

# Phylogenetic inference of contact network parameters with kernel approximate Bayesian computation

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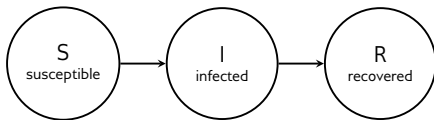
Rosemary M McCloskey<sup>1</sup>   Richard H Liang<sup>1</sup>   Art FY Poon<sup>1,2</sup>

HIV Dynamics & Evolution, Woods Hole, USA, April 25, 2016

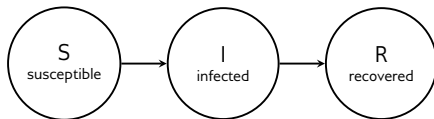
<sup>1</sup>BC Centre for Excellence in HIV/AIDS, Vancouver, Canada

<sup>2</sup>Department of Medicine, University of British Columbia, Vancouver, Canada

# Most epidemiological models assume homogeneous mixing

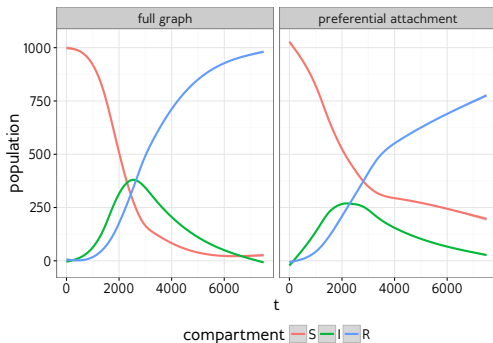
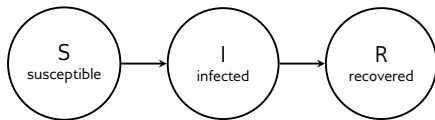


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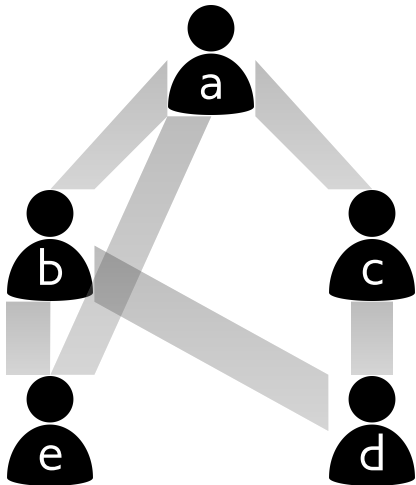
- Often provide a reasonable approximation in practice.

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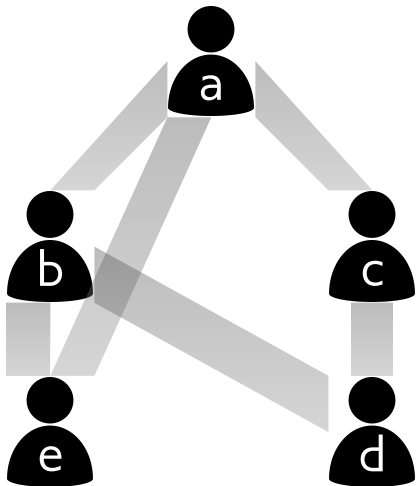


- Often provide a reasonable approximation in practice.
- Can be inaccurate when substantial contact heterogeneity exists.

