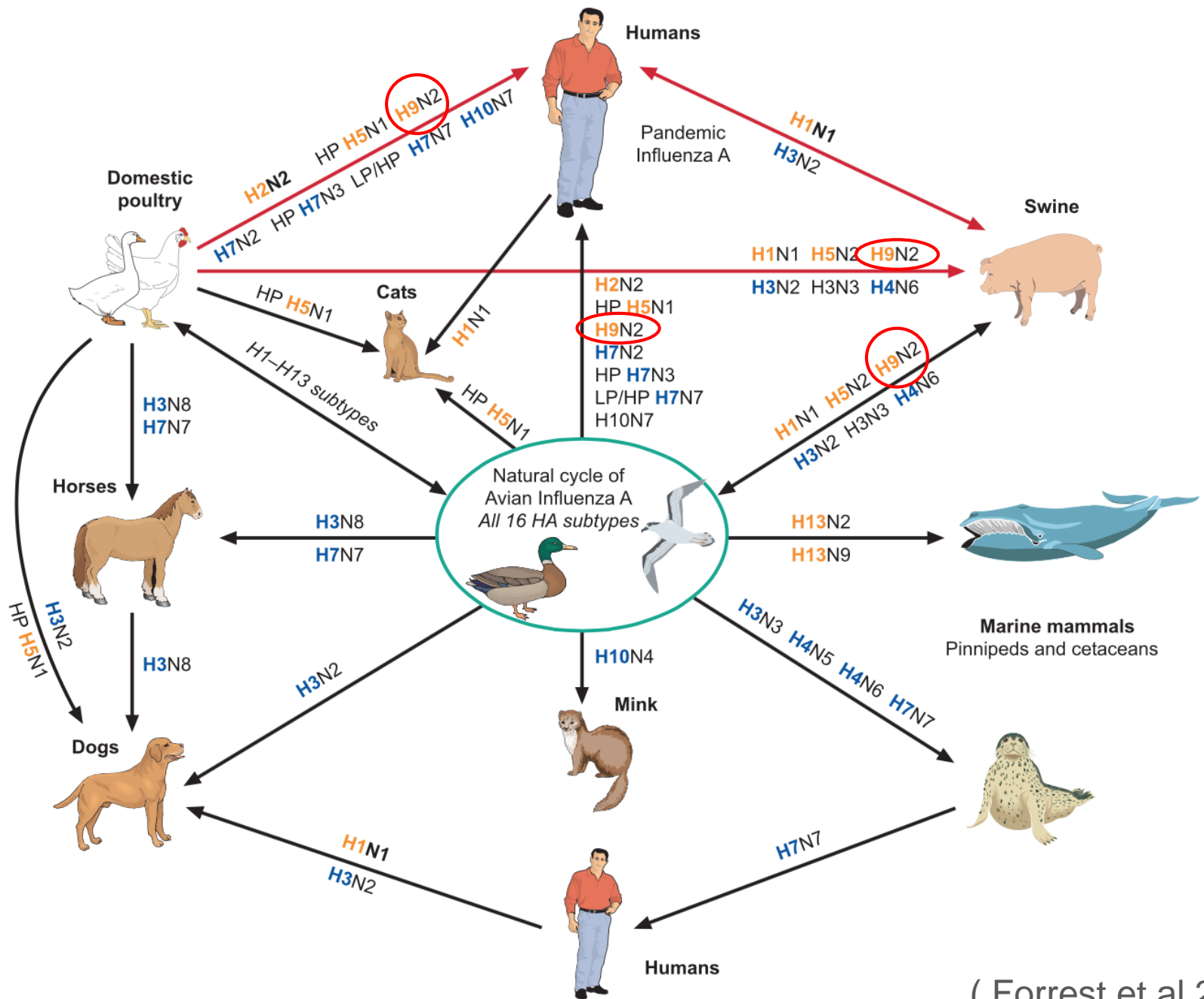


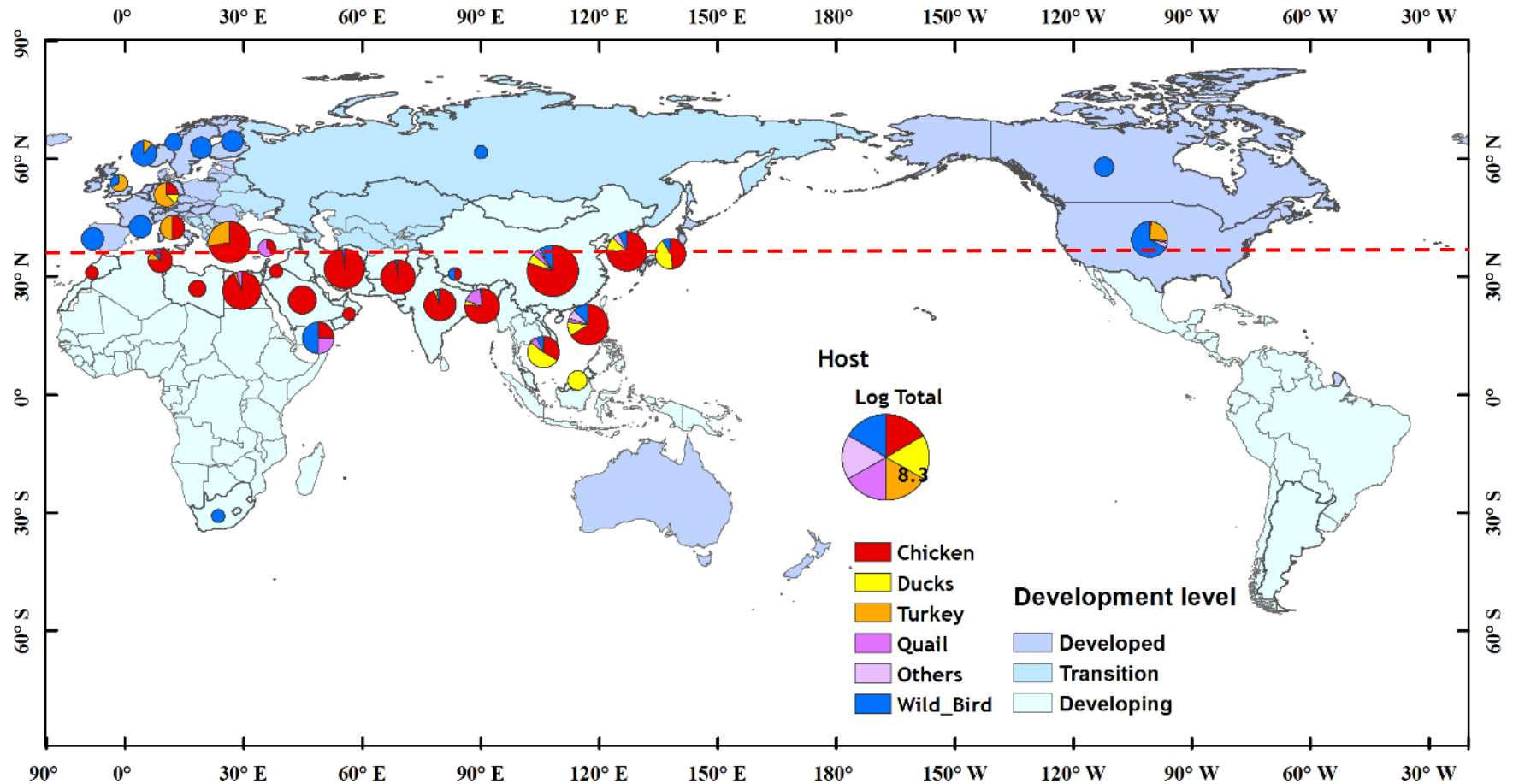
Modeling the global dynamics of avian influenza H9N2

