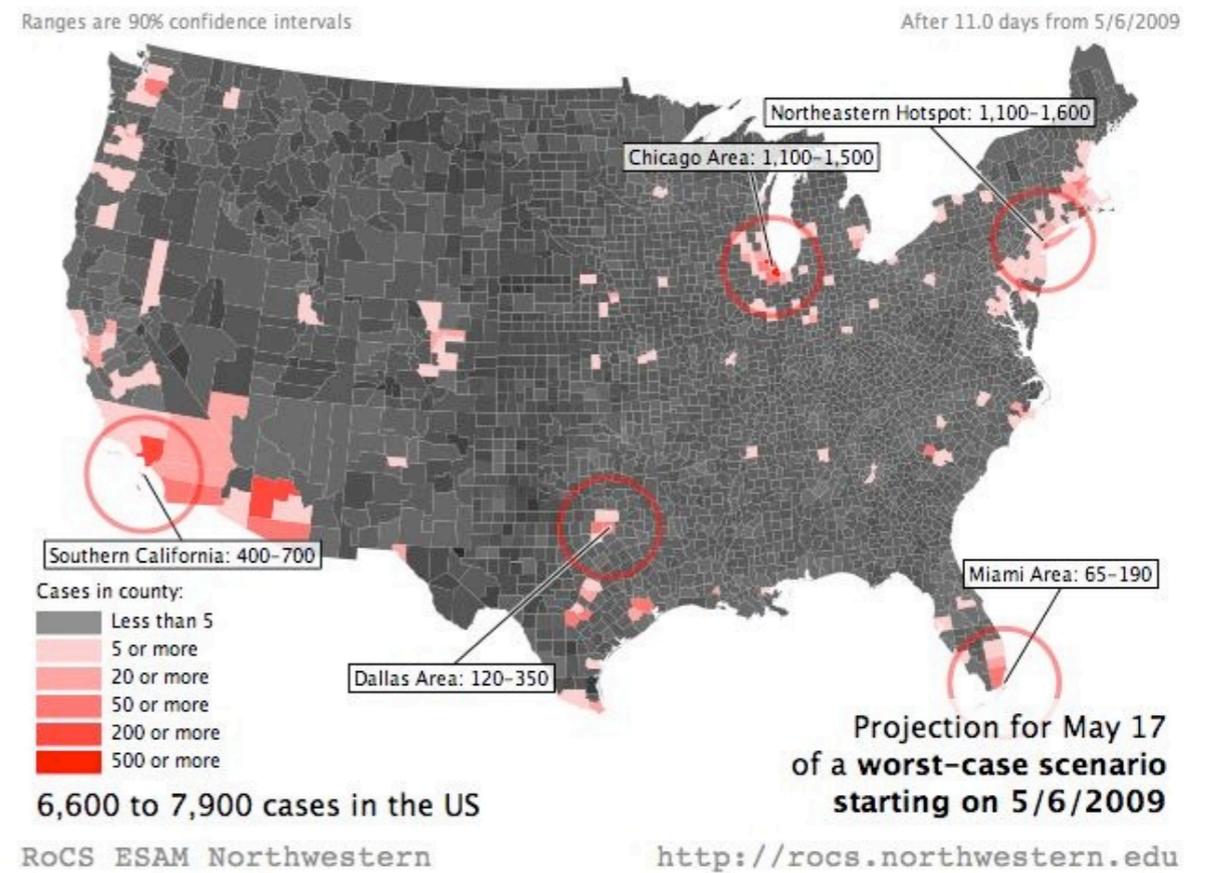
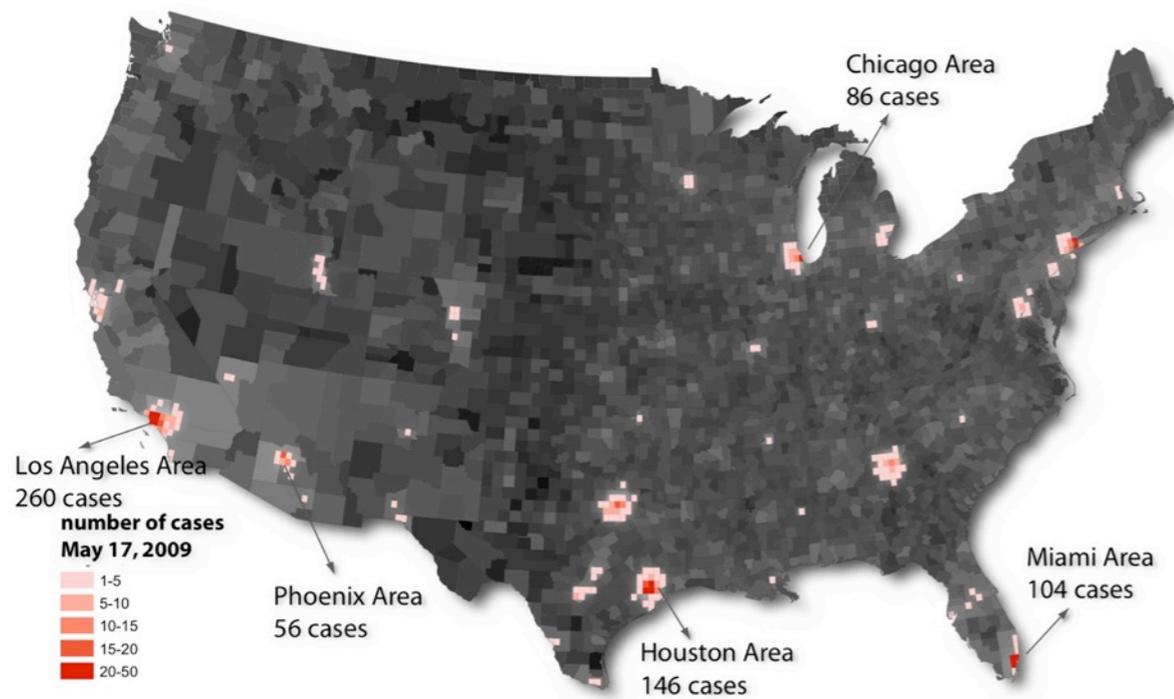
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we arguably are seeing an increasingly large body of work in which sweeping conclusions - "emergent phenomena" - are drawn from the alleged working of a mathematical model, without clear understanding of what is actually going on. I think this can be worrying.

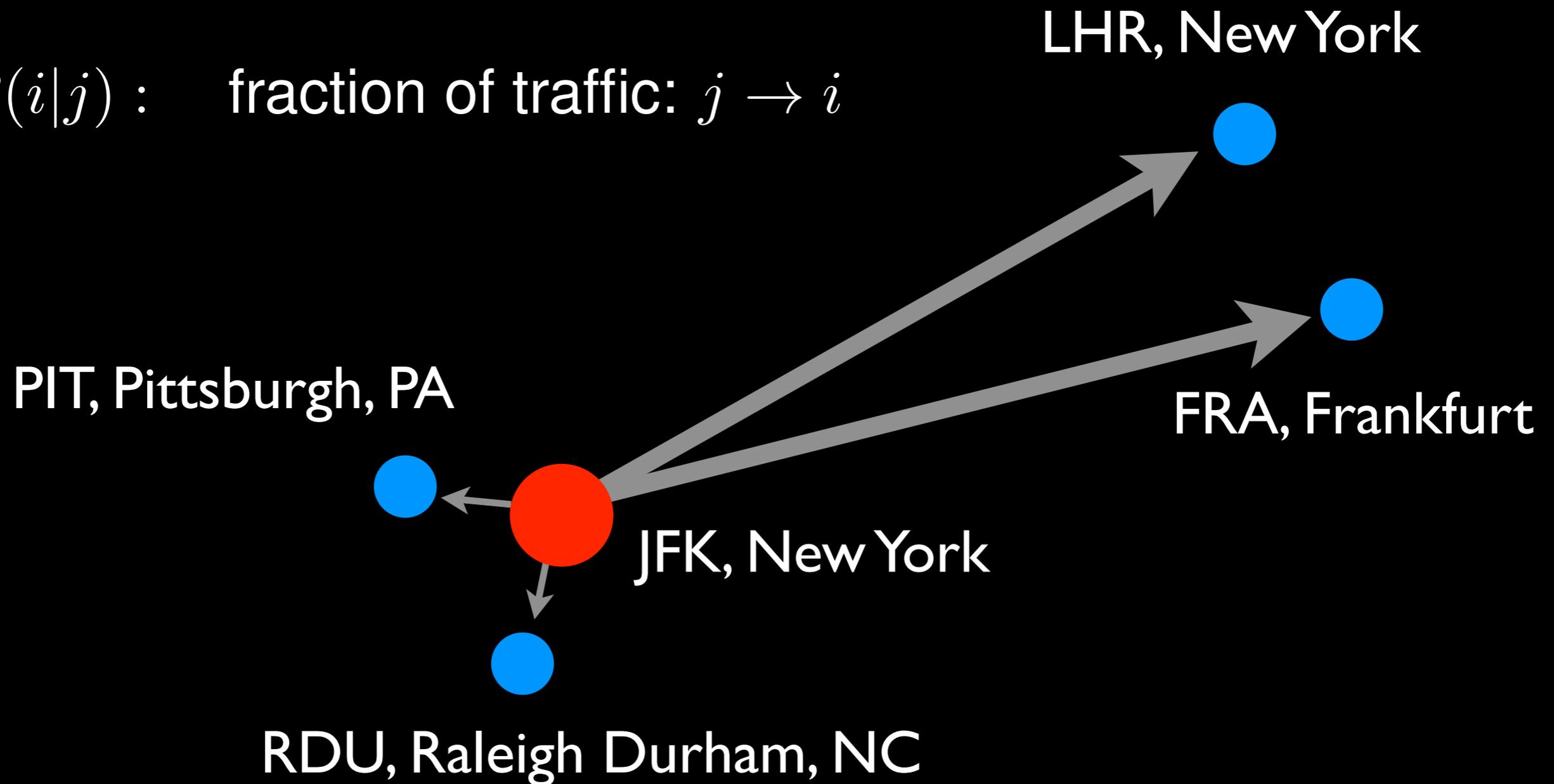


is something more fundamental going on?

