

**What genomic data could tell us**

**...through phylodynamics**

**Philippe Lemey**

**Rega Institute,**

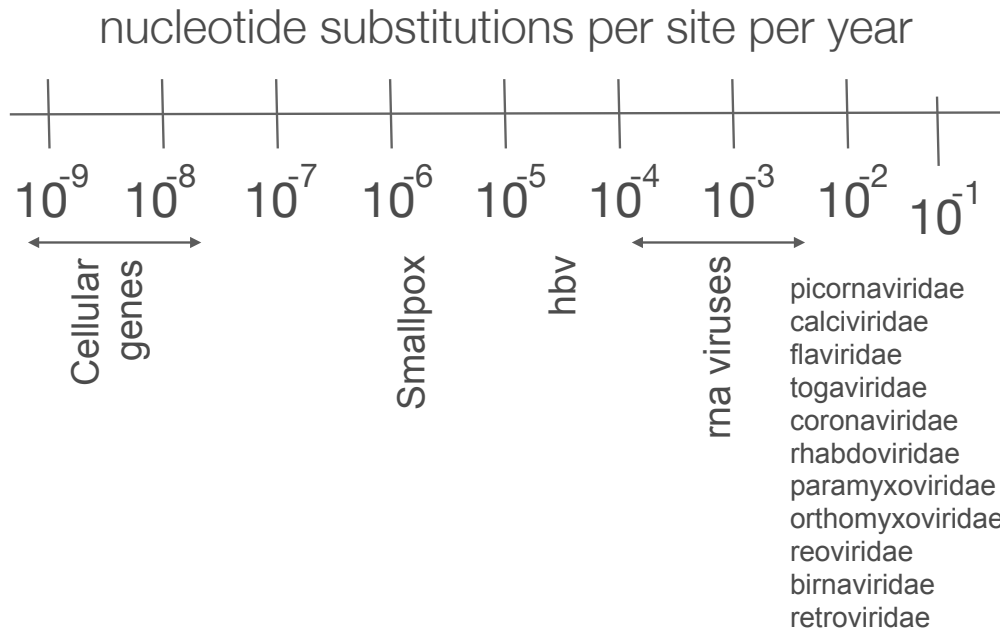
**Department of Microbiology,  
Immunology and Transplantation**

**K.U. Leuven, Belgium.**

# Unifying the epidemiological and evolutionary dynamics of pathogens

“Rapidly evolving pathogens are unique in that their ecological and evolutionary dynamics occur on the same timescale and can therefore potentially interact.”

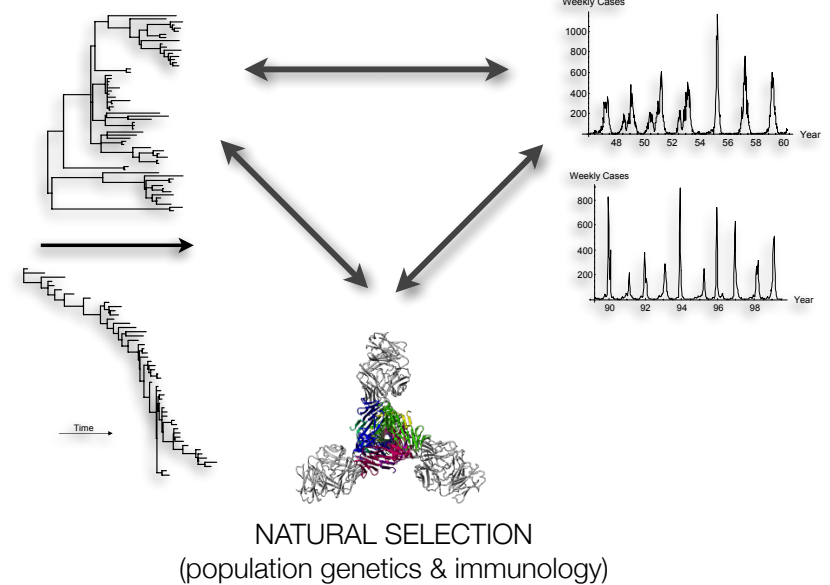
Pybus & Rambaut (2009) Nat. Rev. Genetics 10:540-50



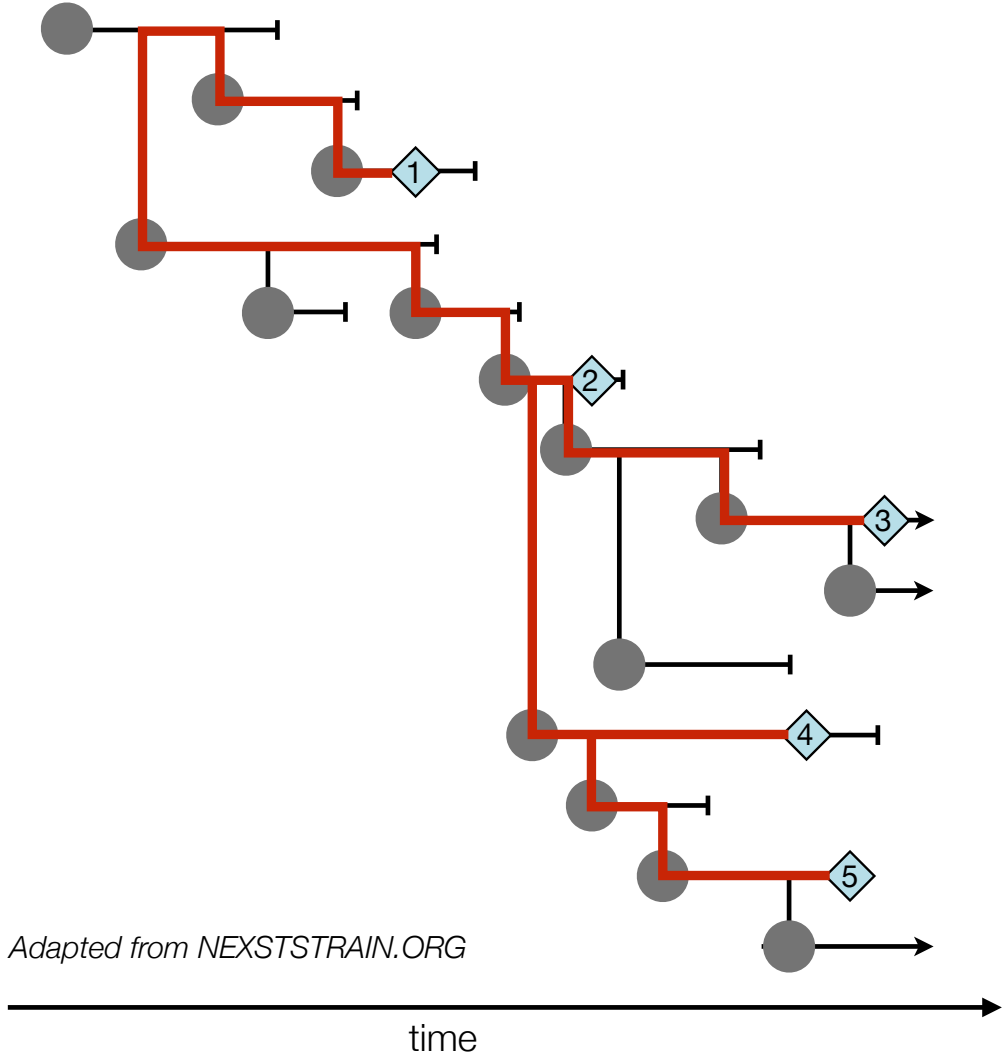
Grenfell (2004) Science 303: 327-30

GENETIC DIVERSITY  
(phylogenetics & molecular evolution)

EPIDEMIC DYNAMICS  
(mathematical epidemiology)