Jing Yang and Alexei Drummond
Center for Computational Evolution,
University of Auckland

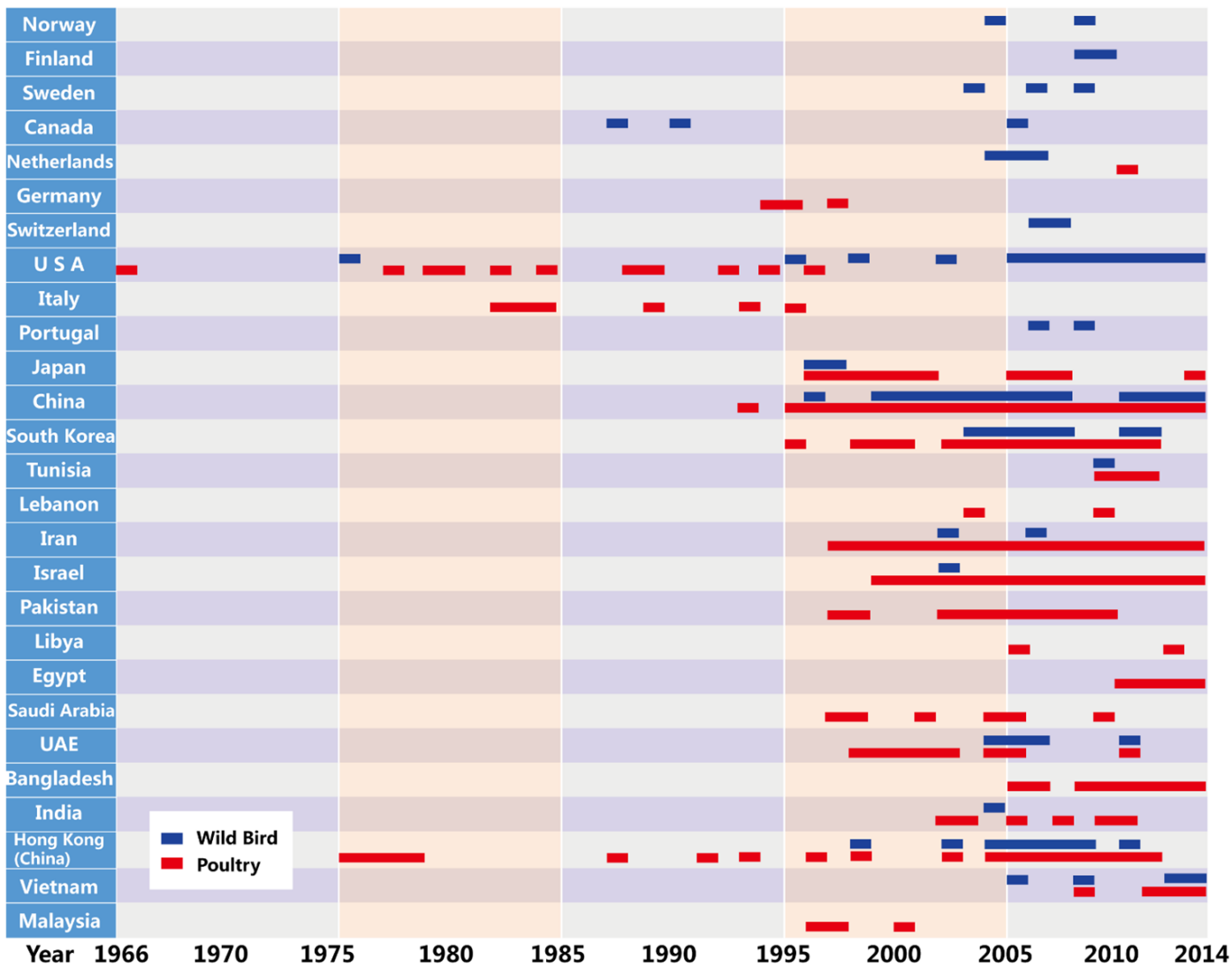


(Forrest et al 2010)

Spatial patterns



Spatial patterns



1. Since the mid 1990s, H9N2 began to spread into multiple countries, especially in Asia.
2. Latitude with different hosts

(y axis – countries were listed from high to low latitude in north hemisphere)

BEAST: Bayesian evolutionary analysis by sampling trees

Alexei J Drummond^{*1,2} and Andrew Rambaut³

BMC Evol Biol,
2007

Bayesian Phylogenetics with BEAUti and the BEAST 1.7

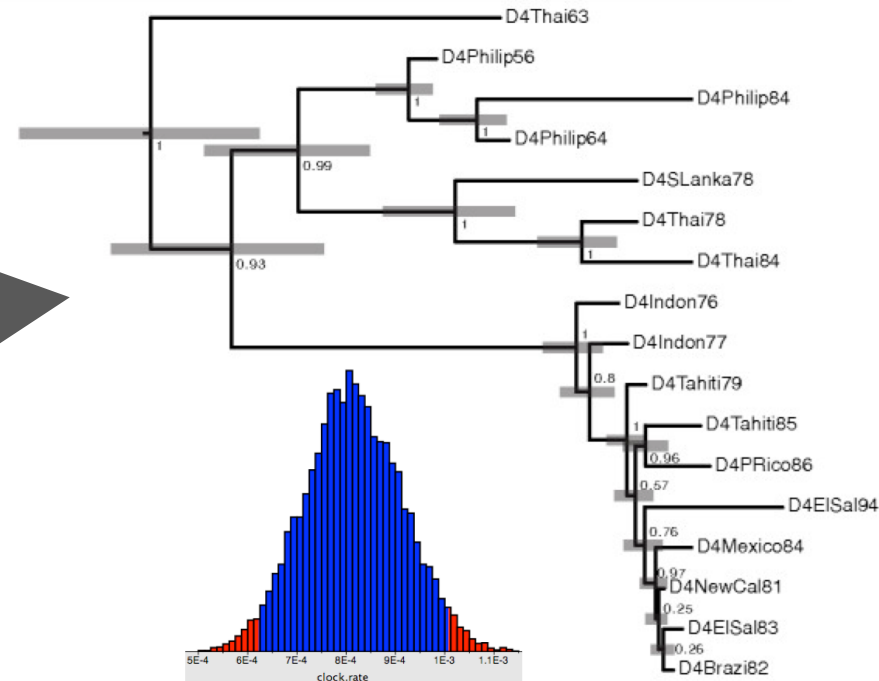
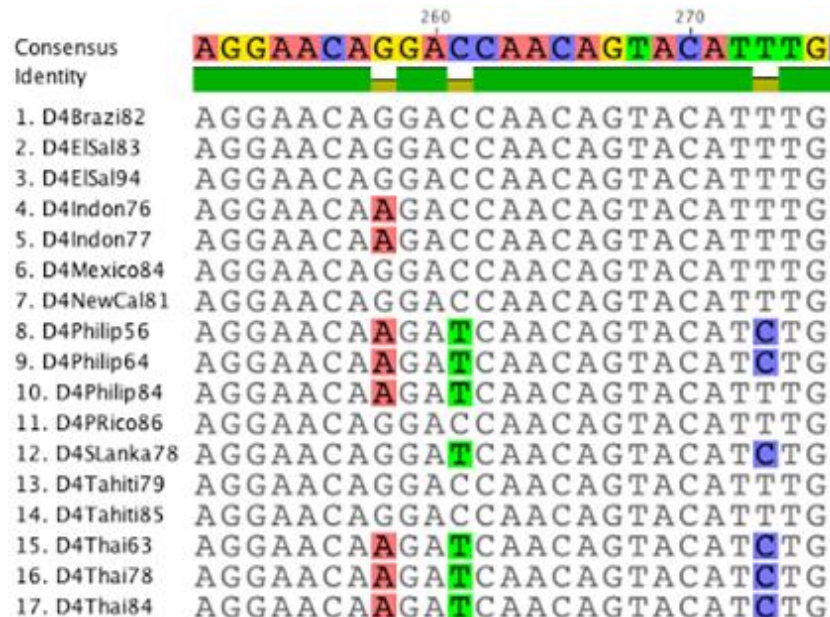
Mol Biol Evol, 2012