redefining the notion of distance

simple idea: effective distance

$P(i|j)$: fraction of traffic: $j \rightarrow i$



a different notion of distance

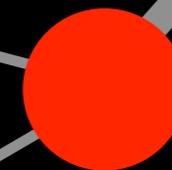
PIT, Pittsburgh, PA



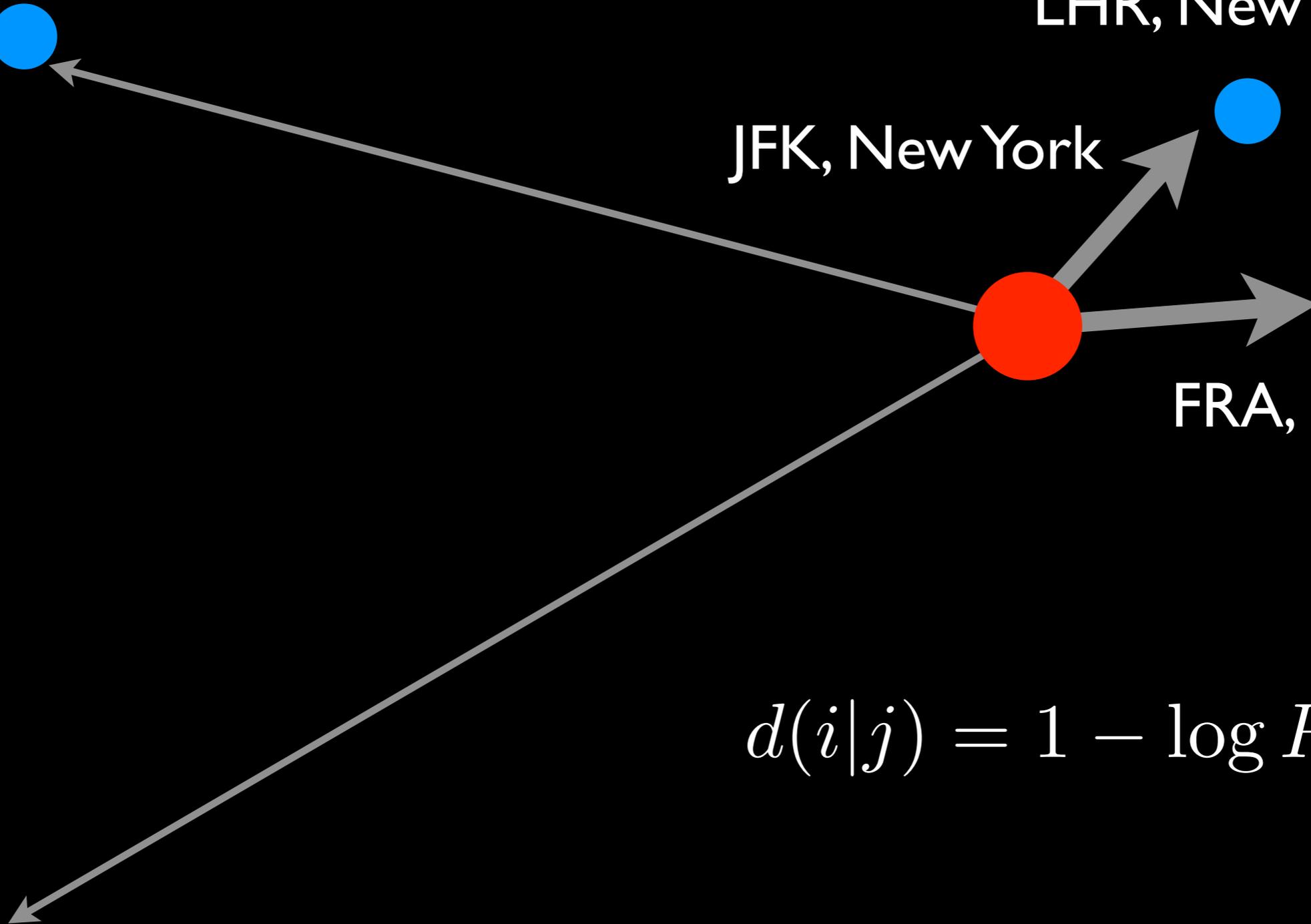
LHR, New York



JFK, New York



FRA, Frankfurt



$$d(i|j) = 1 - \log P(i|j)$$

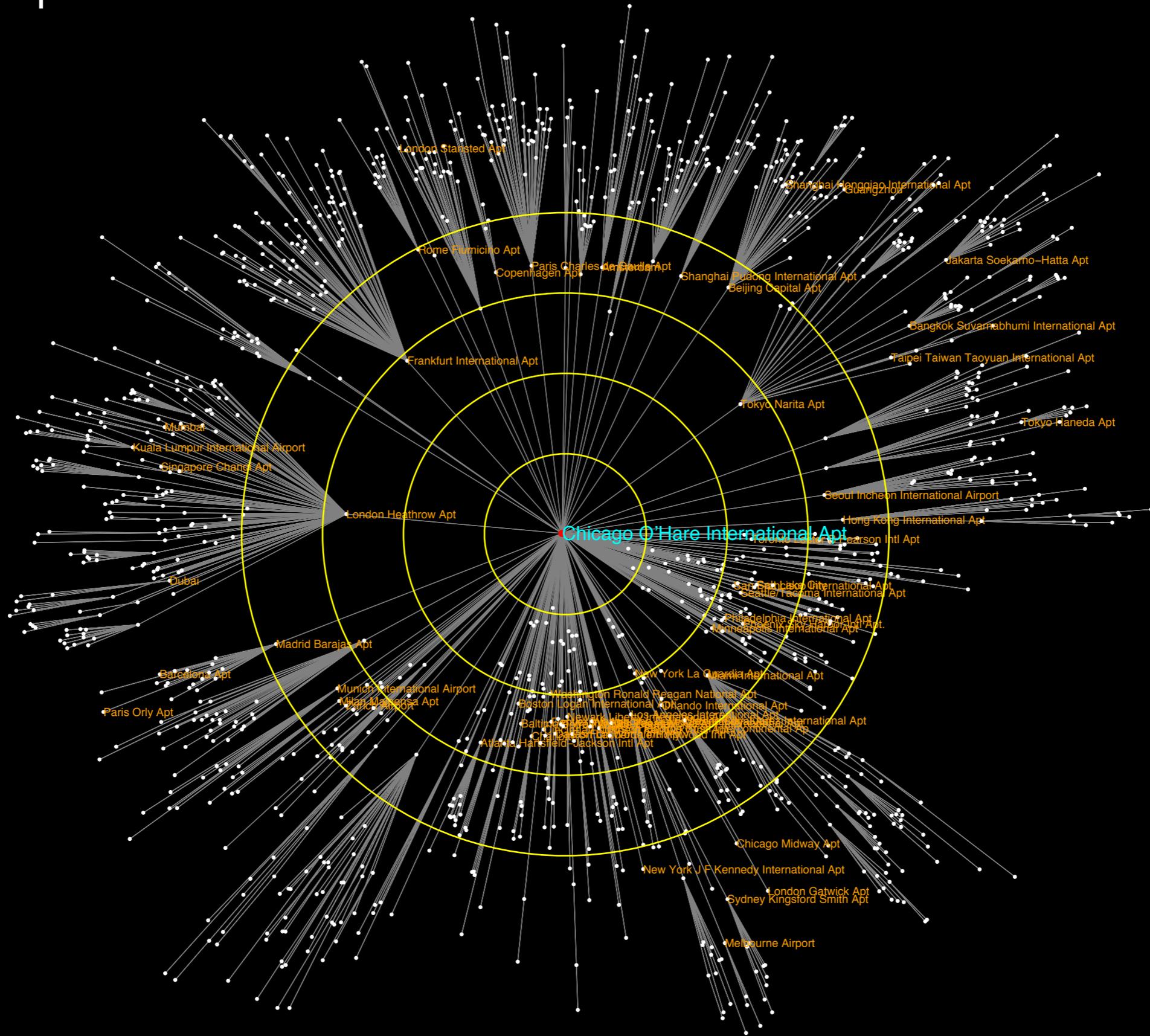
shortest path trees

$$d(i|j) = 1 - \log P(i|j) \quad \lambda(\Gamma) = L - \sum_{i=1}^{L-1} \log P(i|j)$$