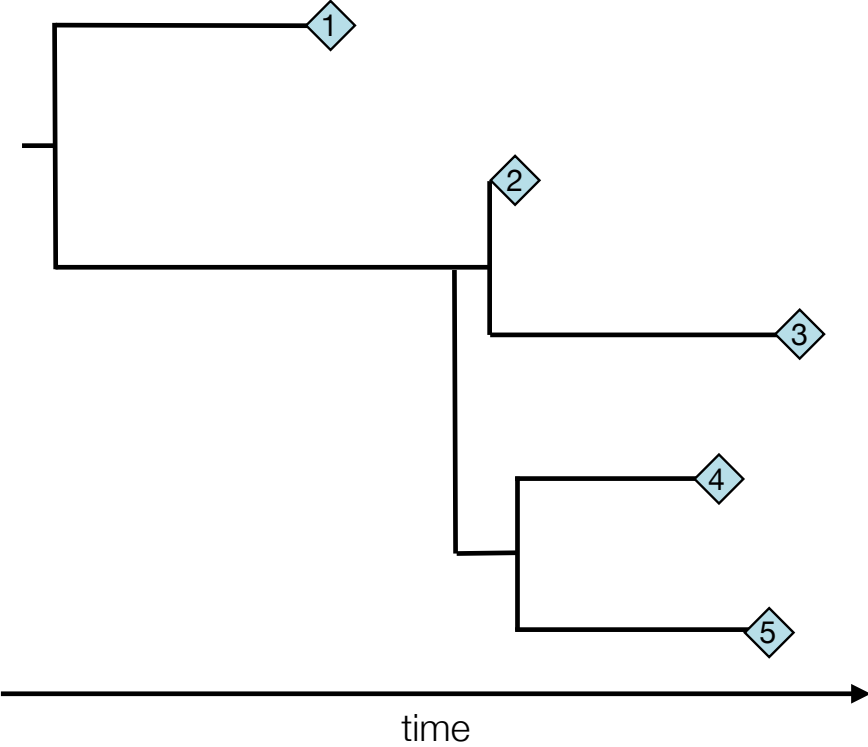


# Transmission history

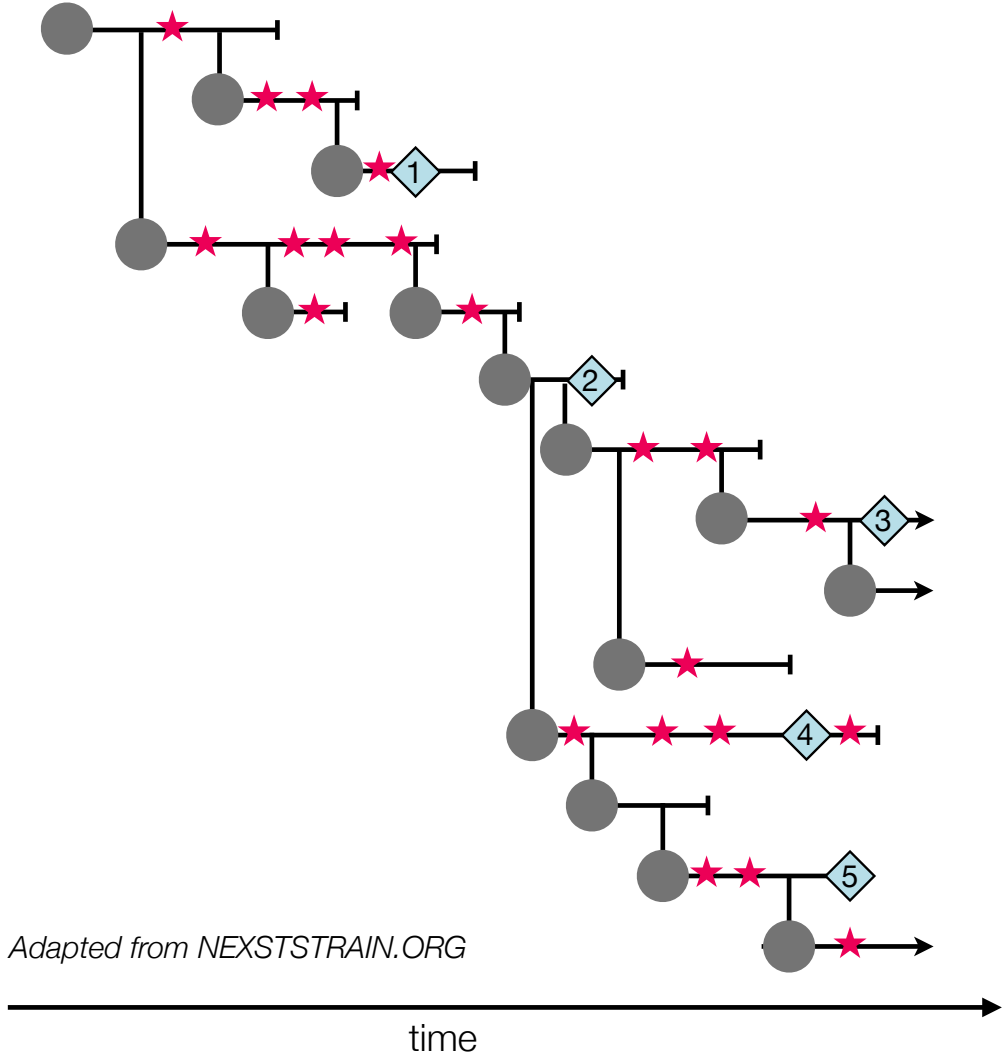


*Adapted from NEXSTSTRAIN.ORG*

# Phylogeny

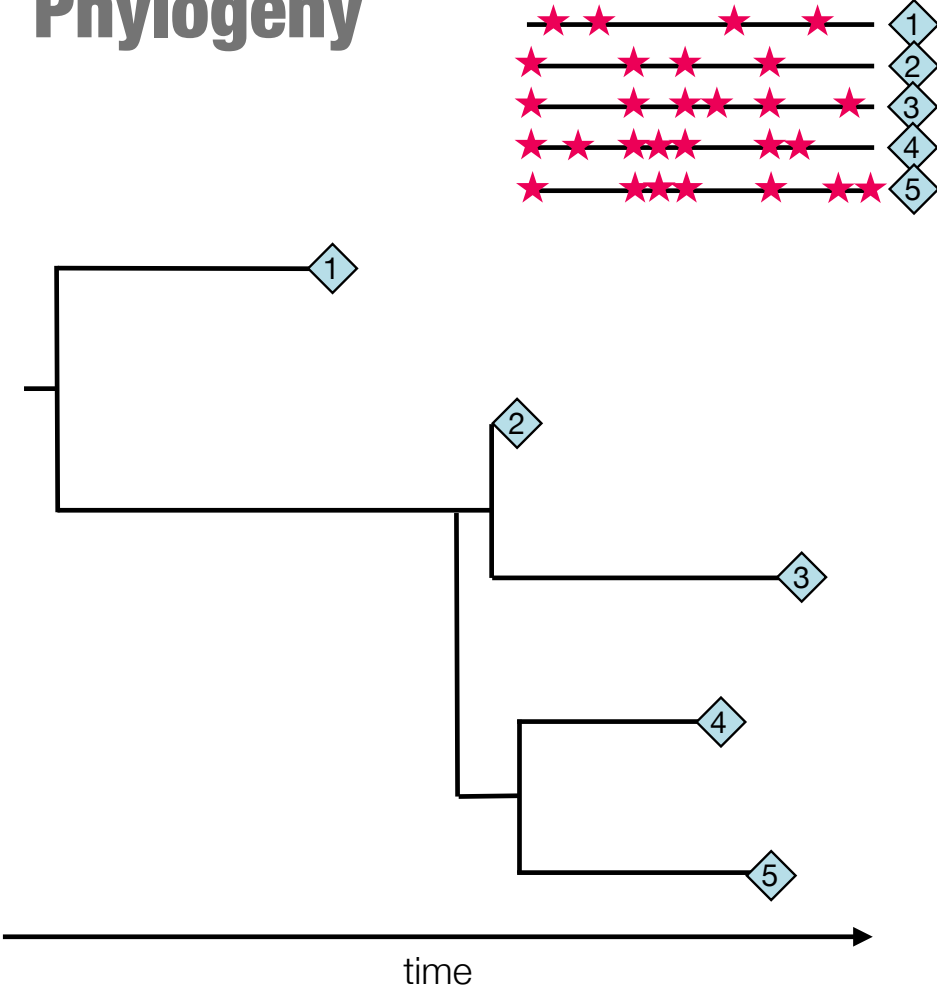


# Transmission history

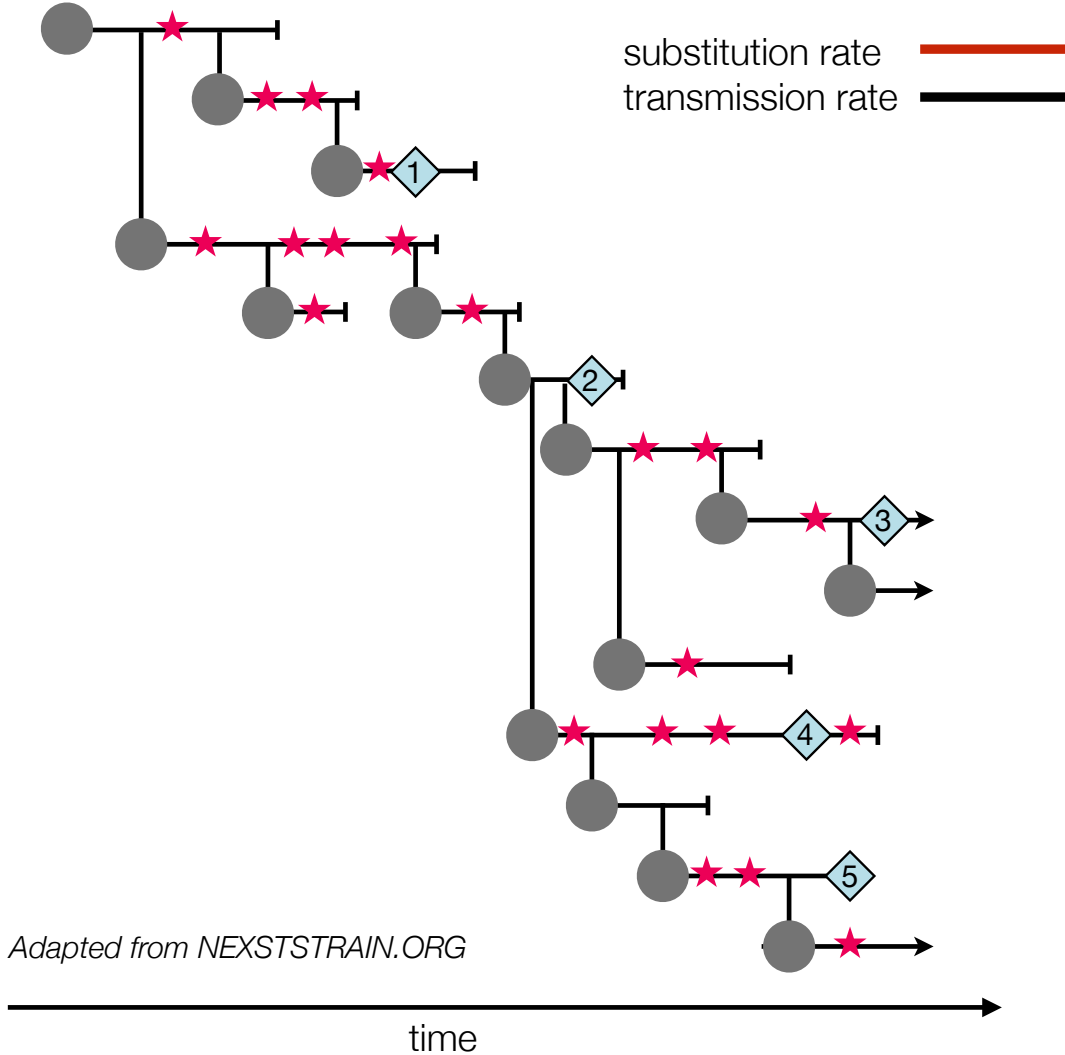


*Adapted from NEXSTSTRAIN.ORG*

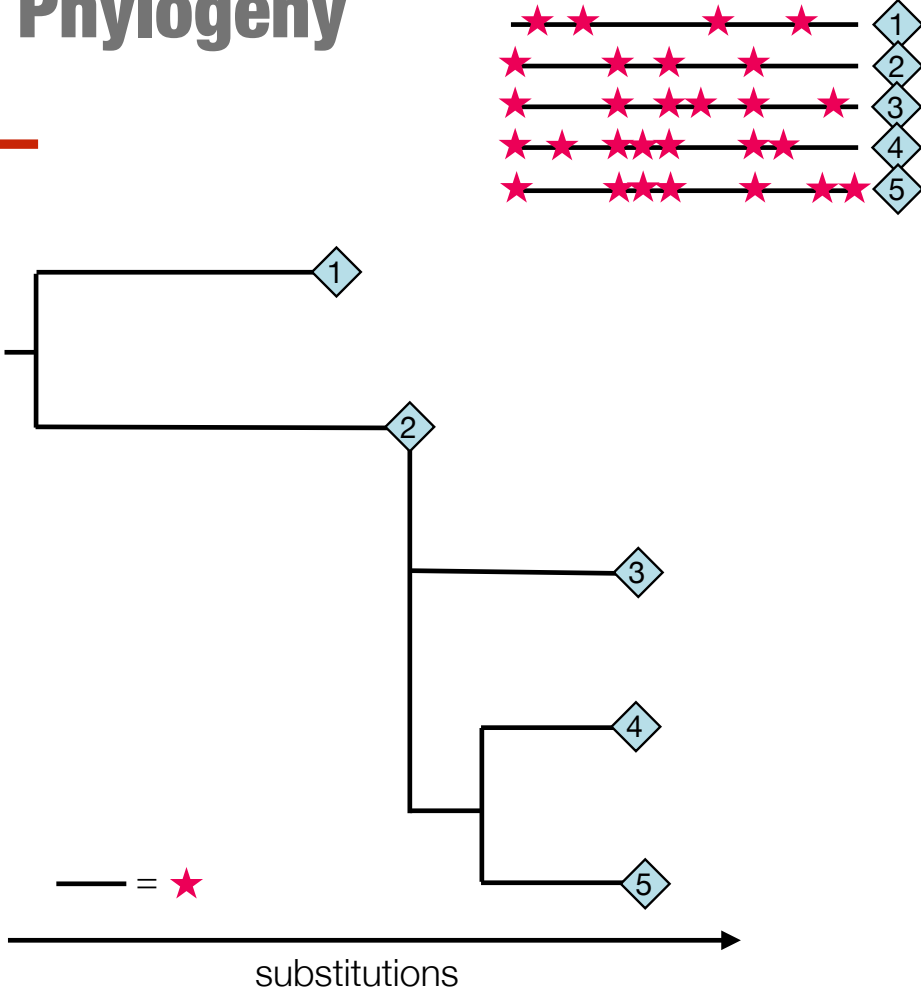
# Phylogeny



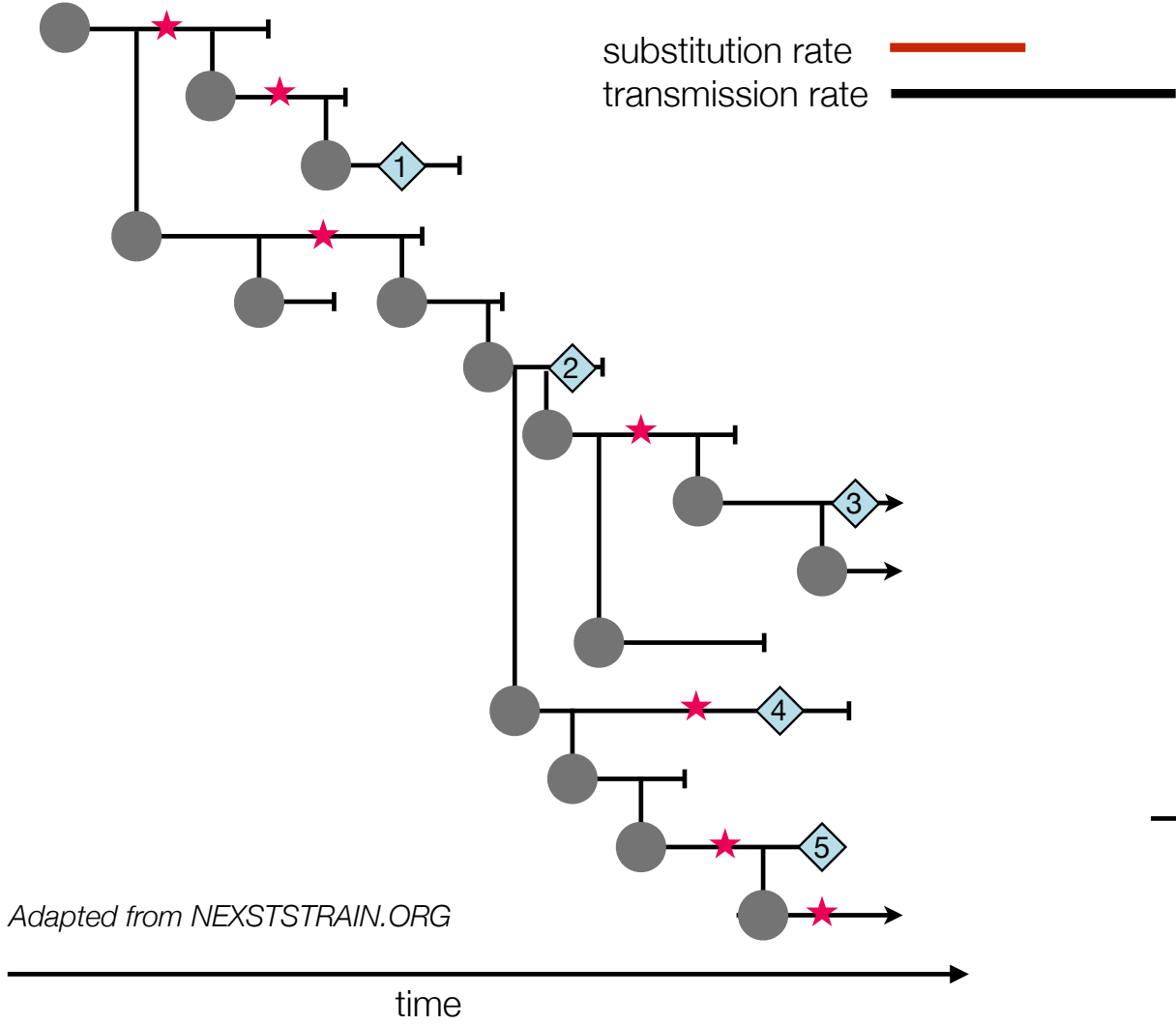