Alexei J. Drummond,^{*1,2} Marc A. Suchard,^{*3,4} Dong Xie,^{1,2} and Andrew Rambaut^{*5}

BEAST 2: A Software Platform for Bayesian Evolutionary Analysis

PLoS Comp Bio, 2014

Remco Bouckaert^{1*}, Joseph Heled¹, Denise Kühnert^{1,2}, Tim Vaughan^{1,3}, Chieh-Hsi Wu¹, Dong Xie¹, Marc A. Suchard^{4,5}, Andrew Rambaut⁶, Alexei J. Drummond^{1,7*}

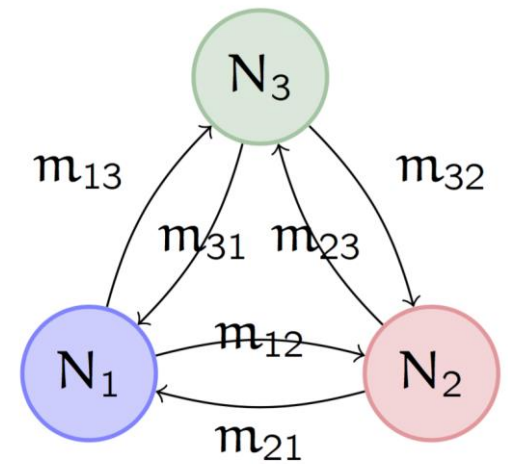
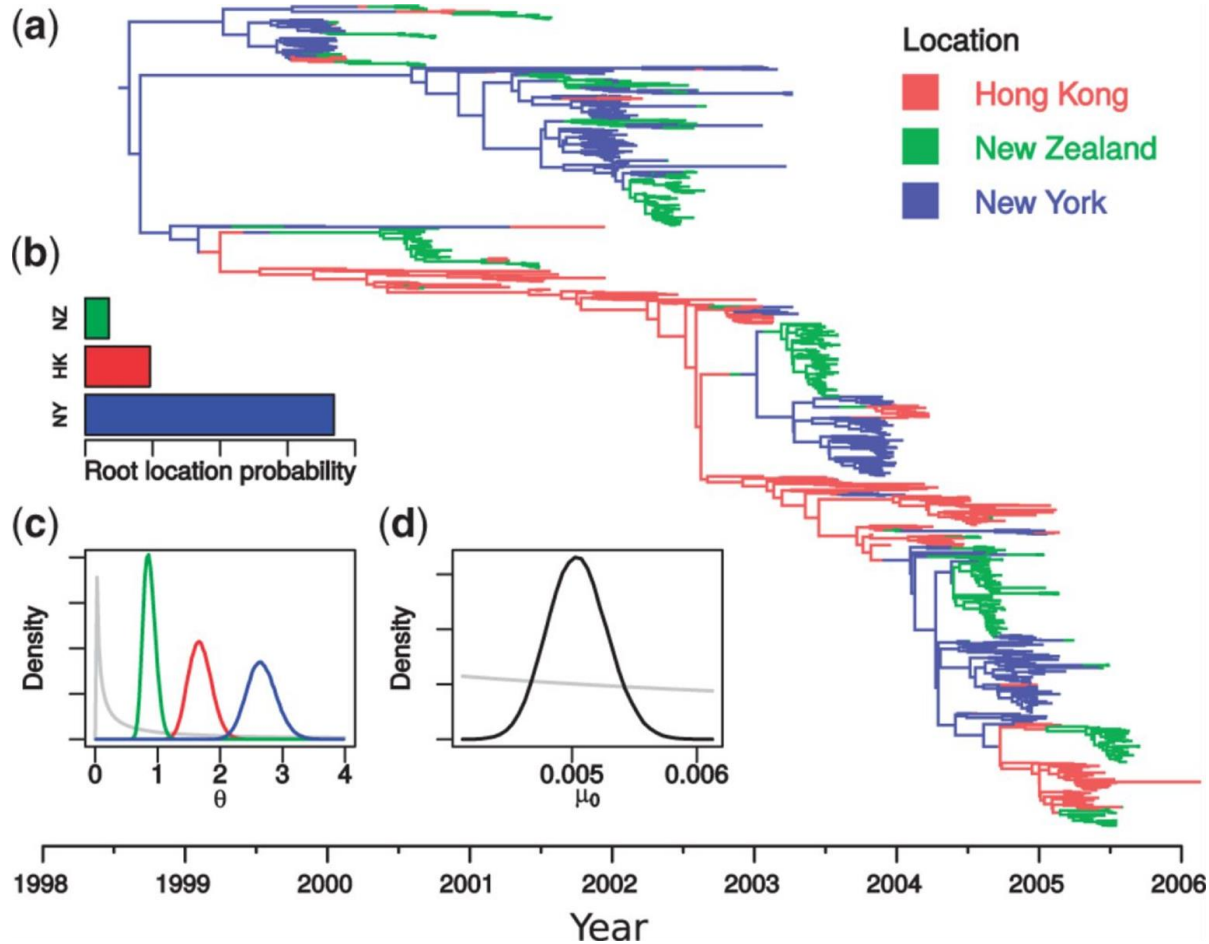


>15,000 scientific studies have used this software since 2007;
> 2,500 studies this year alone (2017)