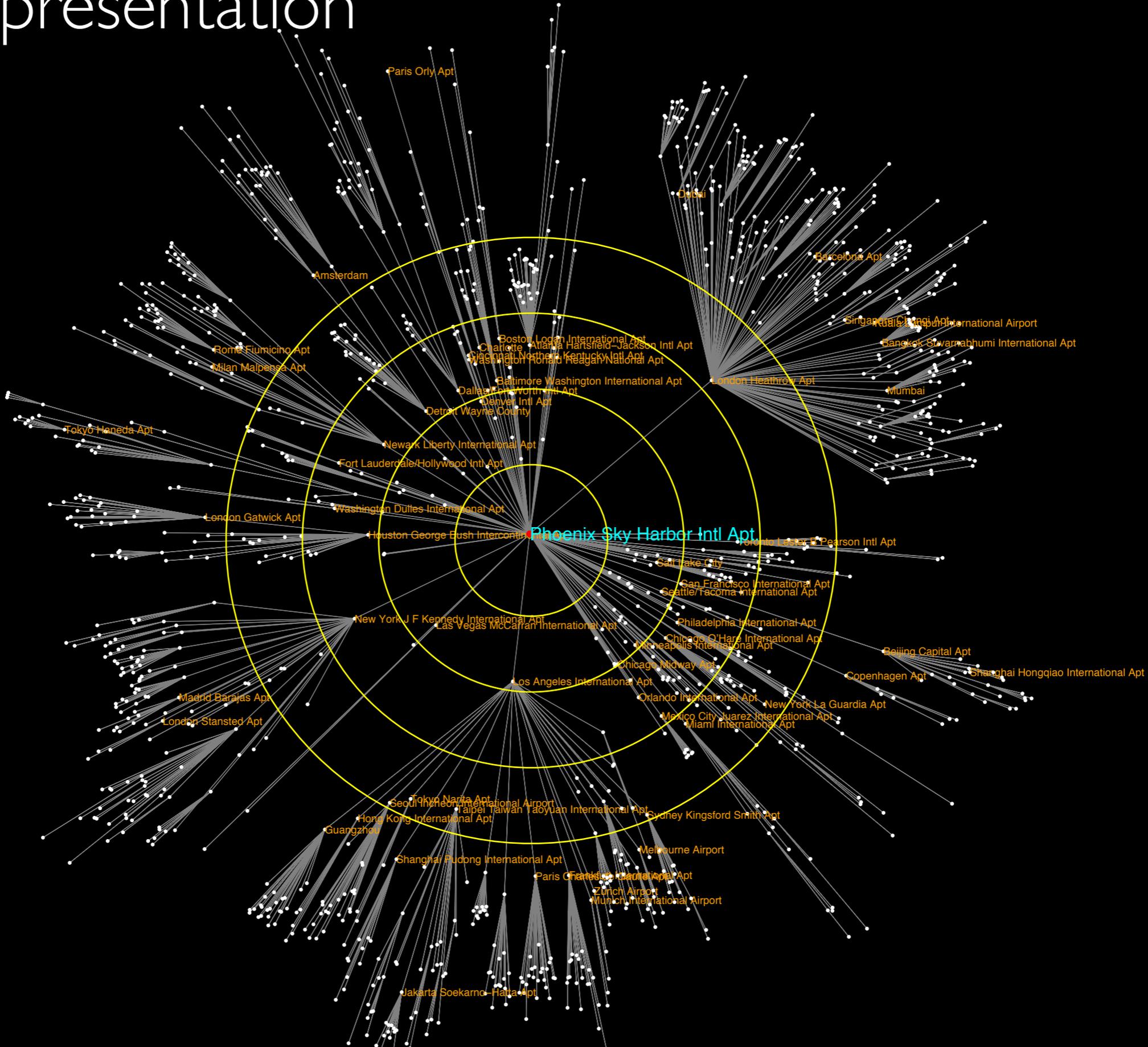


$$D(m|n) = \min_{\Gamma} \lambda(\Gamma)$$

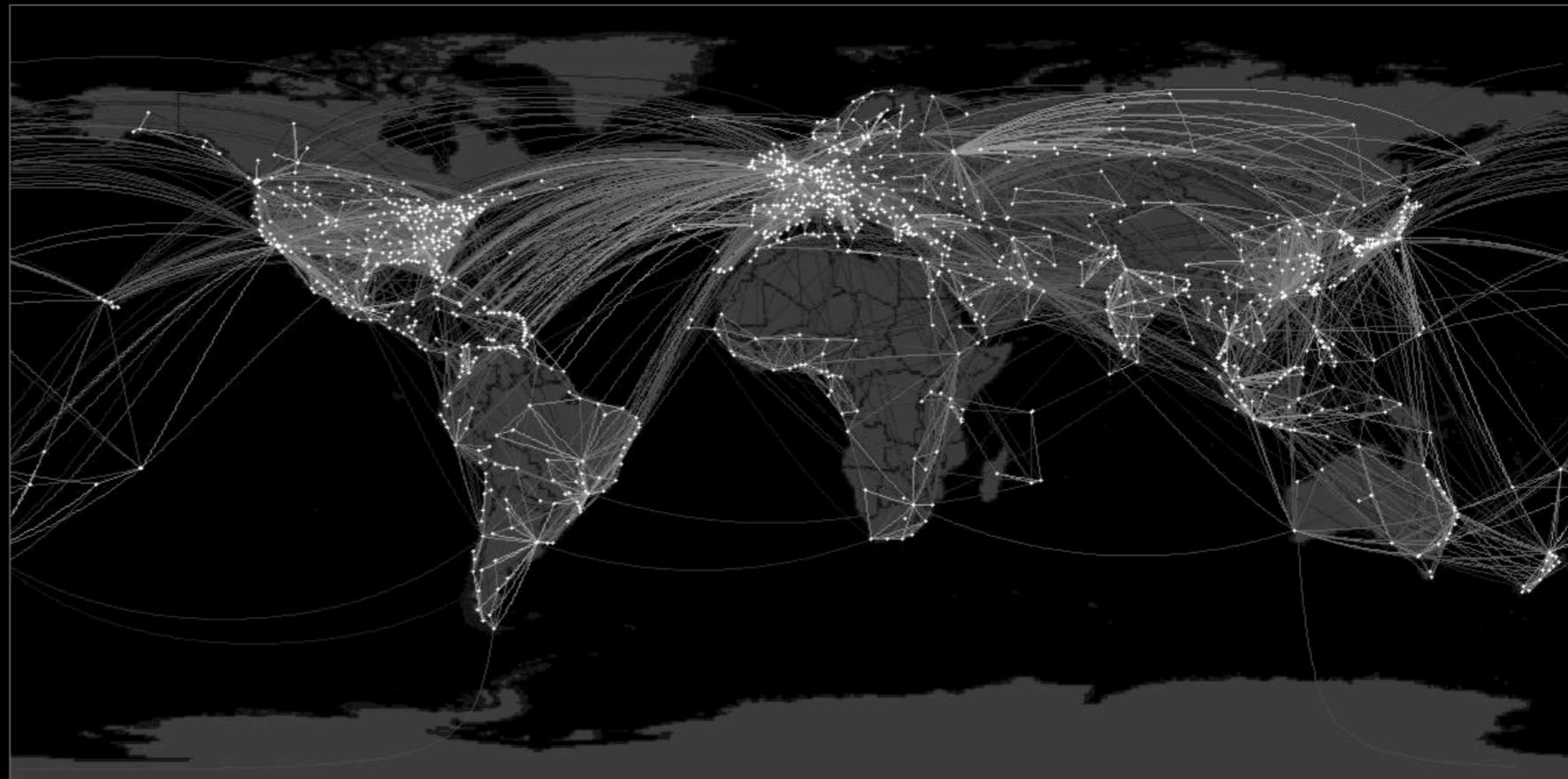