# Transmission history



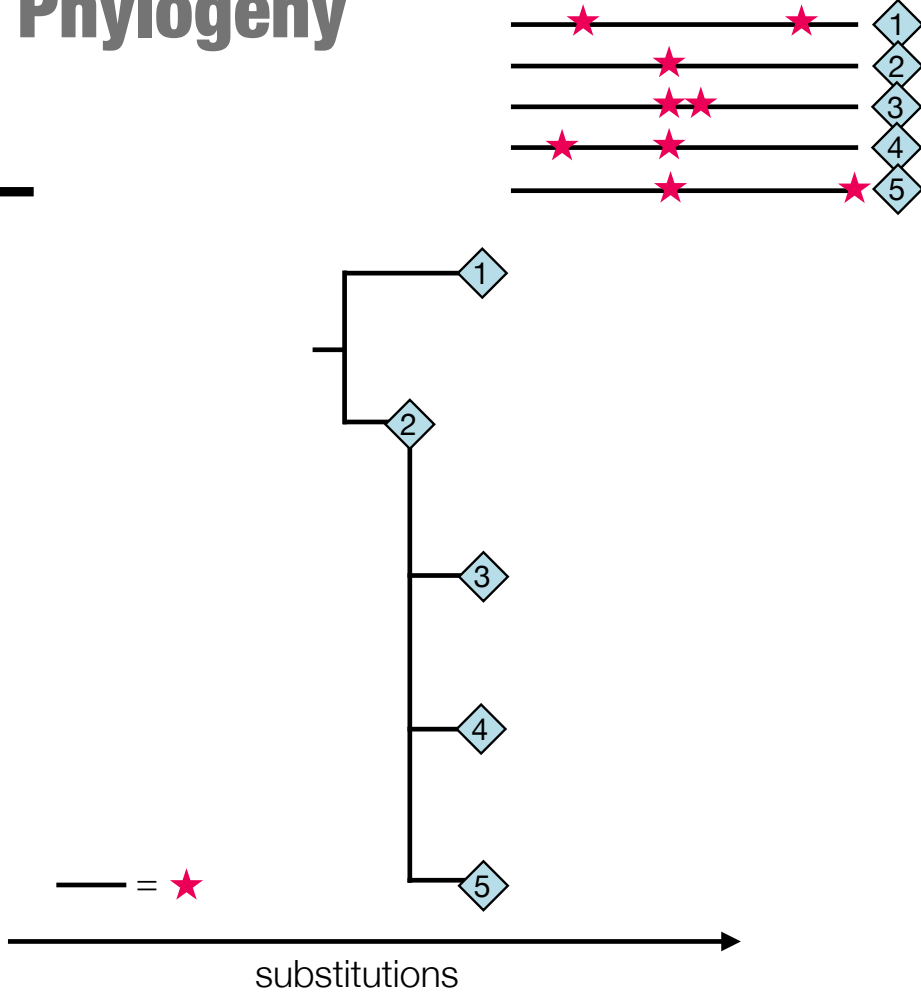
# Phylogeny



# Transmission history

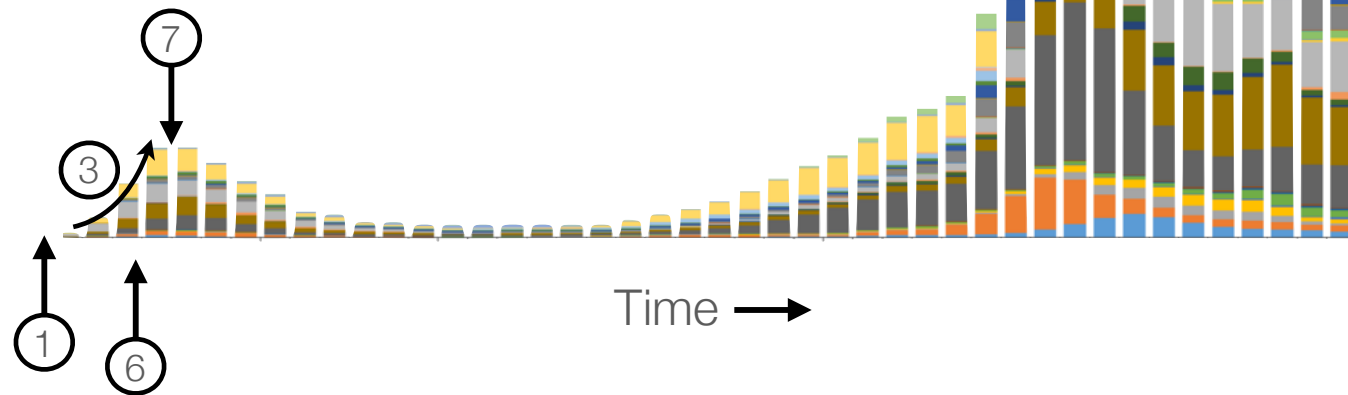


# Phylogeny



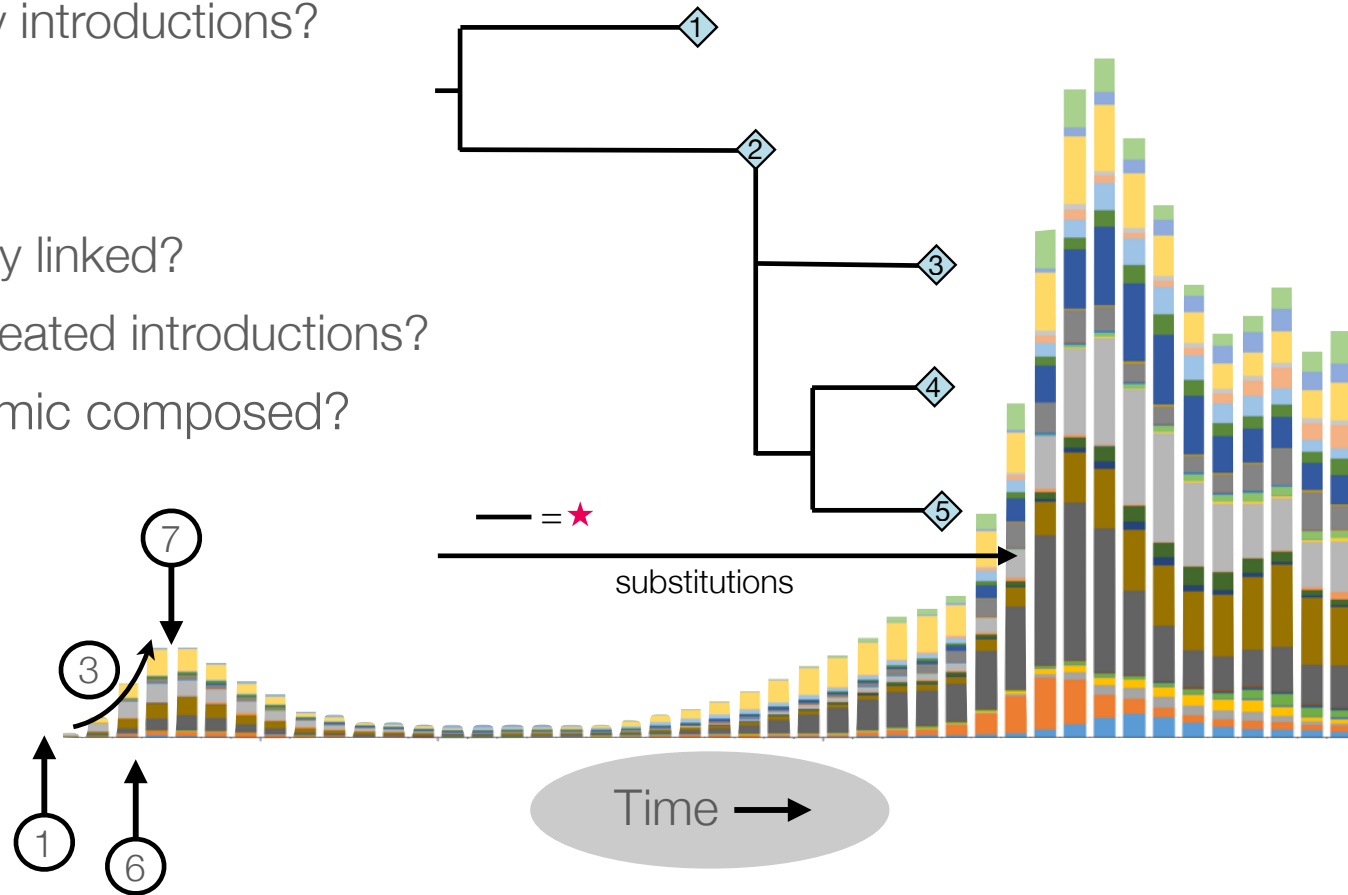
# Epidemiological questions

1. When did an epidemic start?
2. Where did it come from, how many introductions?
3. How fast is it transmitting?