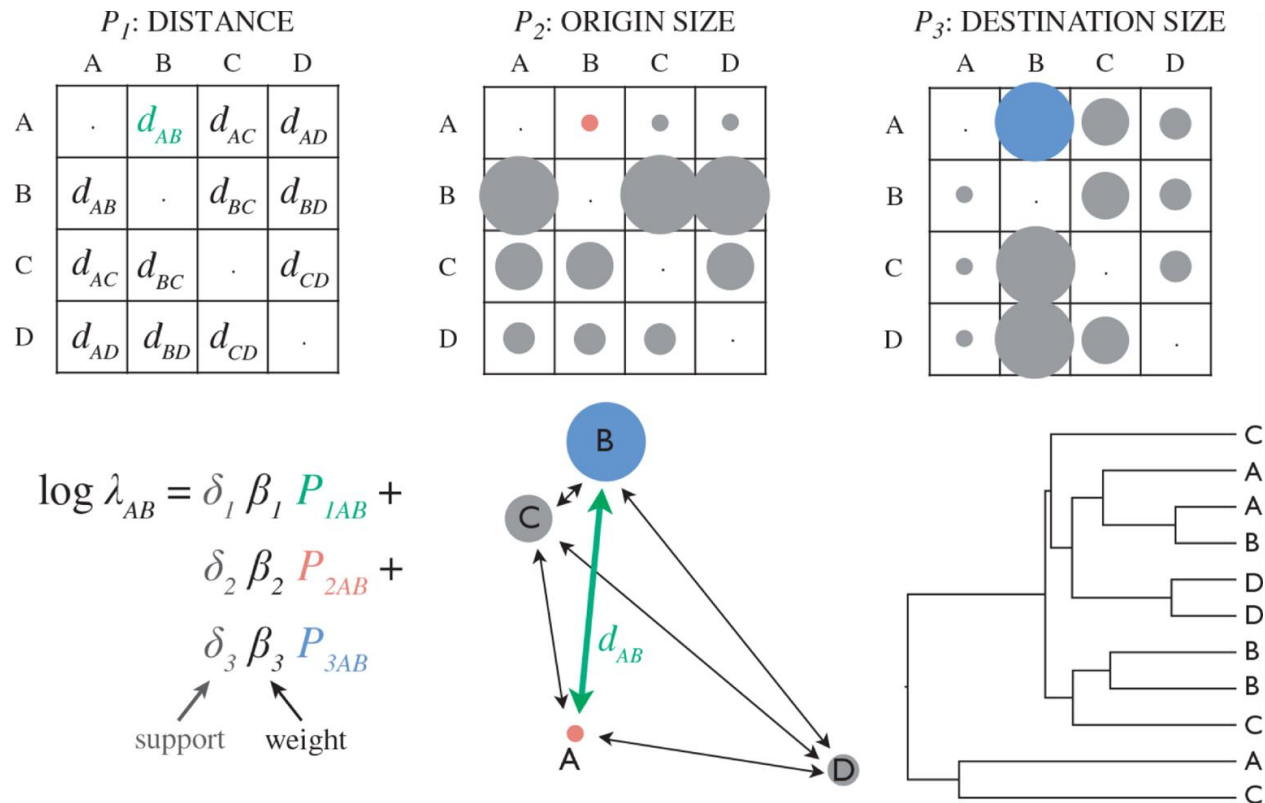
Evolution is happening right now!



Structured coalescent model



General linear models for molecular evolutionary data



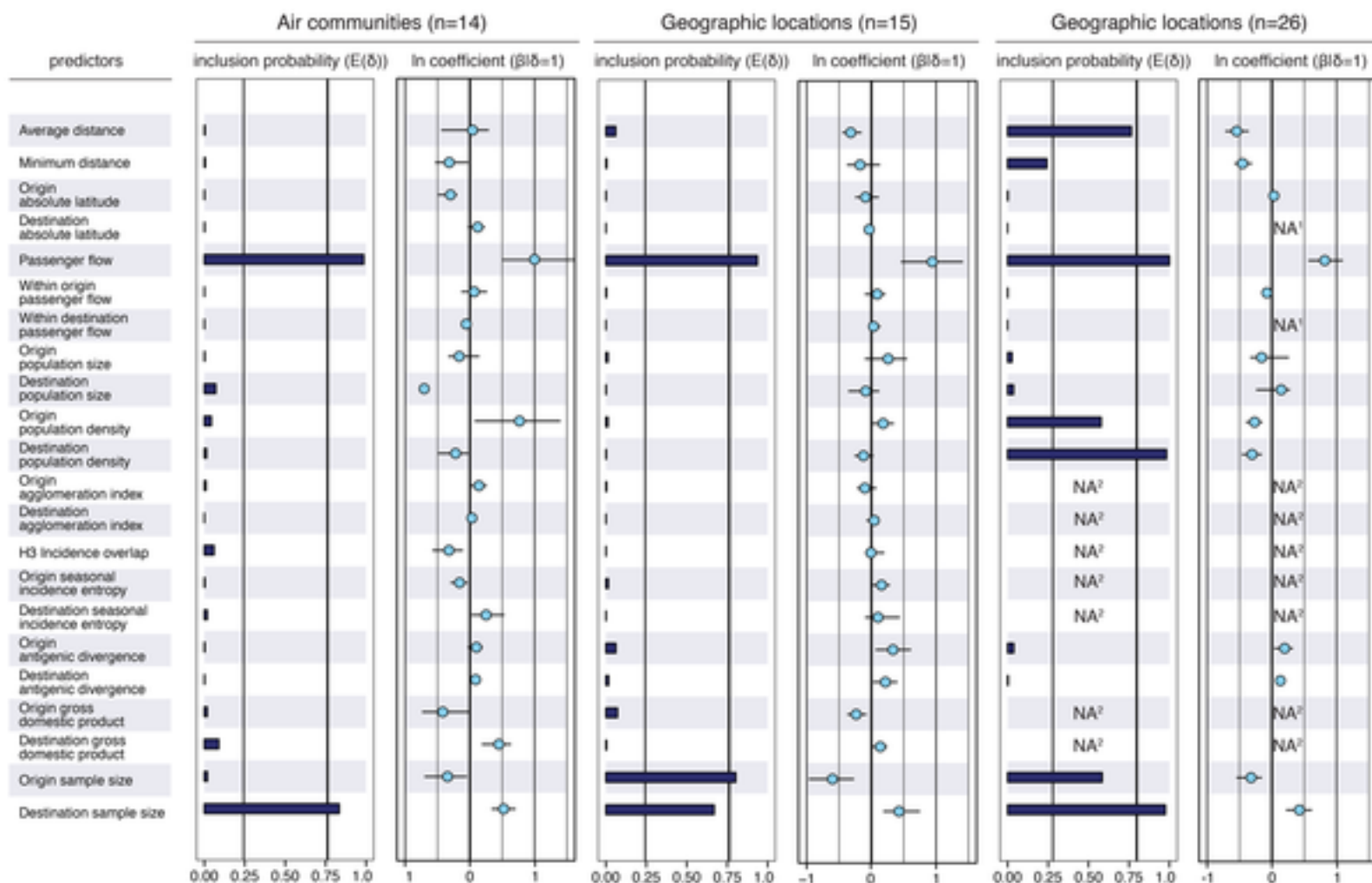
From: Emerging Concepts of Data Integration in Pathogen Phylodynamics

Syst Biol. 2016;66(1):e47-e65. doi:10.1093/sysbio/syw054

Syst Biol | © The Author(s) 2016. Published by Oxford University Press, on behalf of the Society of Systematic Biologists. This is an Open Access article distributed under the terms of the Creative Commons Attribution License

(<http://creativecommons.org/licenses/by/4.0/>), which permits unrestricted reuse, distribution, and reproduction in any medium, provided the original work is properly cited.

Figure 2. Predictors of global H3N2 diffusion among the 14 air communities and the 15 & 26 geographic locations.