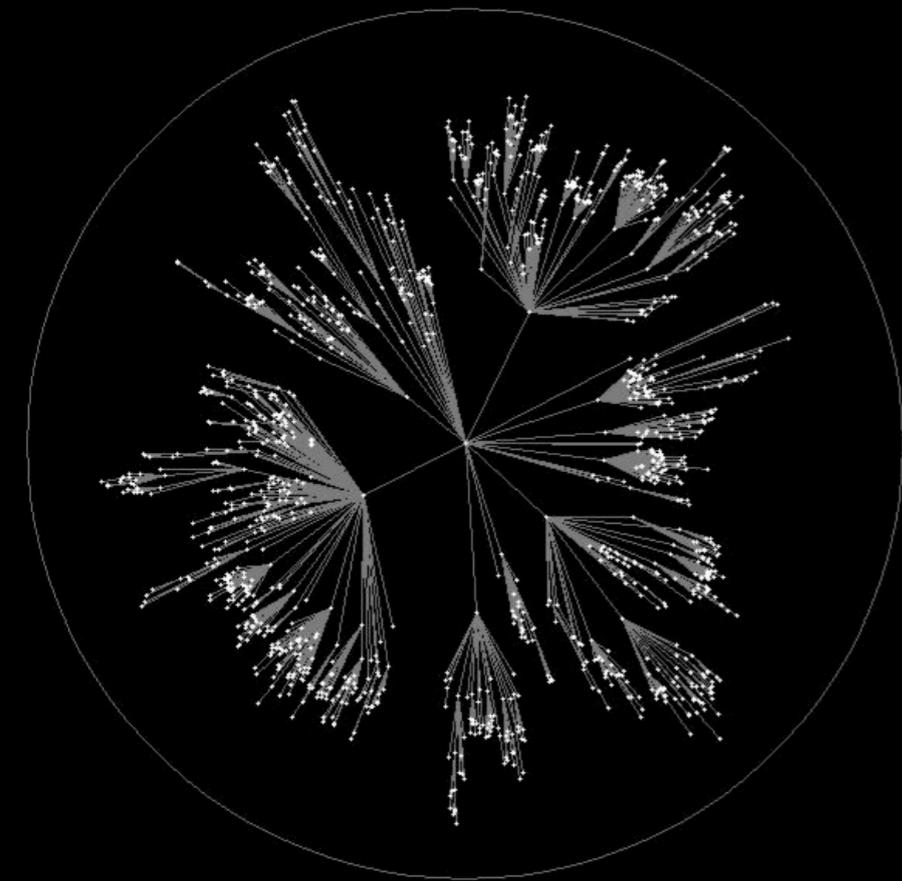
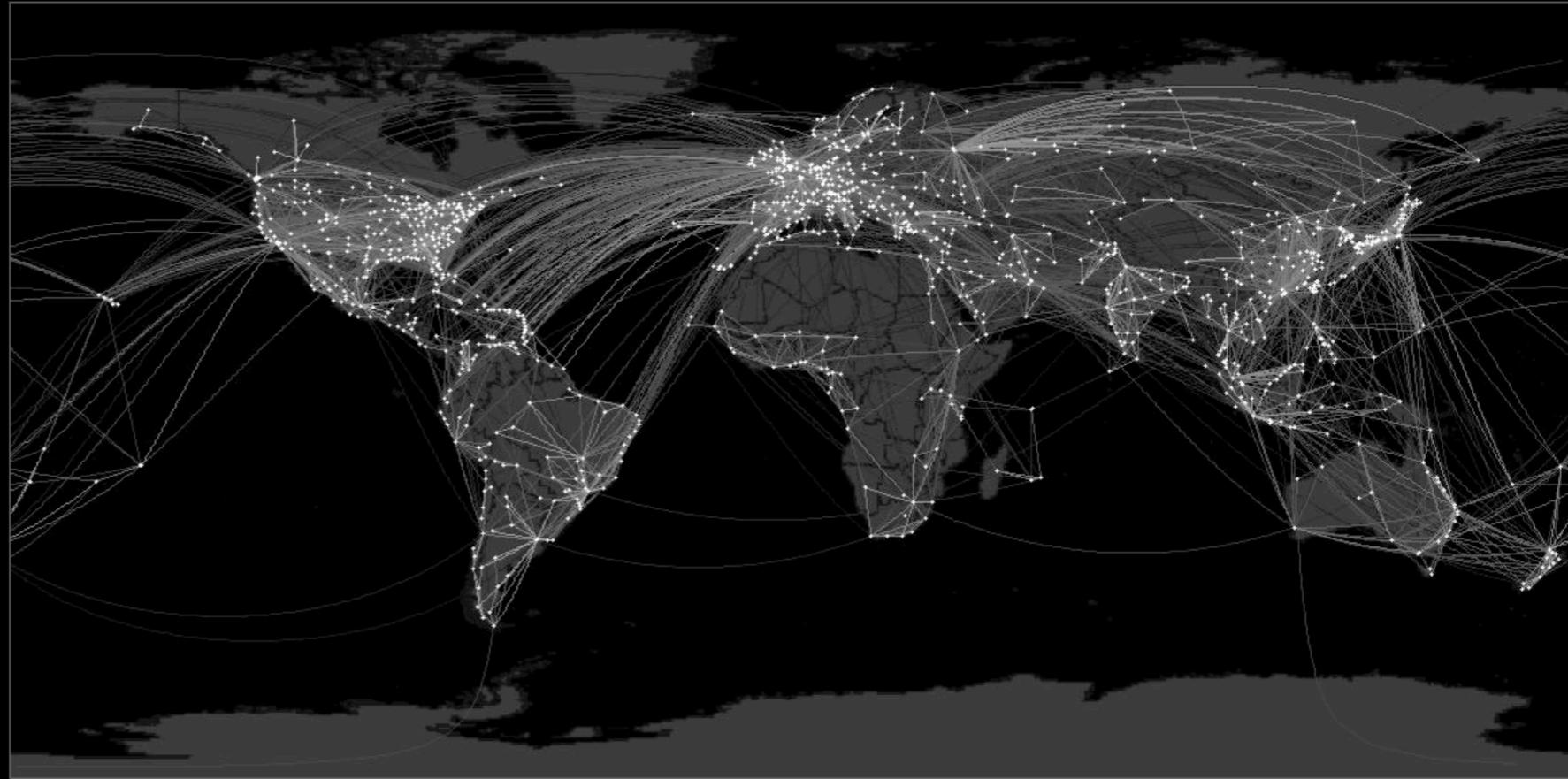
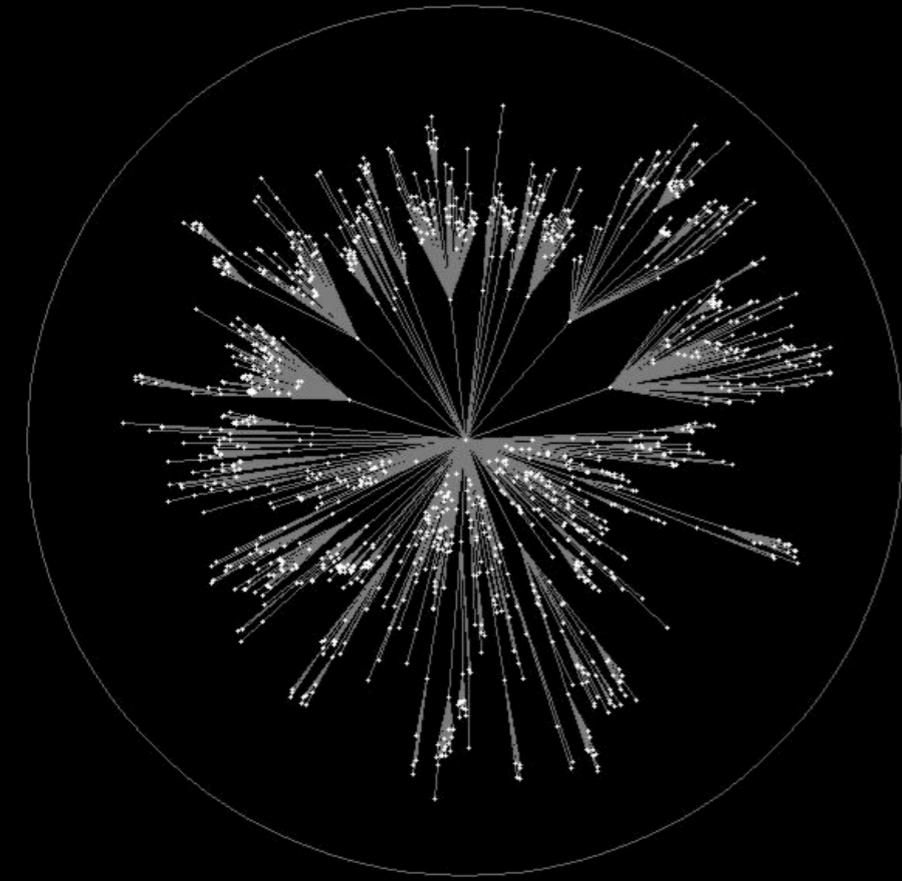
SPT representation