4. In what direction is it spreading?
5. Are hosts X, Y & Z epidemiologically linked?
6. Local transmission or repeated repeated introductions?
7. Of how many strains is the epidemic composed?
8. What adaptations has it accrued?
9. ...



# Epidemiological questions

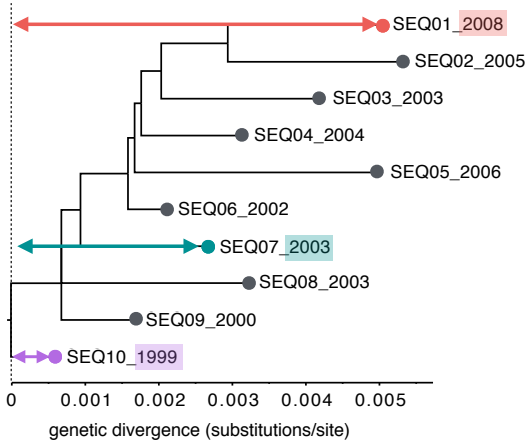
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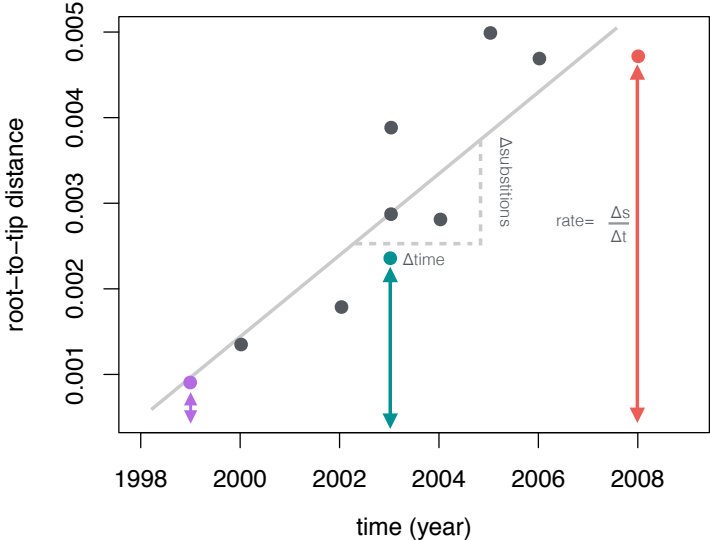