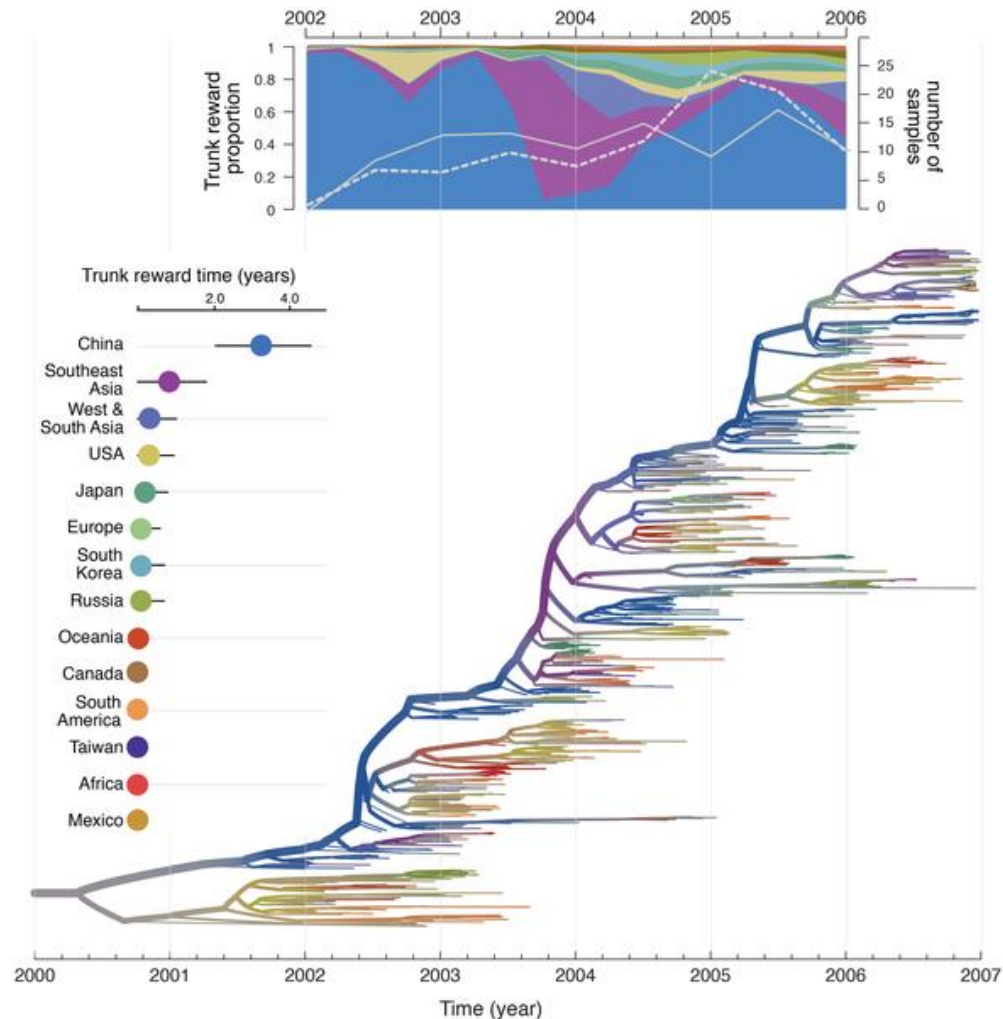


Lemey P, Rambaut A, Bedford T, Faria N, Bielejec F, et al. (2014) Unifying Viral Genetics and Human Transportation Data to Predict the Global Transmission Dynamics of Human Influenza H3N2. PLOS Pathogens 10(2): e1003932.

<https://doi.org/10.1371/journal.ppat.1003932>

<http://journals.plos.org/plospathogens/article?id=10.1371/journal.ppat.1003932>

Figure 3. Phylogeographic reconstruction and spatial history of the trunk lineage.



Lemey P, Rambaut A, Bedford T, Faria N, Bielejec F, et al. (2014) Unifying Viral Genetics and Human Transportation Data to Predict the Global Transmission Dynamics of Human Influenza H3N2. PLOS Pathogens 10(2): e1003932.

<https://doi.org/10.1371/journal.ppat.1003932>

<http://journals.plos.org/plospathogens/article?id=10.1371/journal.ppat.1003932>

Hypothesis

Poultry trade and poultry production will be predictive
of the spatial population dynamics of avian influenza
H9N2

Data

Global avian-origin H9N2 AIV full length HA genes:

n: curated 647 sequences

L: 1683 nucleotides

(East Asia: 288; South Asia: 75; Southeast Asia: 23; West Asia: 149;
Europe:34; North America:47; Africa: 31)

Asian avian-origin H9N2 AIV HA genes: Total: 535 sequences

GenBank Influenza Virus Database

(<http://www.ncbi.nlm.nih.gov/genomes/FLU/FLU.html>)

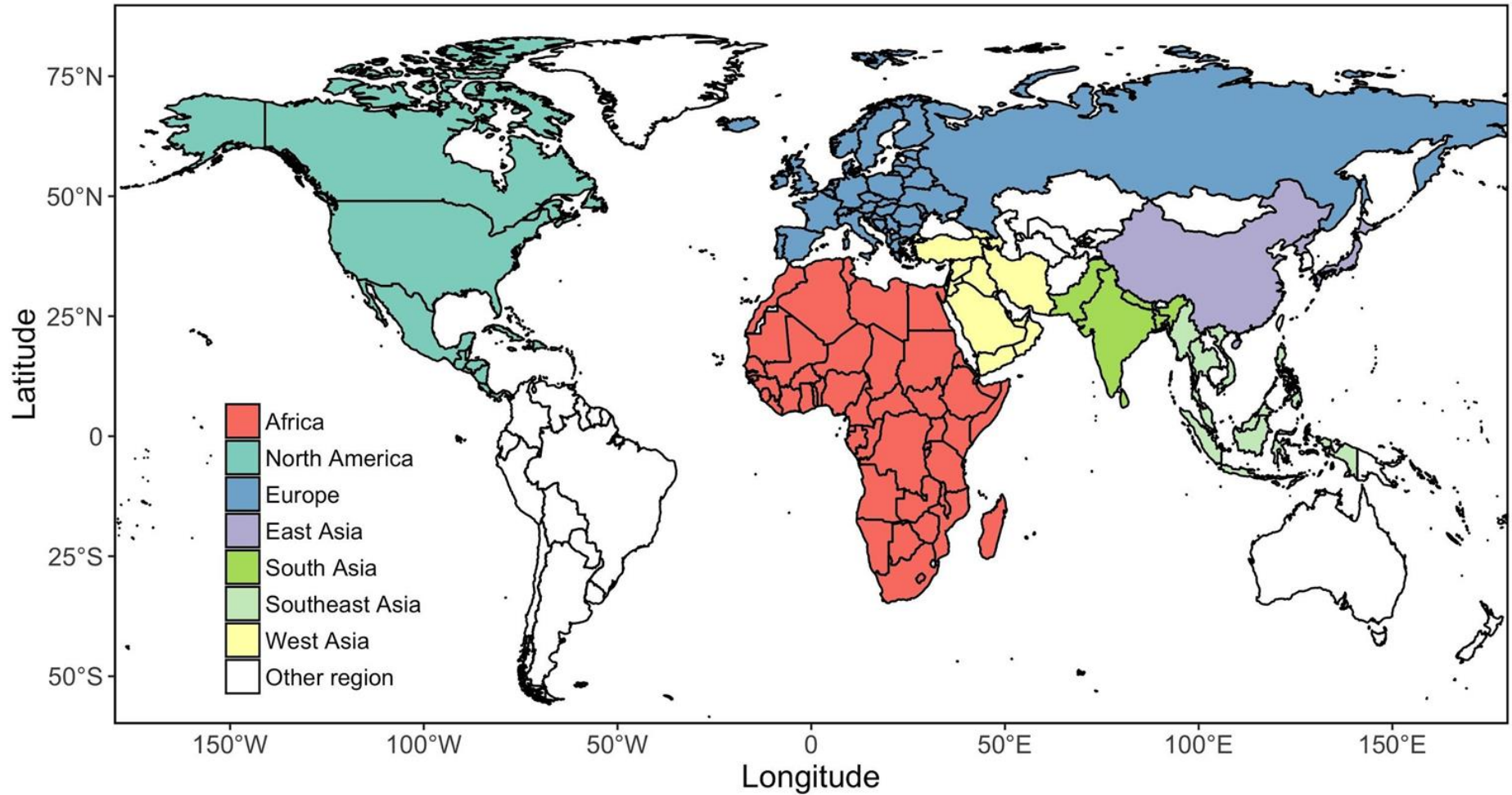
Poultry production and trade data:

Poultry production: 1986-2013

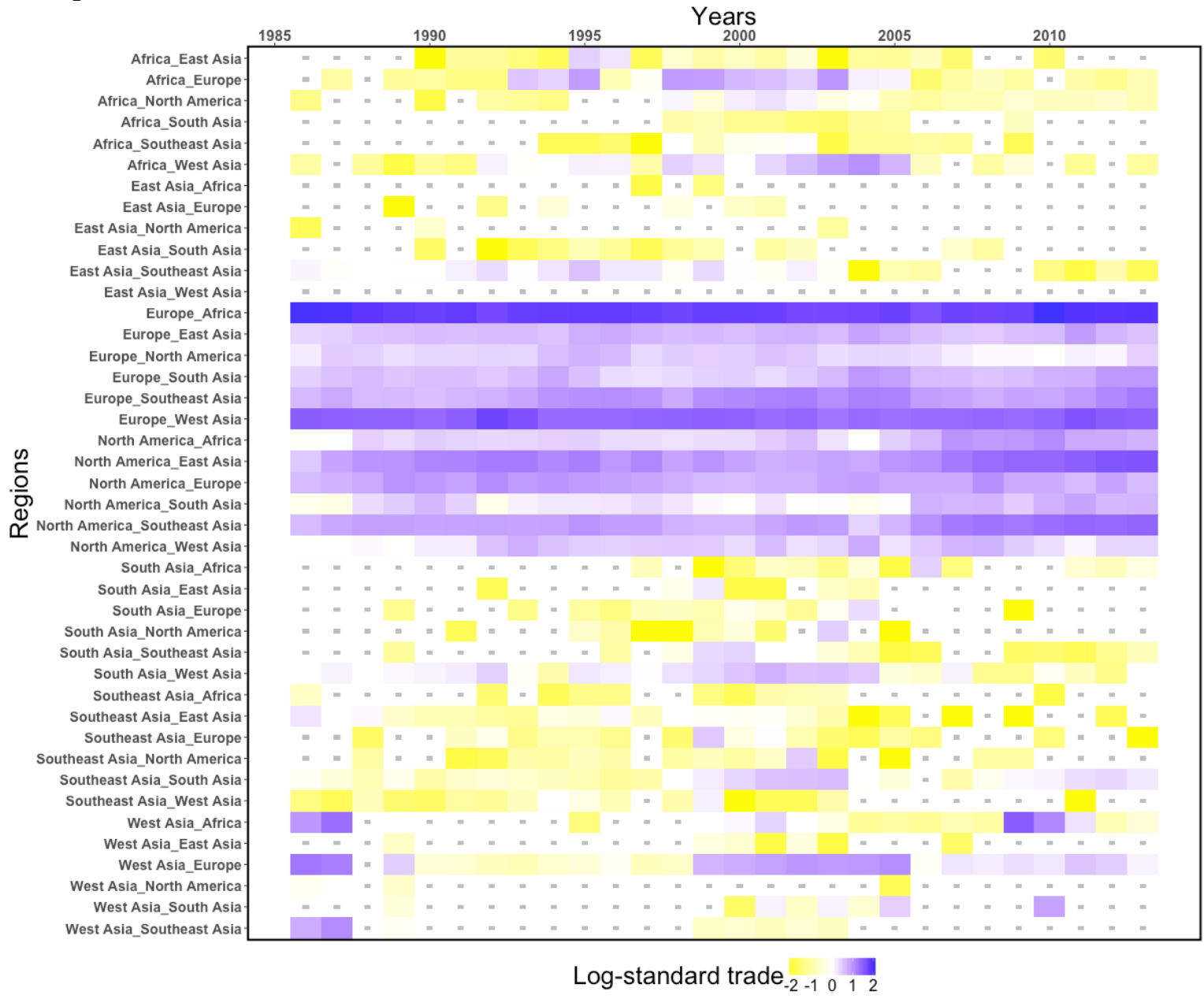
Poultry trade: 1986-2013

FAOSTAT (<http://faostat3.fao.org>)