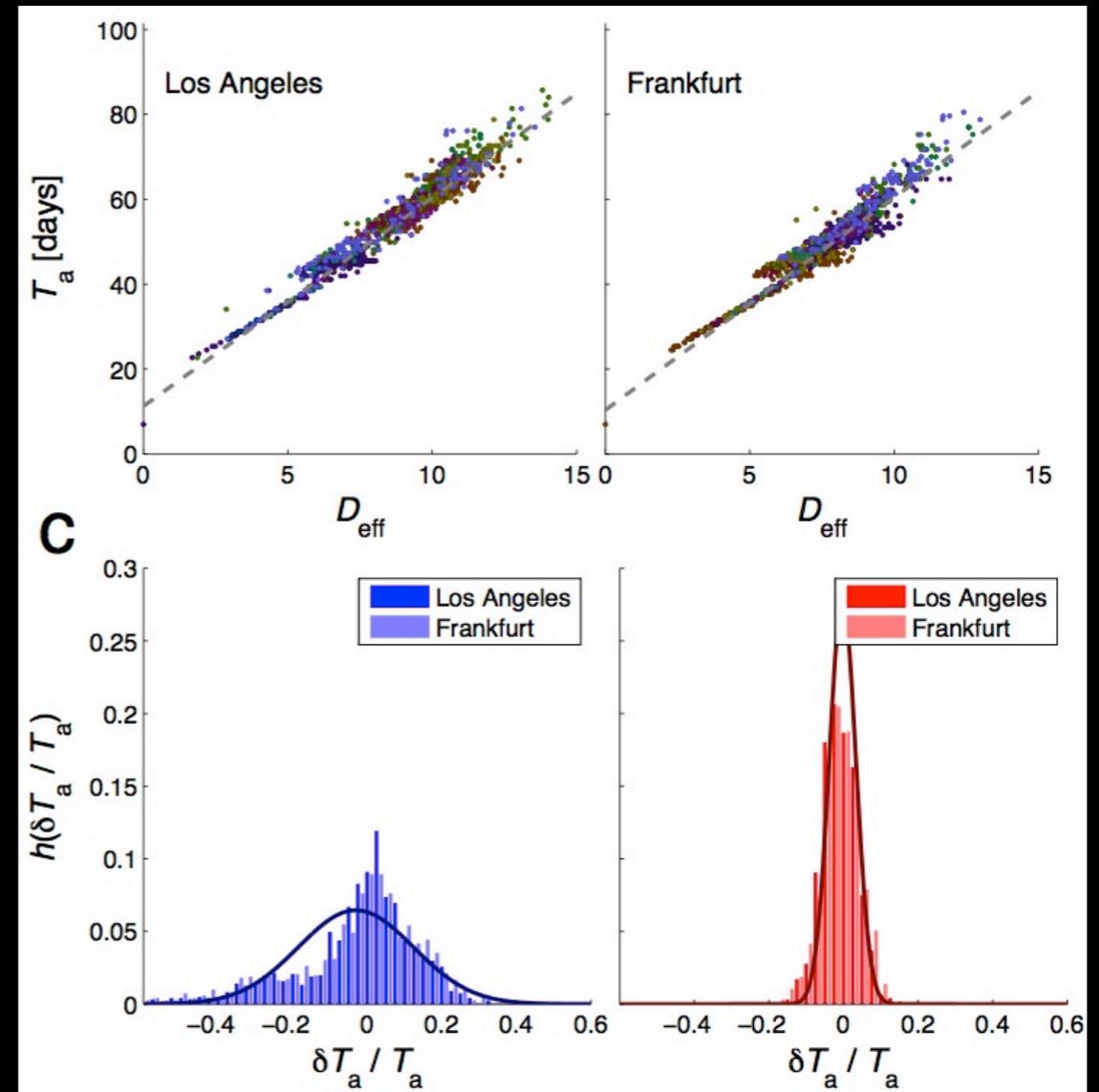
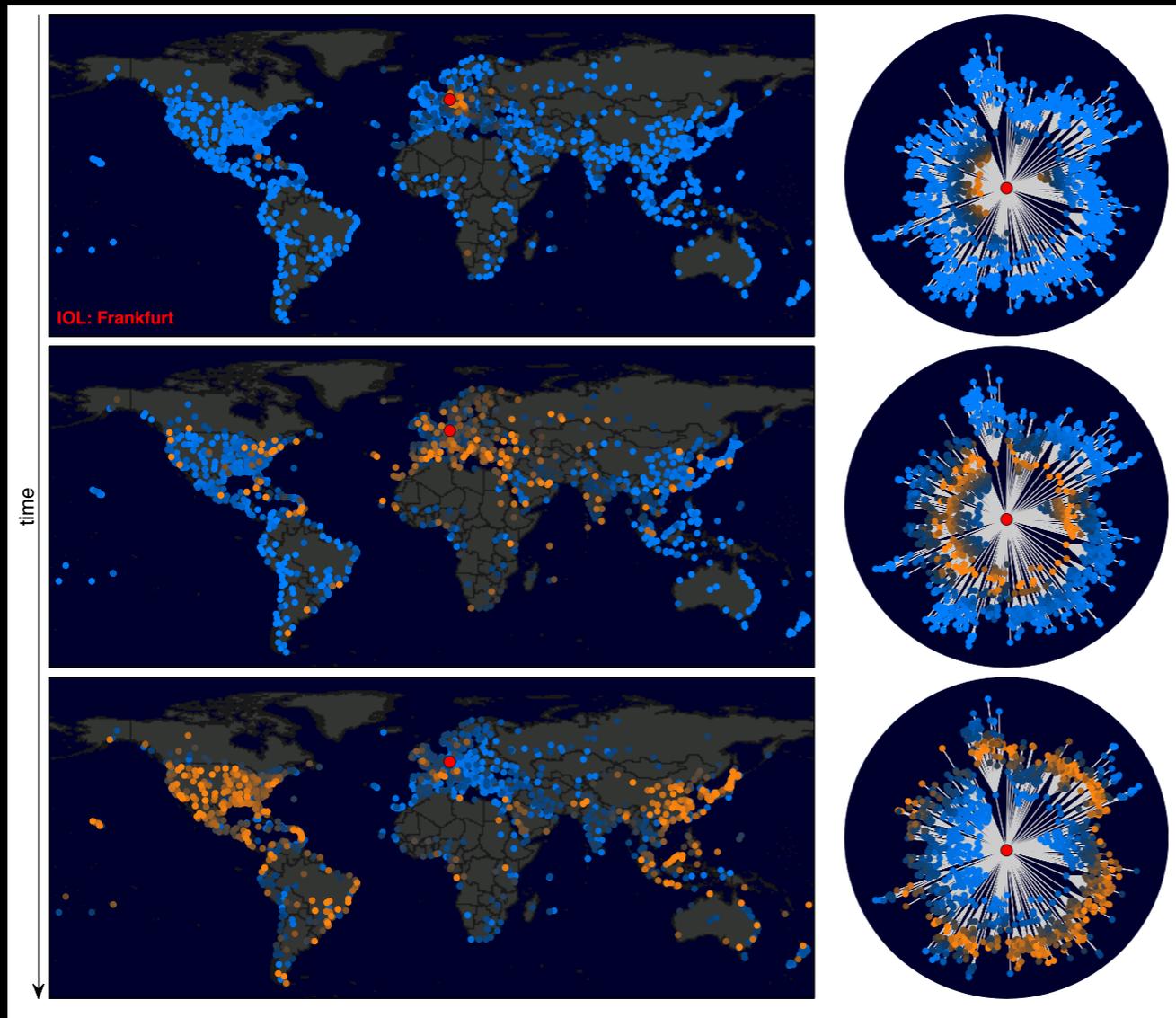
SPT representation