# Phylodynamic inference: time

### Substitutions

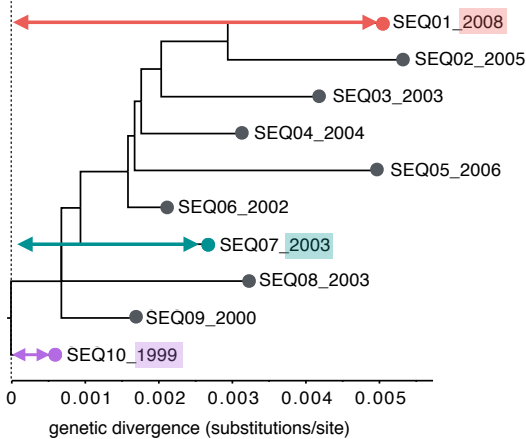


### Evolutionary rate

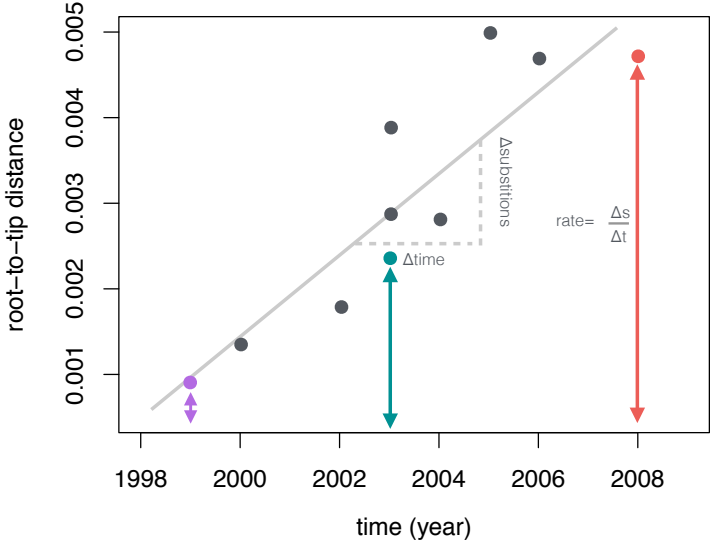


# Phylodynamic inference: time

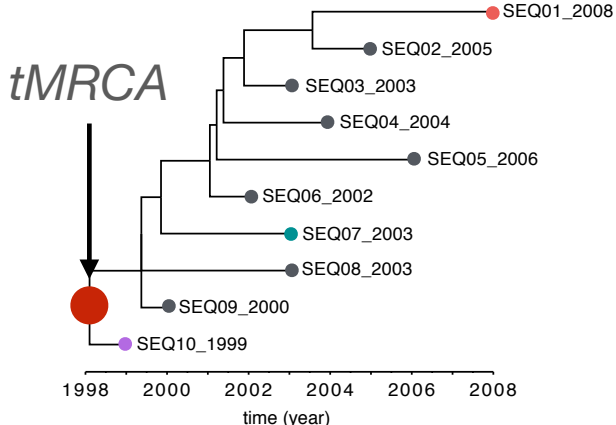
## Substitutions



## Evolutionary rate



## Time

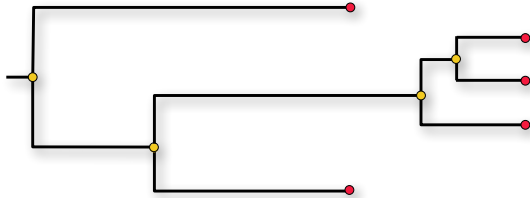
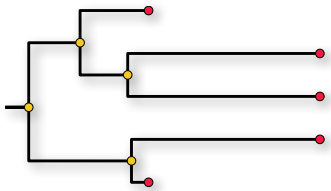