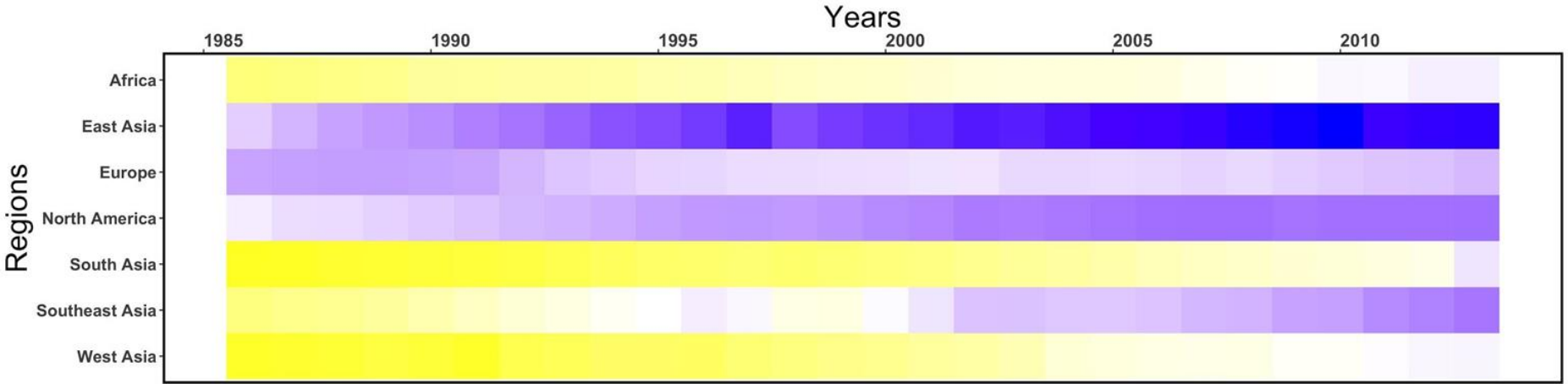
Regions



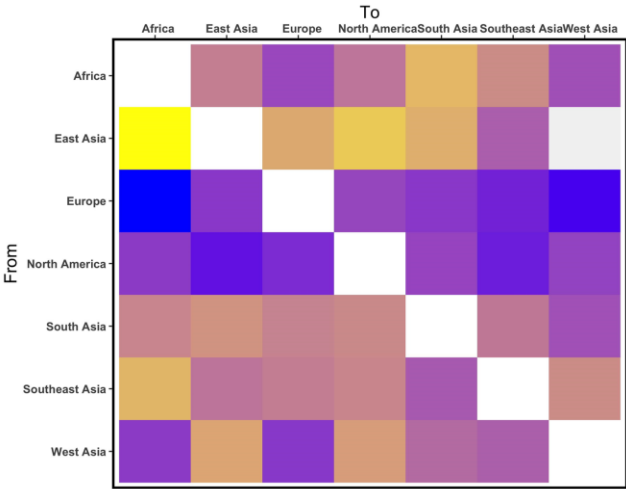
Poultry trade around the world



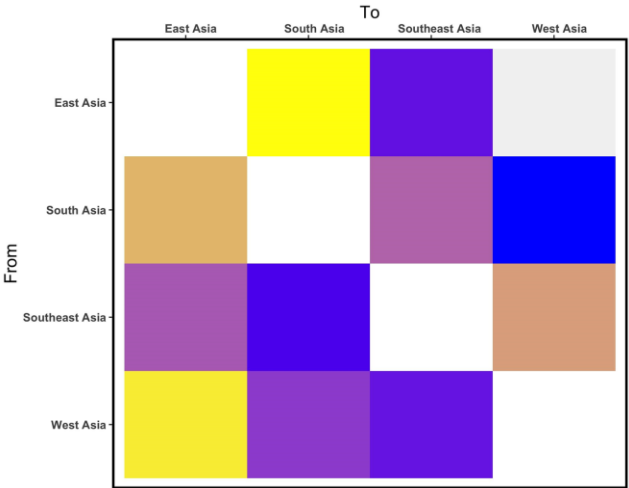
Poultry production around the world



Log-standard production -1 0 1 2



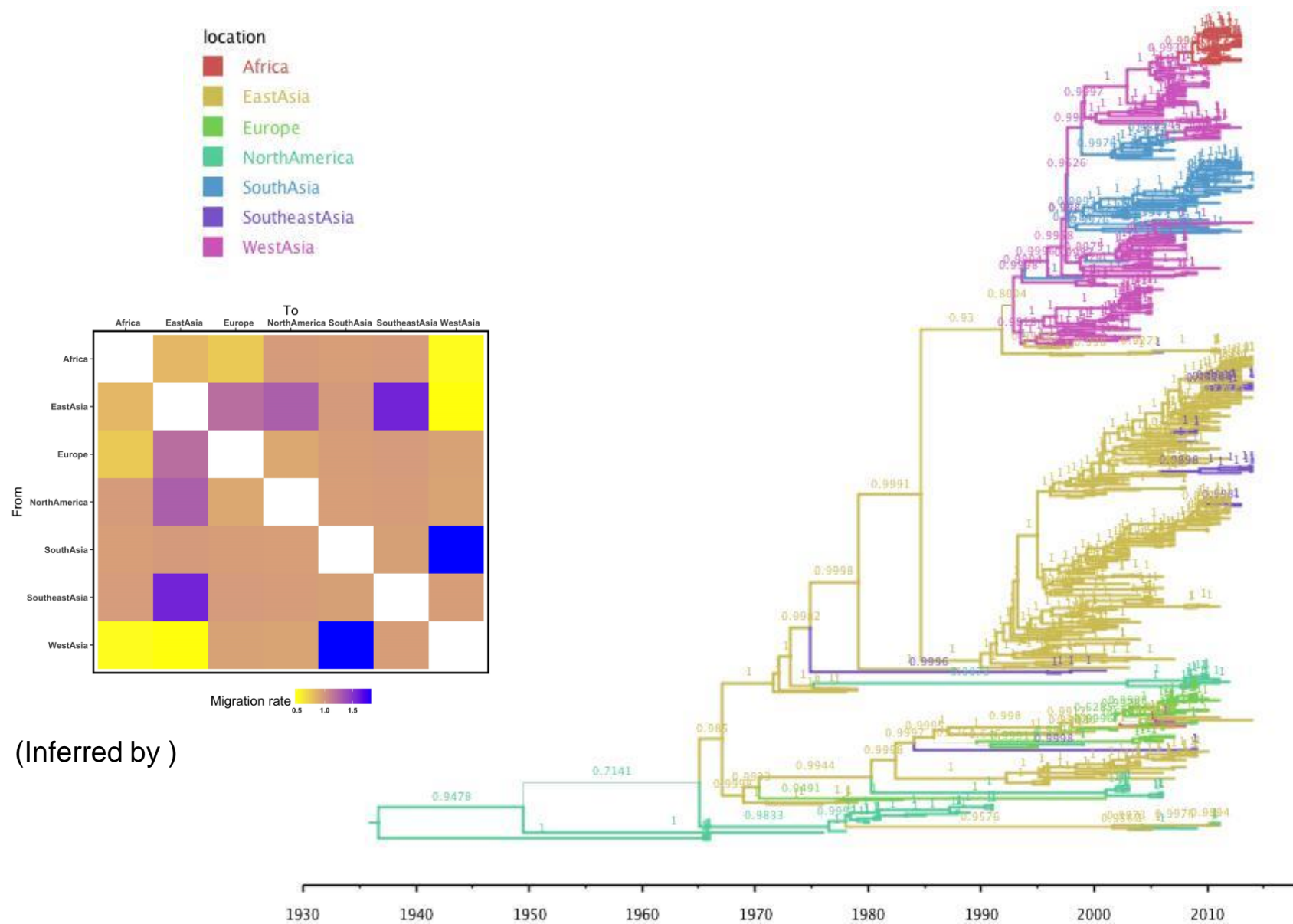
Log-standard average trade -2 -1 0 1 2



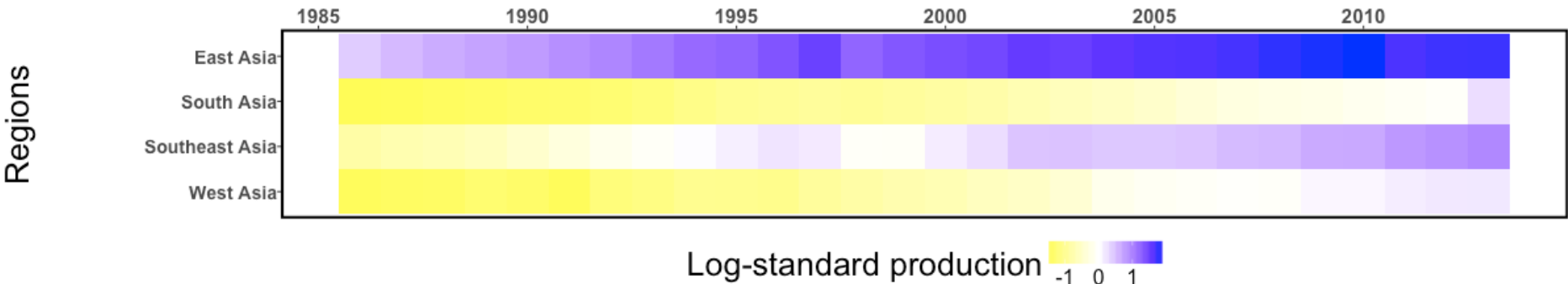
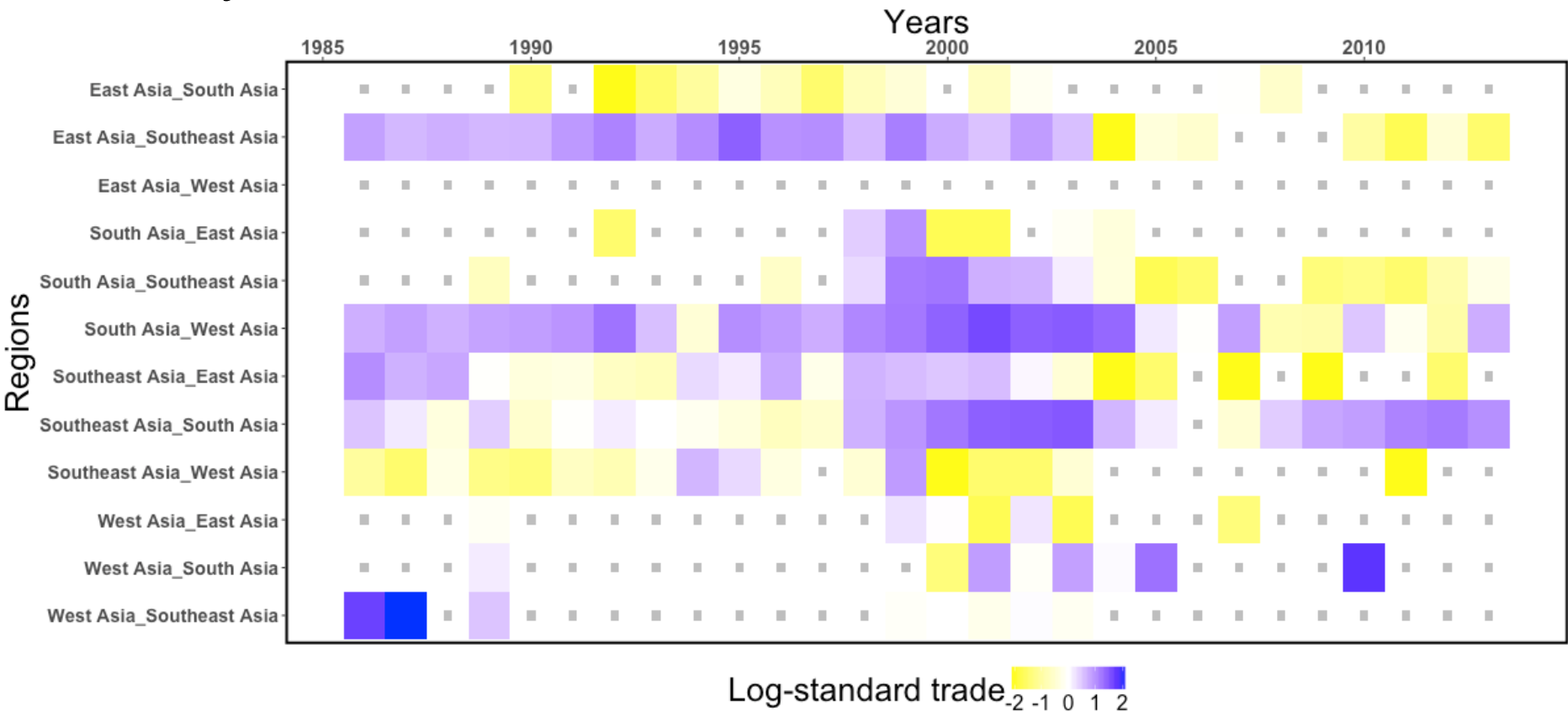
Log-standard average trade -1 0 1

average poultry trade over 28 years on global and asian scale.

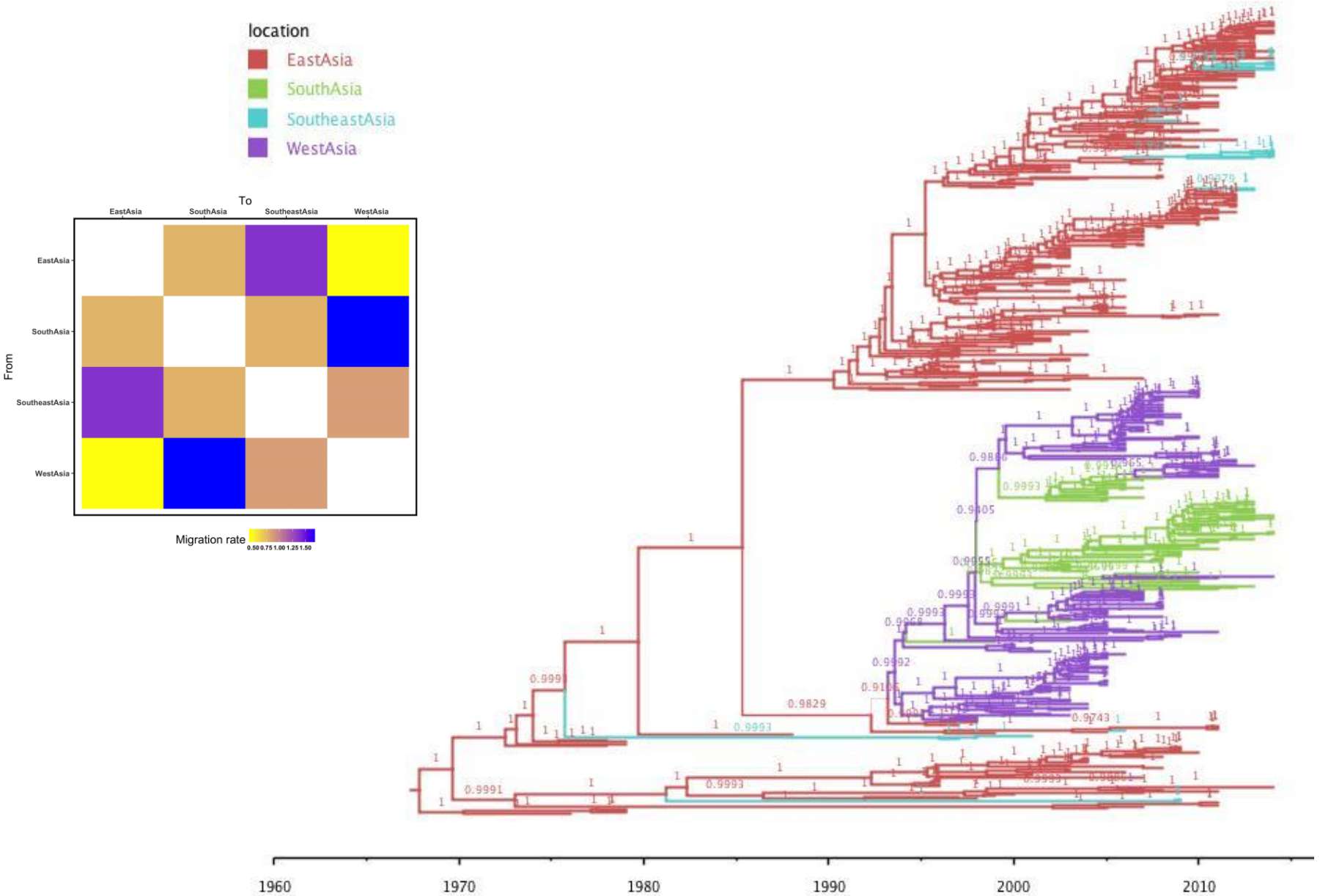
Time-scaled tree of global



Poultry trade in Asia



Time-scaled tree in Asia



Statistical support for poultry trade as explanatory variable

Model	Asia	
	Poultry trade	Poultry production
No covariates / non-structured	No formal test	N/A
Time-averaged covariates / non-structured population model	++	N/A
Time-averaged covariates / structured population model	--	++
Full temporal covariates / structured population model	++	++

Conclusions

- There is weak direct statistical support for poultry trade as a driver for the spread of H9N2 avian influenza.
- Expanding the data set from 600 -> 4000 sequences will improve power but is computationally challenging
- Model granularity in time and space can have a decisive effect on hypothesis testing.
- New methods for molecular epidemiology will increasingly be characterized by data integration efforts.
- Integration of detailed covariate data with molecular epidemiological analyses will allow formal testing of drivers.