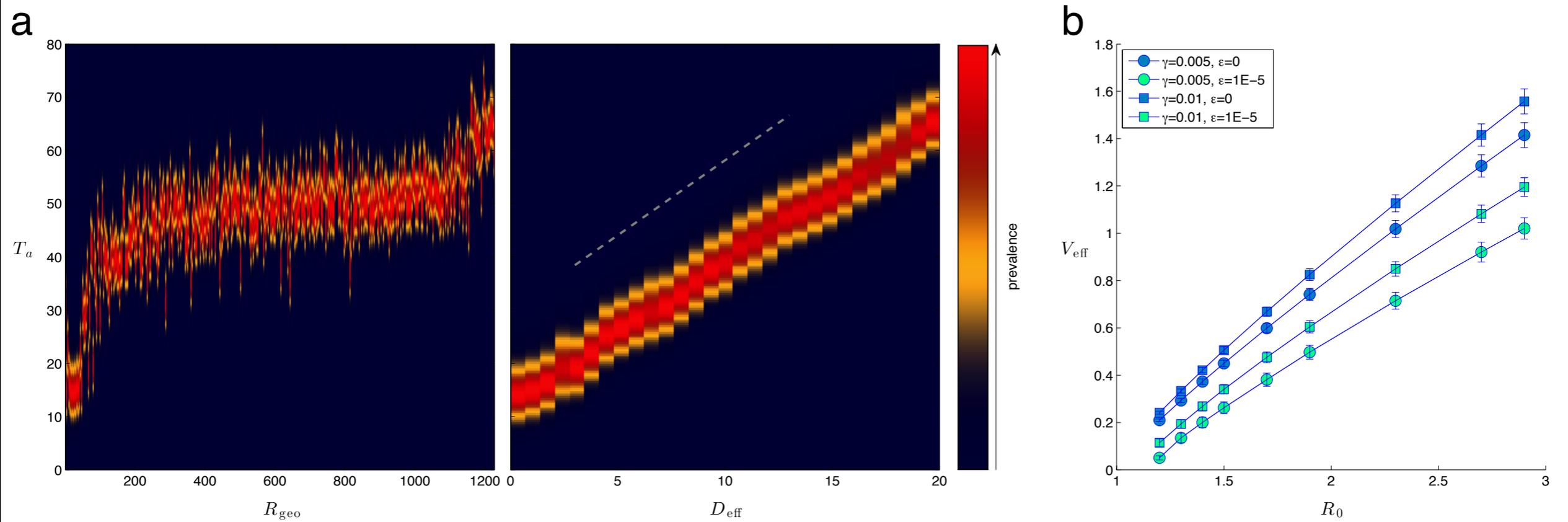


propagating waves



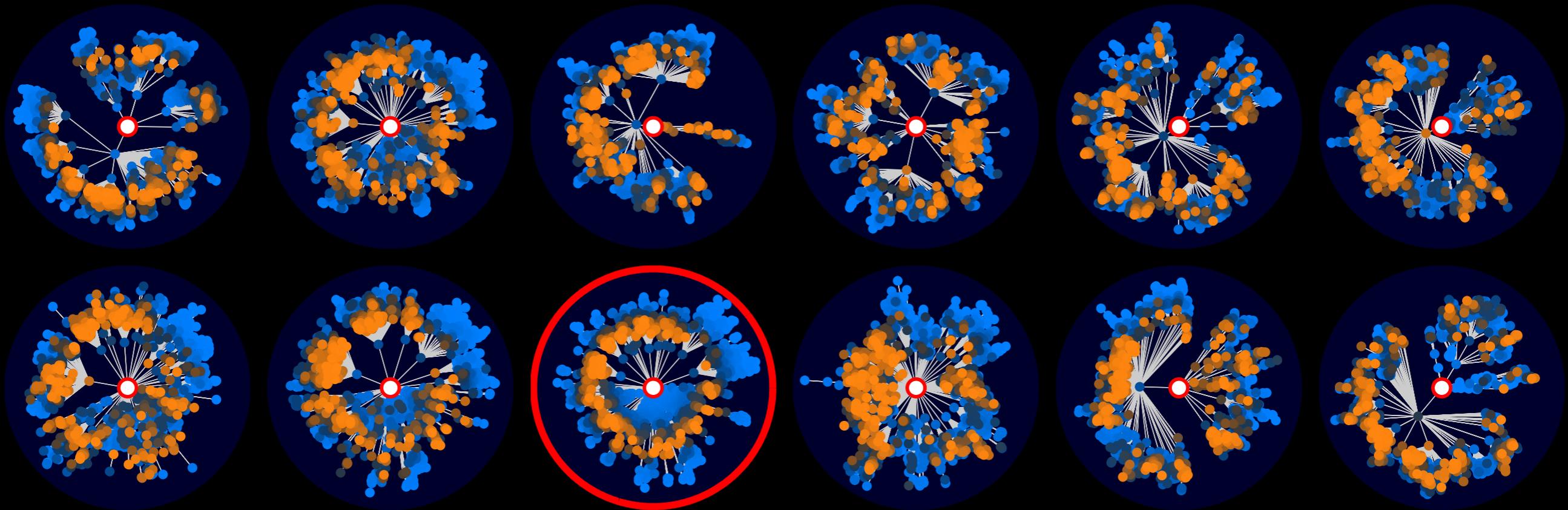
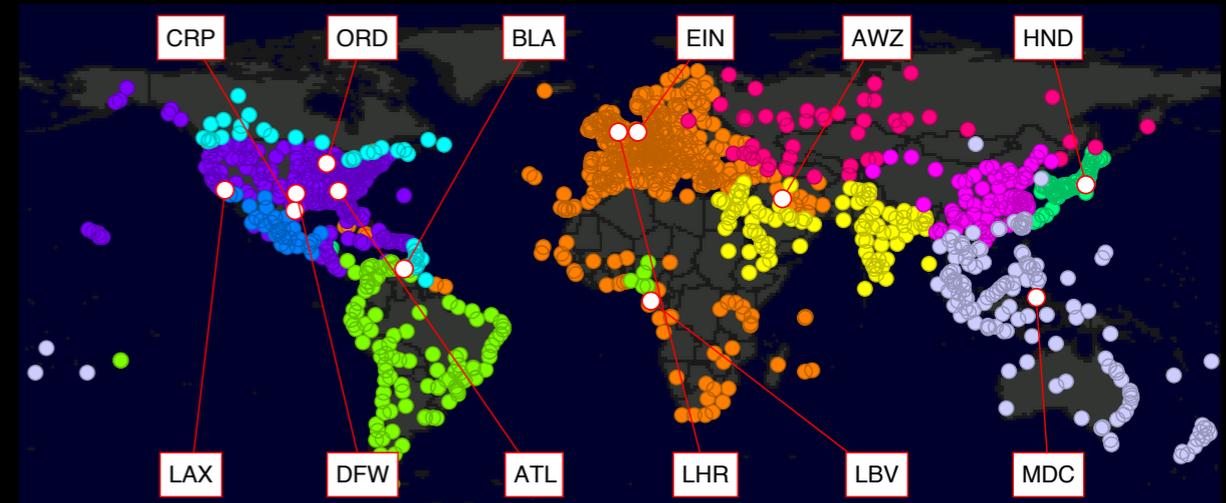
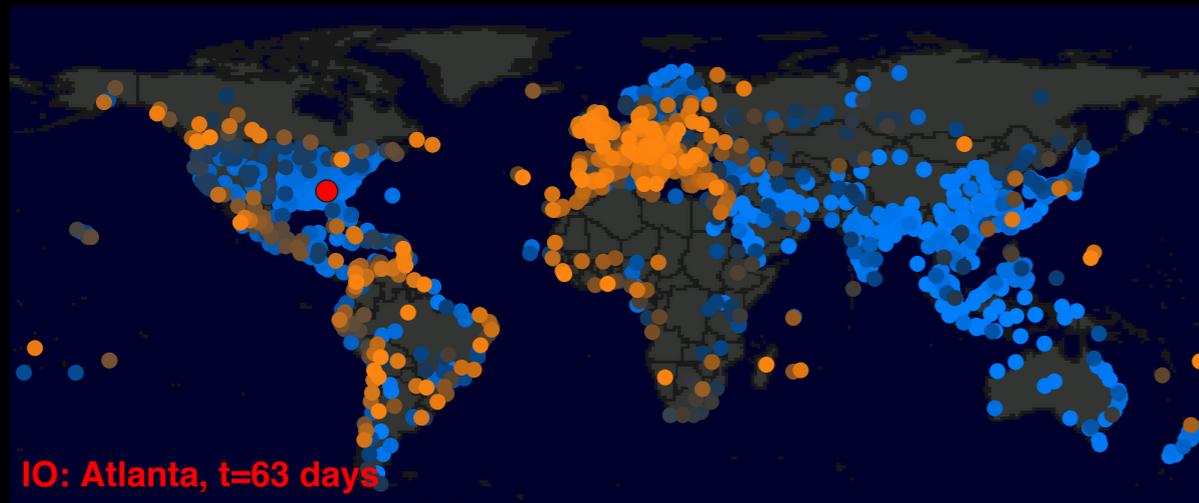
the spread in effective distance



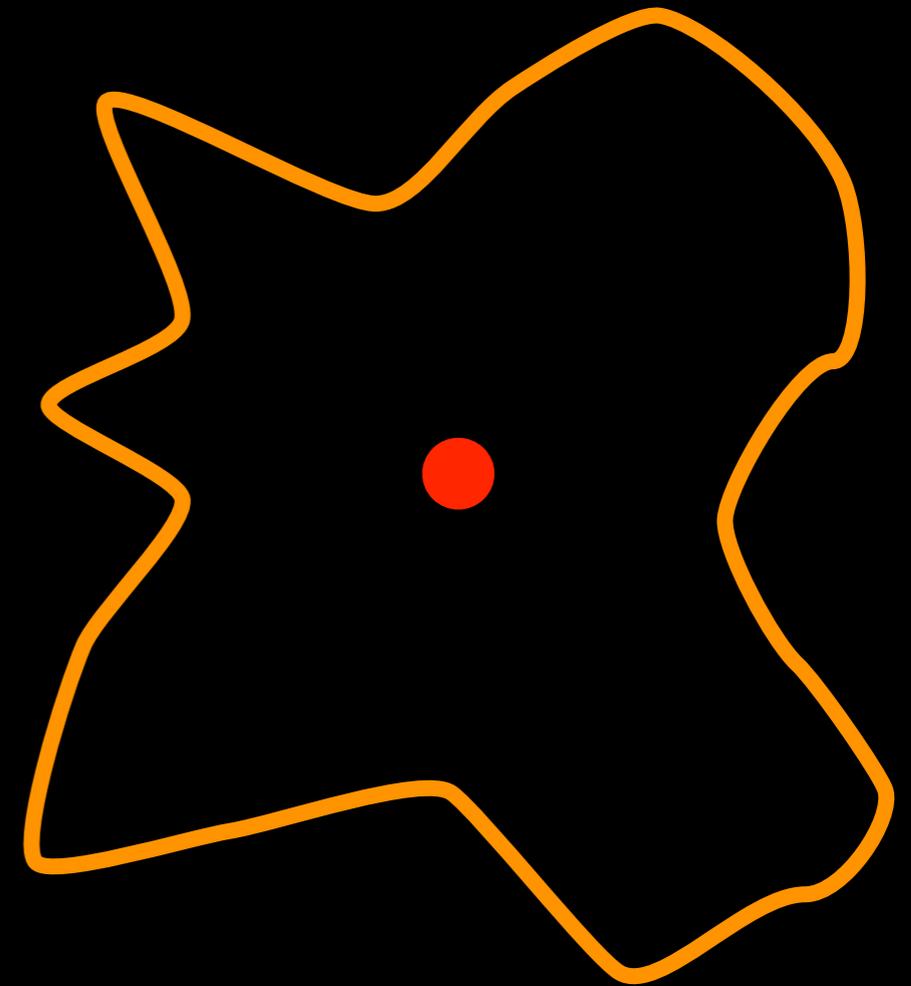
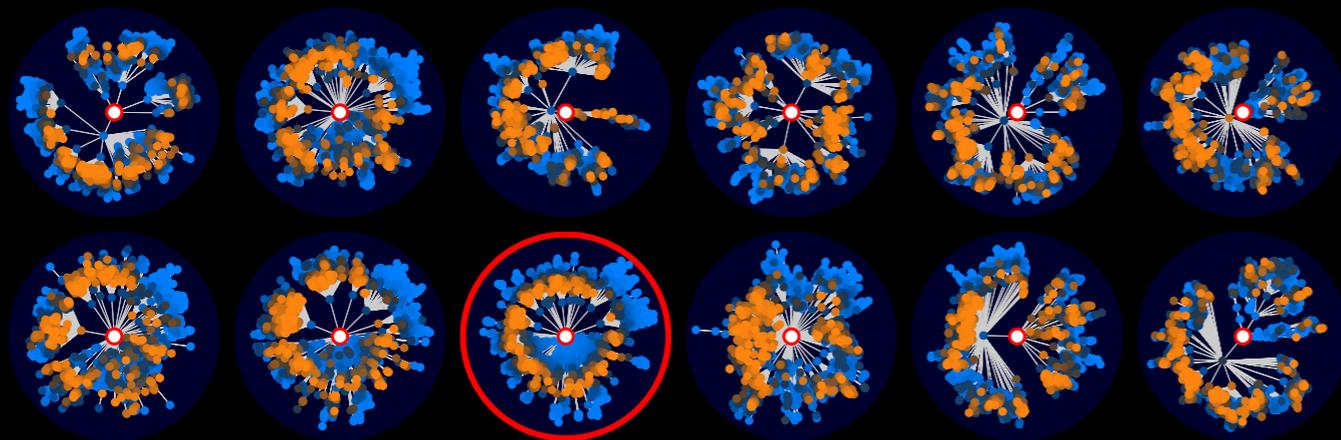
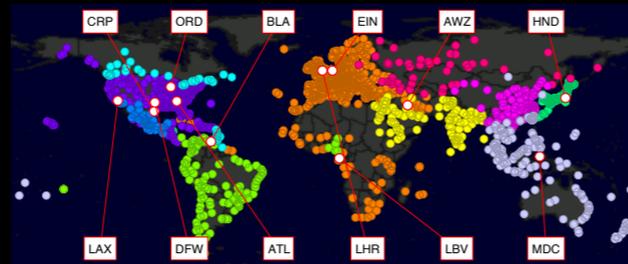
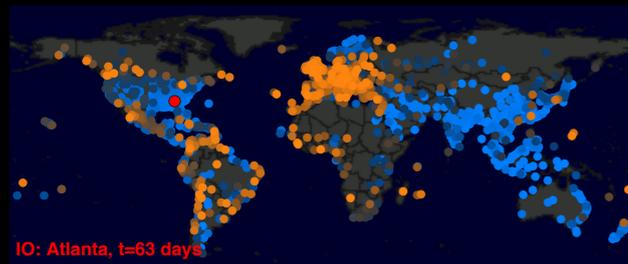