Molecular clock

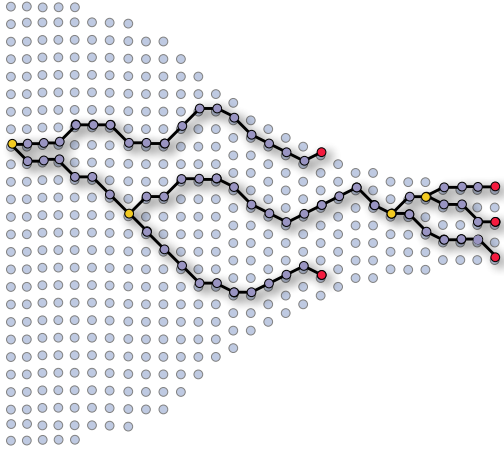
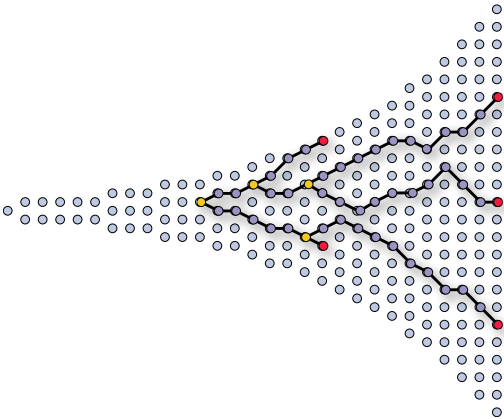




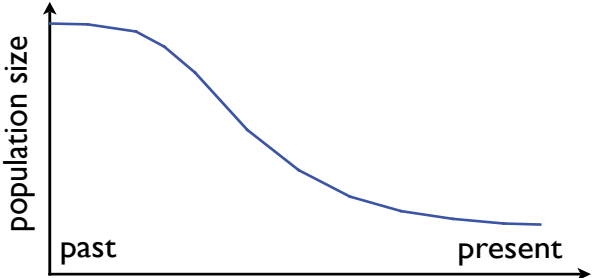
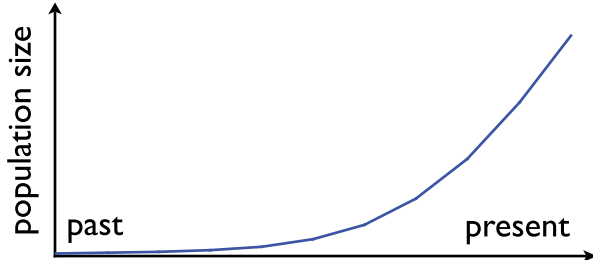
# Phylodynamic inference: the coalescent



Phylogeny or Genealogy



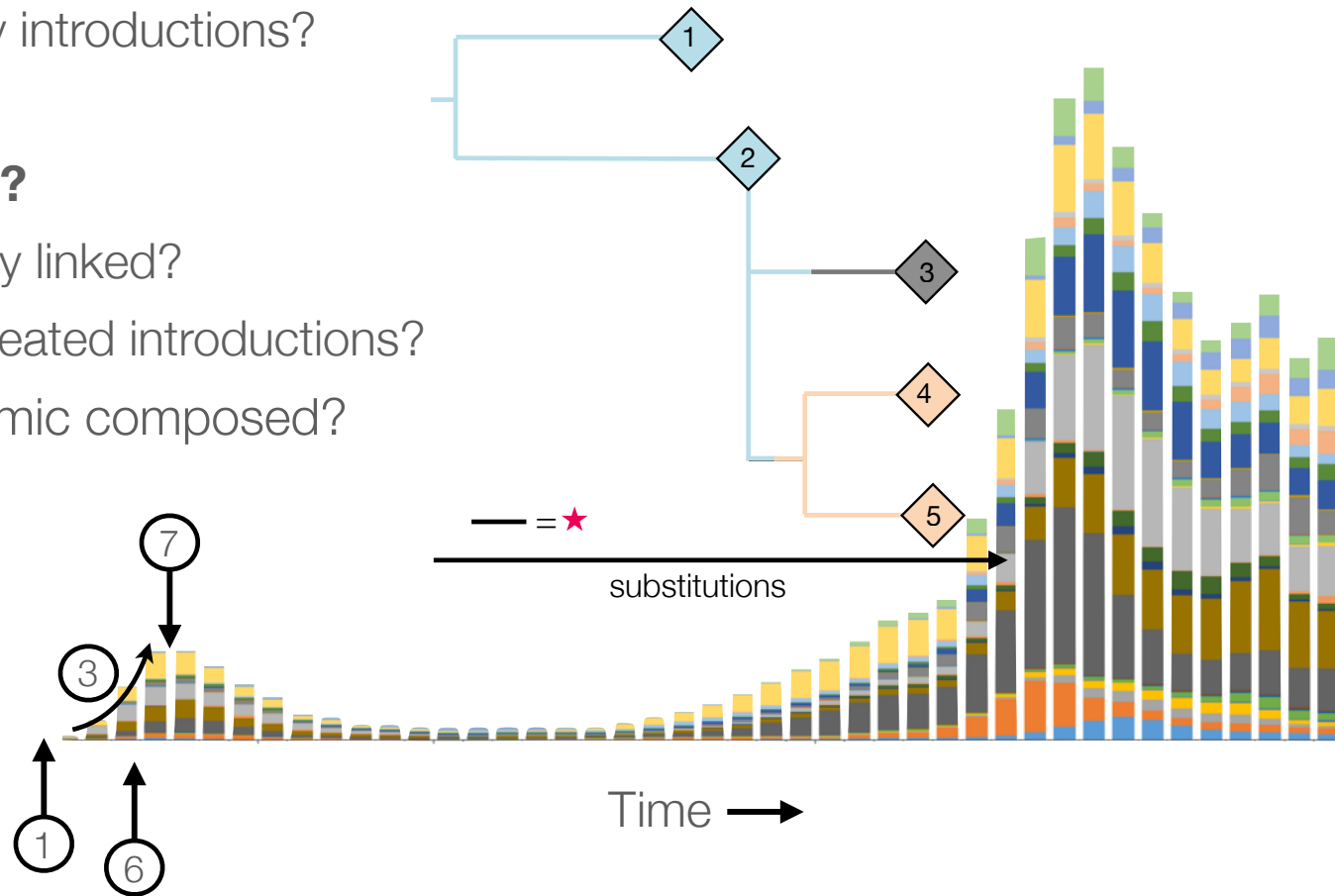
The coalescent relates the rate at which sampled 'individuals' find their common ancestors to population size



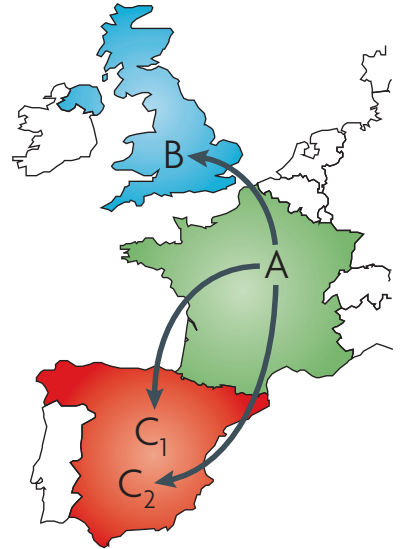
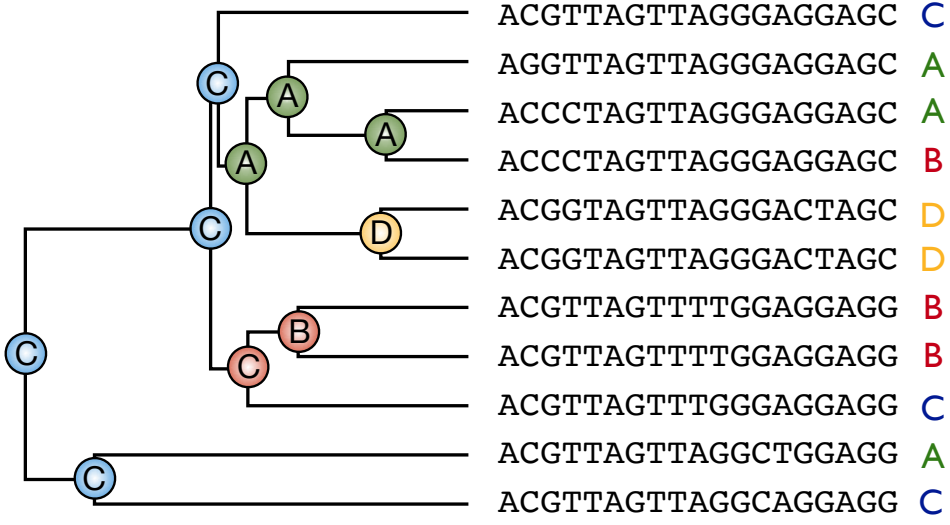
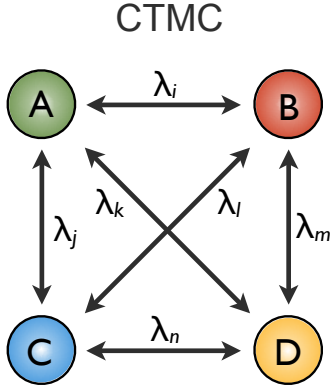
Population size