reconstructing an outbreak origin

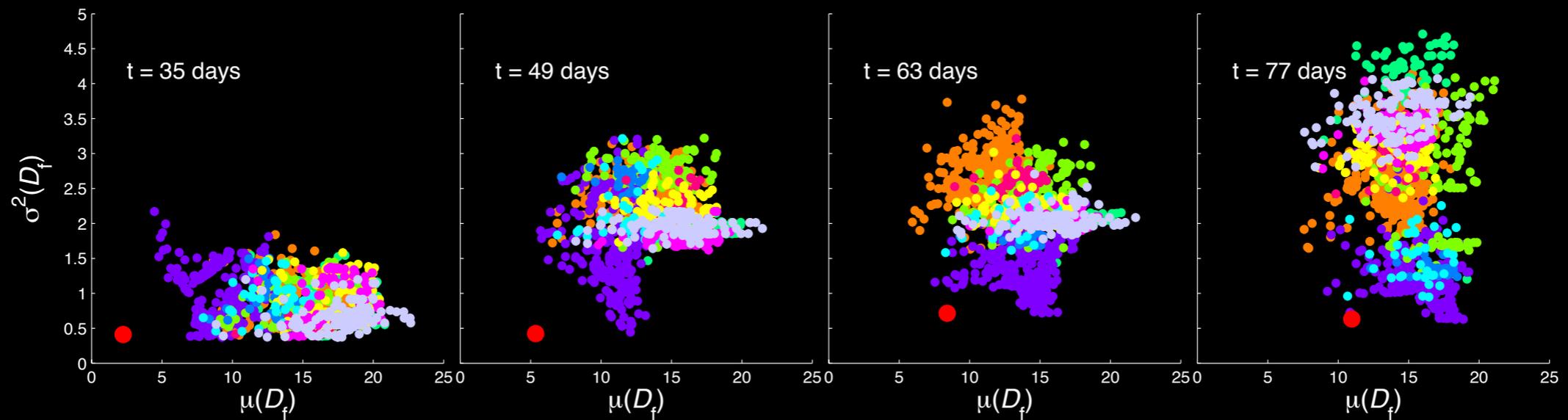
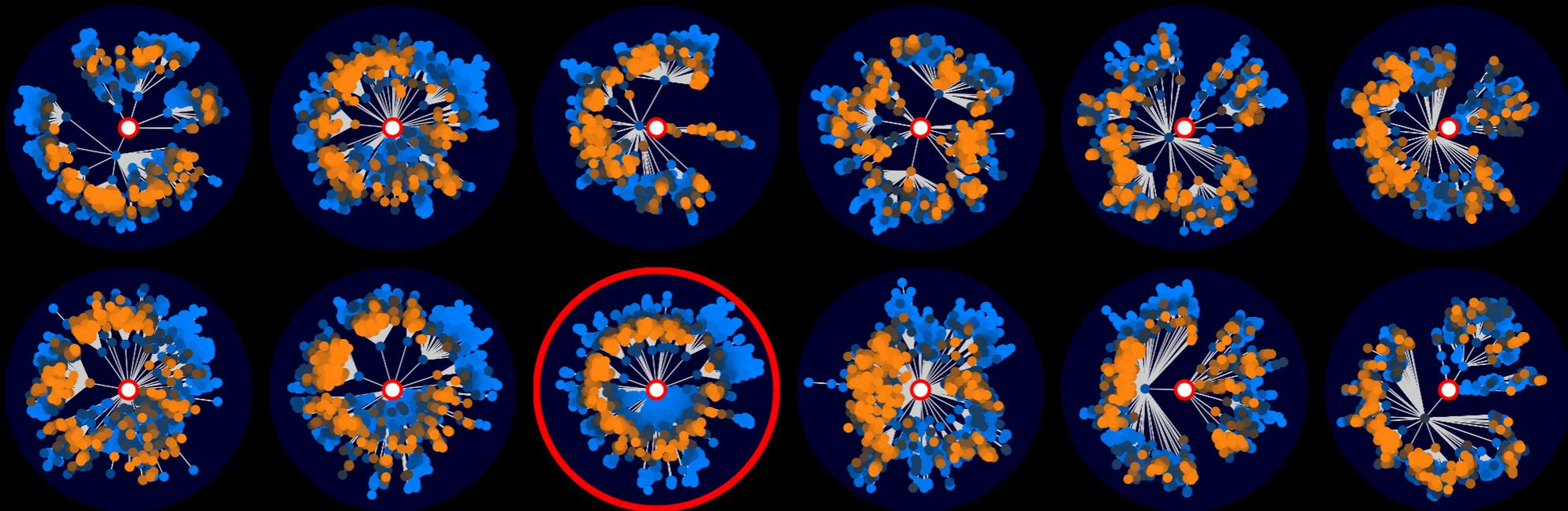
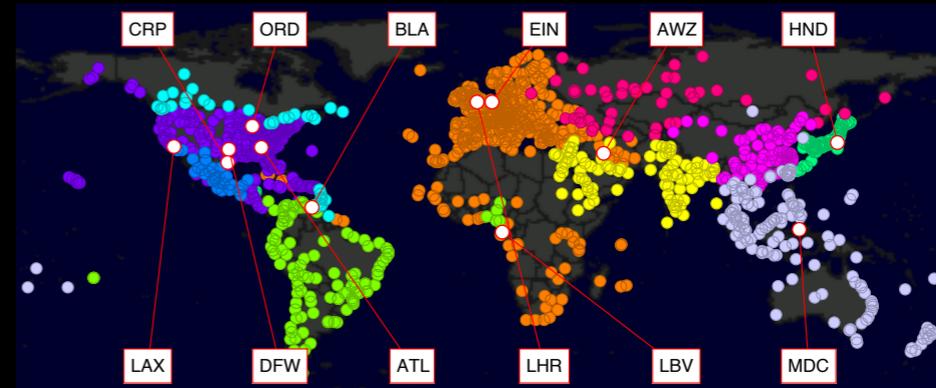
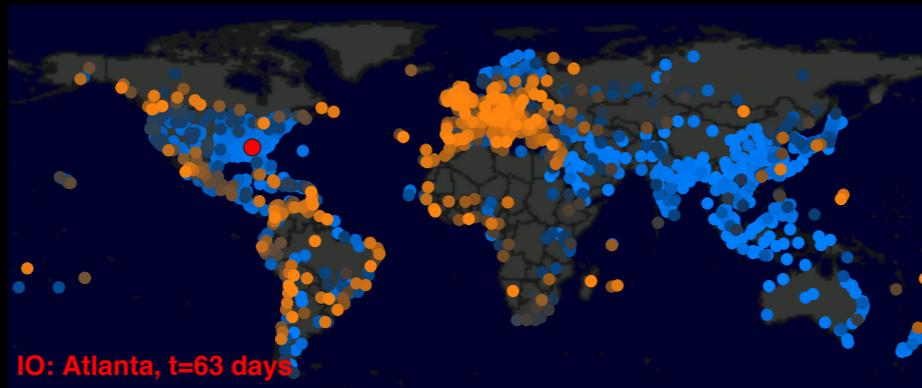
outbreak origin reconstruction