# Epidemiological questions

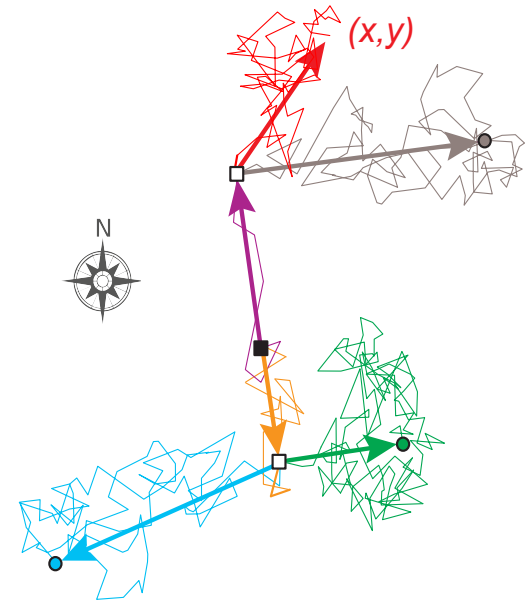
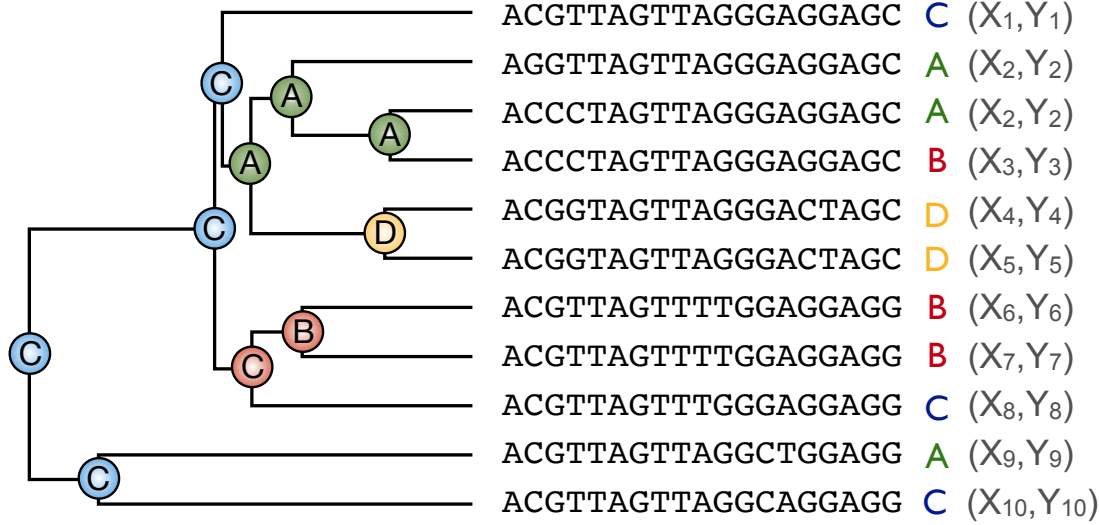
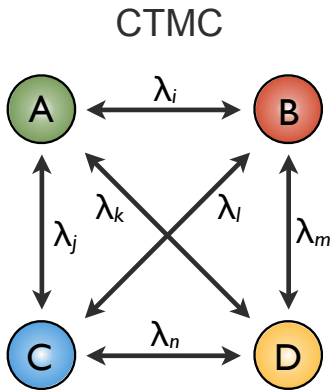
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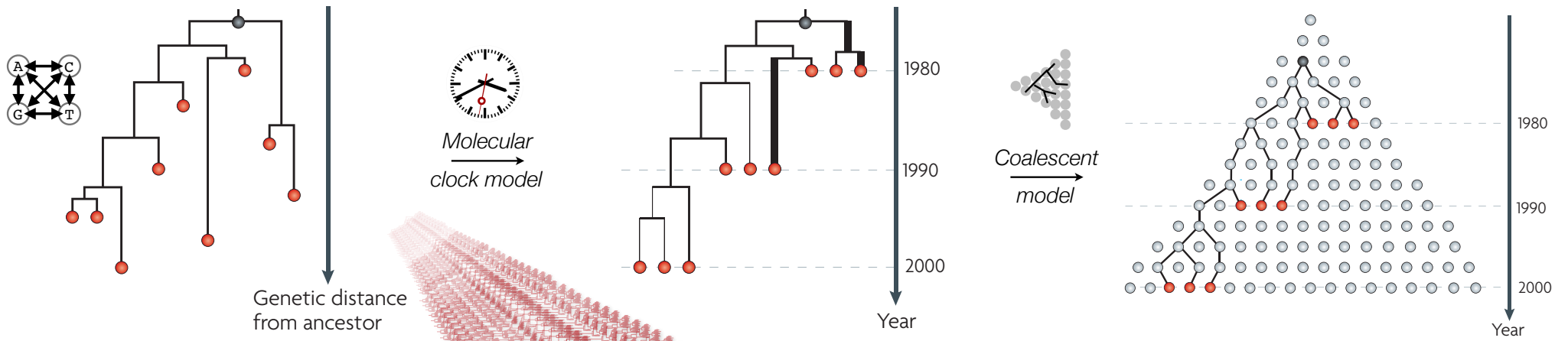


# Phylodynamic inference: spatial diffusion

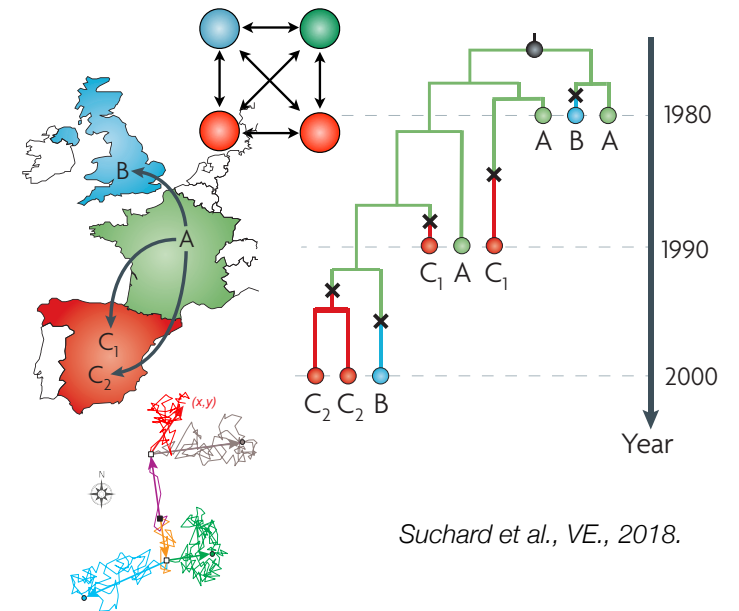
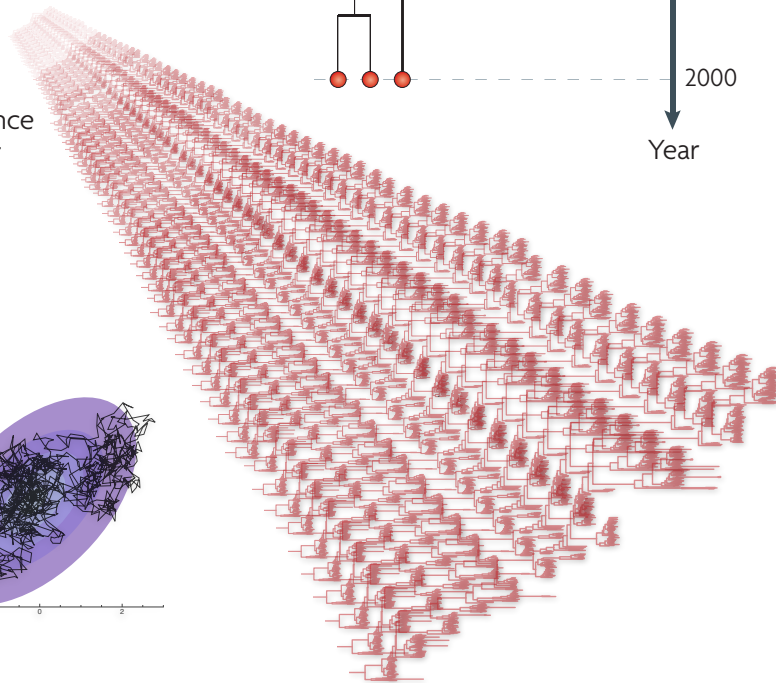
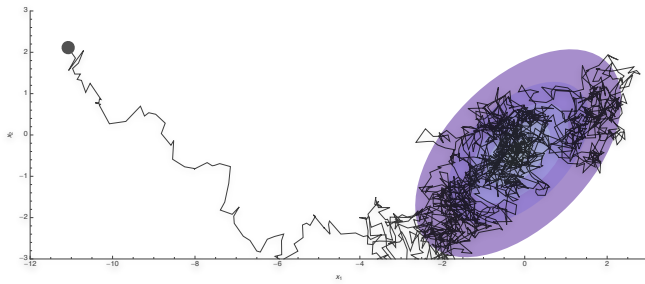


# Phylodynamic inference: spatial diffusion





Pybus and Rambaut, NRG, 2009.