outbreak origin reconstruction



outbreak origin reconstruction



a key component of predictability



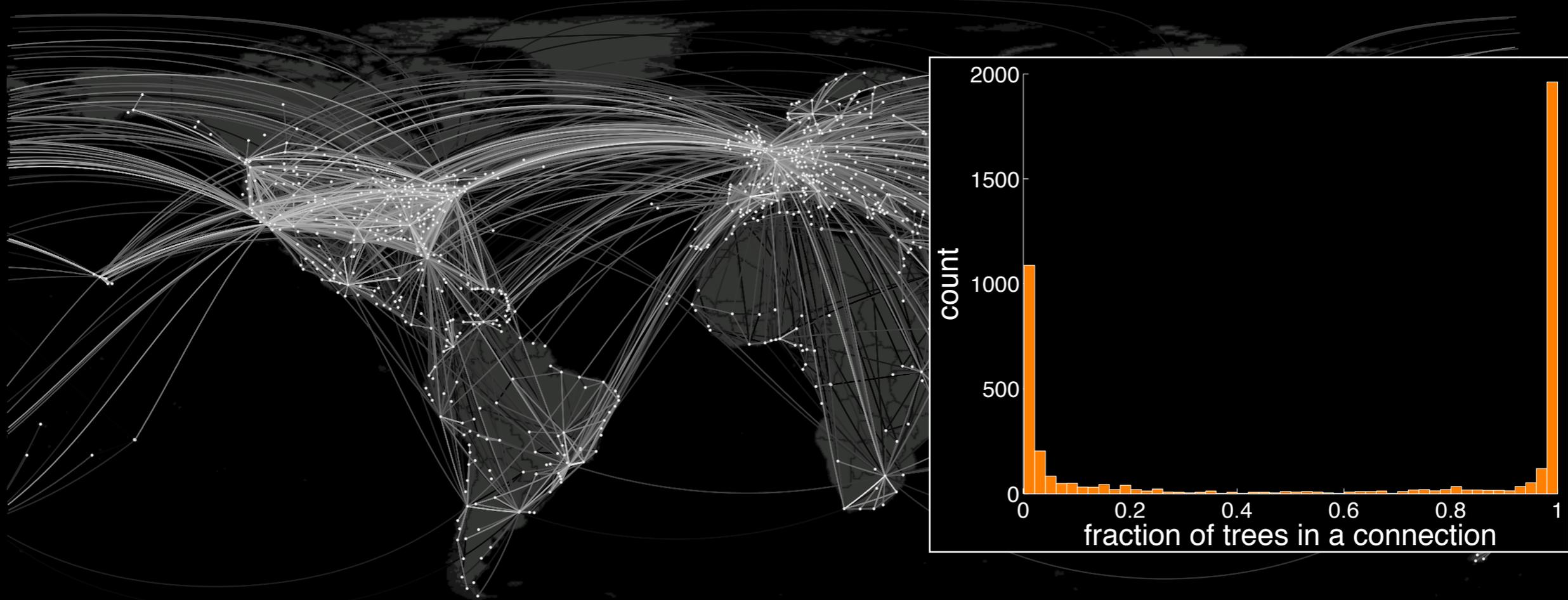
Madrid

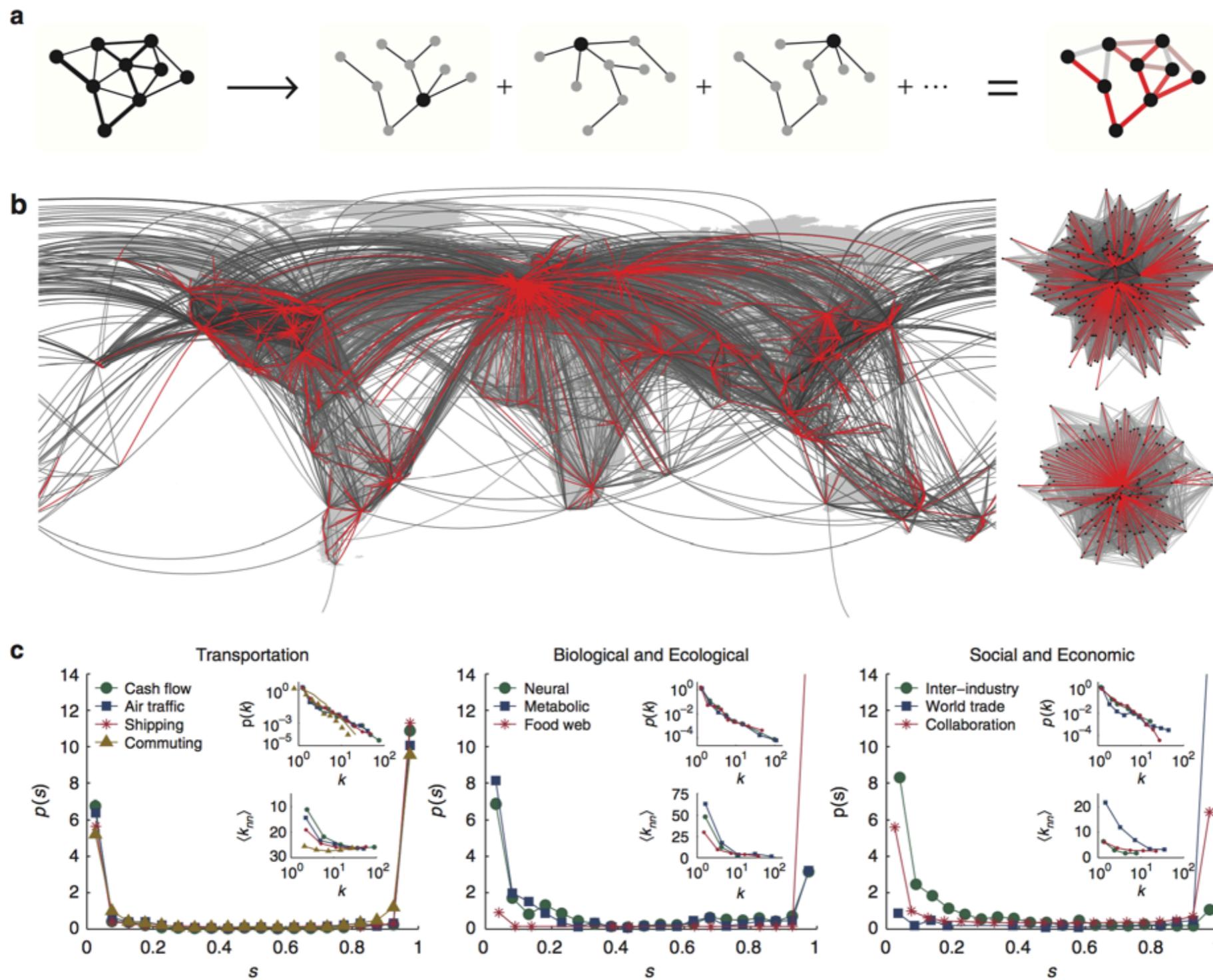


NY



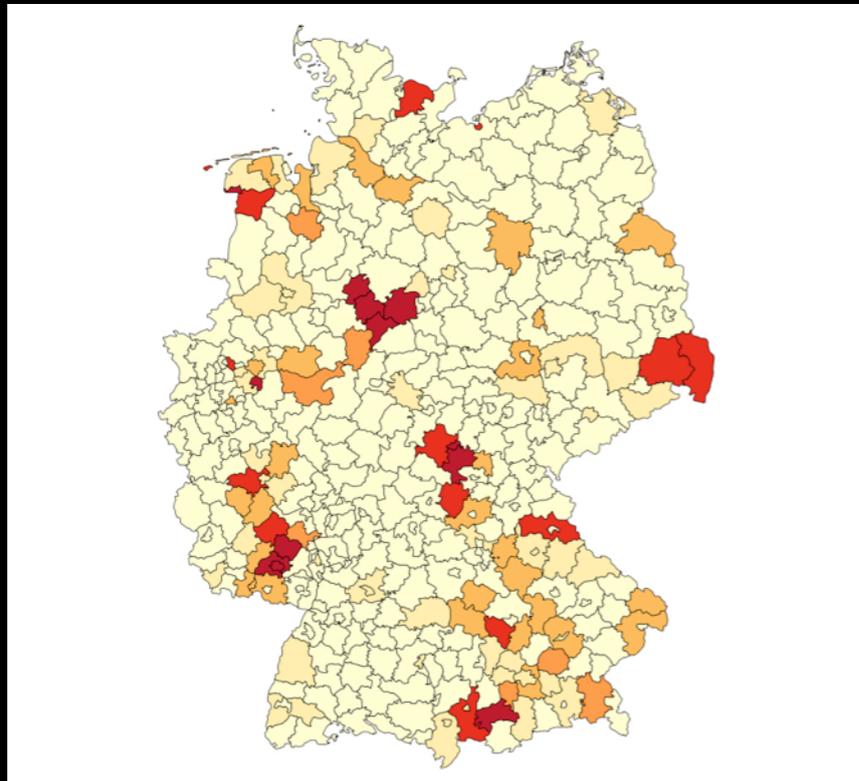
Frankfurt





ongoing projects

HUS/EHEC 2011



neolithic spread of farming