Suchard et al., VE., 2018.

# BEAST: Bayesian phylodynamics



# Epidemiological questions

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9. ...

## The emergence and ongoing convergent evolution of the SARS-CoV-2 N501Y lineages

Darren P. Martin,<sup>1,\*</sup> Steven Weaver,<sup>2</sup> Houriiyah Tegally,<sup>3</sup> James Emmanuel San,<sup>3</sup> Stephen D. Shank,<sup>2</sup> Eduan Wilkinson,<sup>3</sup> Alexander G. Lucaci,<sup>2</sup> Jennifer Giandhari,<sup>3</sup> Sureshnee Naidoo,<sup>3</sup> Yeshnee Pillay,<sup>3</sup> Lavanya Singh,<sup>3</sup> Richard J. Lessells,<sup>3</sup> NGS-SA<sup>4,15</sup>, COVID-19 Genomics UK (COG-UK),<sup>5,16</sup> Ravindra K. Gupta,<sup>5,7</sup> Joel O. Wertheim,<sup>8</sup> Anton Nekturenko,<sup>9</sup> Ben Murrell,<sup>10</sup> Gordon W. Harkins,<sup>11</sup> Philippe Lemeys,<sup>12</sup> Oscar A. MacLean,<sup>13</sup> David L. Robertson,<sup>13</sup> Tulio de Oliveira,<sup>3,14,17,\*</sup> and Sergei L. Kosakovsky Pond<sup>2,\*</sup>

## CORONAVIRUS

### Timing the SARS-CoV-2 index case in Hubei province

Jonathan Pekar<sup>1,2</sup>, Michael Worobey<sup>3,\*</sup>, Niema Moshiri<sup>4</sup>, Konrad Scheffler<sup>5</sup>, Joel O. Wertheim<sup>6,\*</sup>

### The Huanan Seafood Wholesale Market in Wuhan was the early epicenter of the COVID-19 pandemic

Michael Worobey<sup>1\*</sup>, Joshua I. Levy<sup>2</sup>, Lorena Malpica Serrano<sup>1</sup>, Alexander Crits-Christoph<sup>3</sup>, Jonathan E. Pekar<sup>4,5</sup>, Stephen A. Goldstein<sup>6</sup>, Angela L. Rasmussen<sup>7,8</sup>, Moritz U. G. Kraemer<sup>9</sup>, Chris Newman<sup>10</sup>, Marion P. G. Koopmans<sup>11,12</sup>, Marc A. Suchard<sup>13,14,15</sup>, Joel O. Wertheim<sup>16</sup>, Philippe Lemeys<sup>17,18</sup>, David L. Robertson<sup>19</sup>, Robert F. Garry<sup>18,20,21</sup>, Edward C. Holmes<sup>22</sup>, Andrew Rambaut<sup>23</sup>, Kristian G. Andersen<sup>2,24\*</sup>

## CORONAVIRUS

### Cryptic transmission of SARS-CoV-2 in Washington state

Trevor Bedford<sup>1,2,3,\*</sup>, Alexander L. Greninger<sup>4,5</sup>, Pavitra Roychoudhury<sup>4,5</sup>, Lea M. Staritz<sup>2,3,1</sup>, Michael Famulare<sup>6,7</sup>, Meei-Li Huang<sup>4</sup>, Arun Nalla<sup>4</sup>, Gregory Pepper<sup>4</sup>, Adam Reinhardt<sup>4</sup>, Hong Xie<sup>4</sup>, Lasata Shrestha<sup>4</sup>, Truong N. Nguyen<sup>4</sup>, Amanda Adler<sup>4</sup>, Elisabeth Brandstetter<sup>4</sup>, Shari Cho<sup>4,8</sup>, Danielle Giroux<sup>4</sup>, Peter D. Hart<sup>4</sup>, Kaitsten Fay<sup>4</sup>, Chris D. Frazar<sup>4</sup>, Misja Ijzerman<sup>4</sup>, Kirsten Lacombe<sup>4</sup>, Jover Lee<sup>4</sup>, Anahita Kiyavand<sup>2,3</sup>, Matthew Richardson<sup>4</sup>, Thomas R. Sibley<sup>4</sup>, Melissa Truong<sup>2,3</sup>, Caitlin R. Wolf<sup>4</sup>, Deborah A. Nickerson<sup>2,3</sup>, Mark J. Rieder<sup>2,3</sup>, Janet A. Englund<sup>2,6,8</sup>, The Seattle Flu Study Investigators<sup>4</sup>, James Hadfield<sup>4</sup>, Emma B. Hodcroft<sup>4,10</sup>, John Huddleston<sup>11</sup>, Louise H. Moncla<sup>4</sup>, Nicola F. Müller<sup>4</sup>, Richard A. Neher<sup>4,10</sup>, Xianding Deng<sup>2,3</sup>, Wei Gu<sup>2</sup>, Scot Federman<sup>2</sup>, Charles Chiu<sup>2</sup>, Jeffrey S. Duchin<sup>2,3</sup>, Romesh Gautam<sup>4</sup>, Geoff Melly<sup>4</sup>, Brian Hiatt<sup>4</sup>, Philip Dykema<sup>4</sup>, Scott Lindquist<sup>4</sup>, Krista Queen<sup>2</sup>, Ying Tao<sup>2</sup>, Anna Uehara<sup>4</sup>, Susiang Tong<sup>2</sup>, Duncan MacCannell<sup>4</sup>, Gregory L. Armstrong<sup>4</sup>, Geoffrey S. Baird<sup>4</sup>, Helen Y. Chu<sup>2,7</sup>, Jay Shendure<sup>2,3,7</sup>, Keith R. Jerome<sup>4</sup>

## Article

### Untangling introductions and persistence in COVID-19 resurgence in Europe

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Philippe Lemeys<sup>12,13</sup>, Nick Ruktanonchai<sup>14</sup>, Samuel L. Hong<sup>1</sup>, Vittoria Colizza<sup>1</sup>, Chiara Poletto<sup>1</sup>, Frederik Van den Broeck<sup>15</sup>, Mandev S. Gill<sup>1</sup>, Xiang Ji<sup>1</sup>, Anthony Levasseur<sup>1</sup>, Bas B. Oude Munnink<sup>1</sup>, Marion Koopmans<sup>1</sup>, Adam Sadleir<sup>16</sup>, Shengjie Liu<sup>1</sup>, Andrew J. Tatem<sup>1</sup>, Guy Baele<sup>1</sup>, Marc A. Suchard<sup>13,14</sup> & Simon Delloo<sup>14,15</sup>

## Article

### Spread of a SARS-CoV-2 variant through Europe in the summer of 2020

<https://doi.org/10.1038/s41586-021-03677-y>

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Emma B. Hodcroft<sup>1,2,3</sup>, Moira Zuber<sup>1</sup>, Sarah Nadeau<sup>1,4</sup>, Timothy G. Vaughan<sup>1,4</sup>, Katharine H. D. Crawford<sup>4,5</sup>, Christian L. Althaus<sup>1</sup>, Martina L. Reichmuth<sup>1</sup>, John E. Bowen<sup>1</sup>, Alexandra C. Walls<sup>1</sup>, Davide Corti<sup>1</sup>, Jesse D. Bloom<sup>1,6</sup>, David Veissler<sup>1</sup>, David Mateo<sup>1</sup>, Alberto Hernandez<sup>1</sup>, Itzaki Comas<sup>1,7</sup>, Fernando González-Candelas<sup>1,8</sup>, SeqCOVID-SPAIN consortium<sup>1</sup>, Tanja Stadler<sup>4,9</sup> & Richard A. Neher<sup>1,2,10,11</sup>