# Network models capture contact heterogeneity

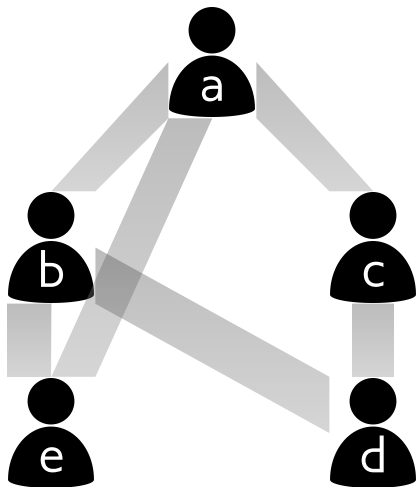


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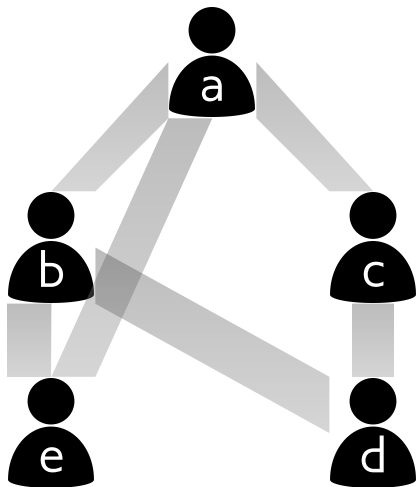
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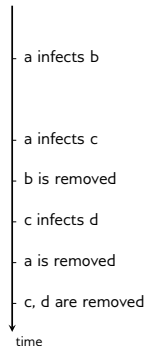
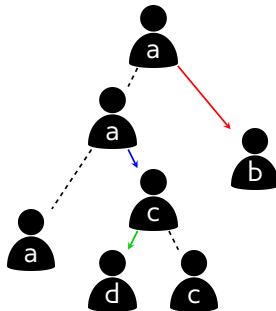
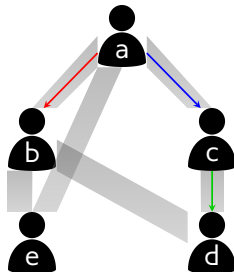
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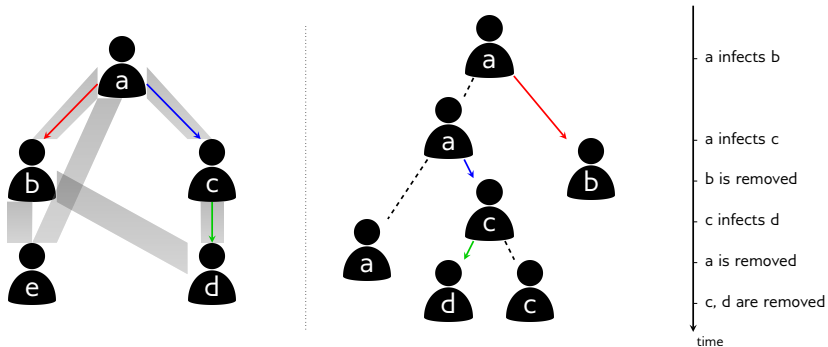
- May offer more accurate predictions for highly structured populations.
- Network parameters may be of interest for their own sake, e.g. are there superspreaders?
- Extremely difficult to estimate in practice.



# Contact networks shape transmission trees

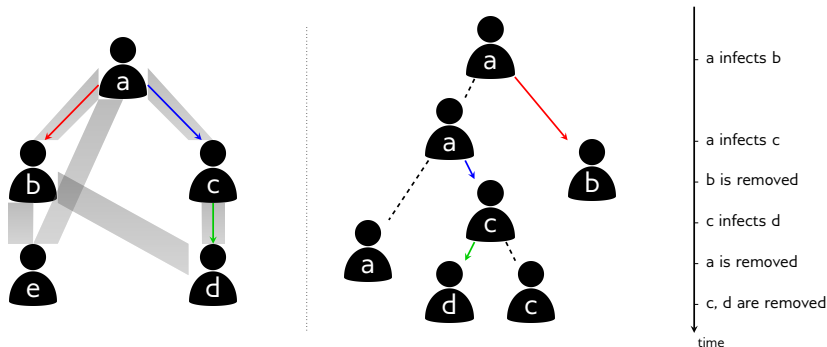


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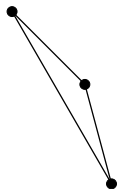
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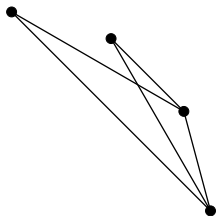
- Transmission trees in turn shape viral phylogenies.
- **Aim:** estimate contact network parameters from viral phylogenies.

# Barabási-Albert model incorporates preferential attachment



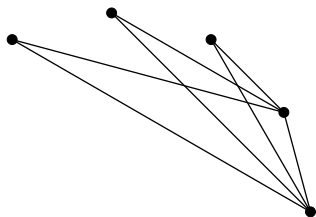
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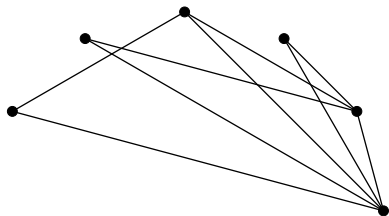
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- Attach new nodes with  $m$  edges.

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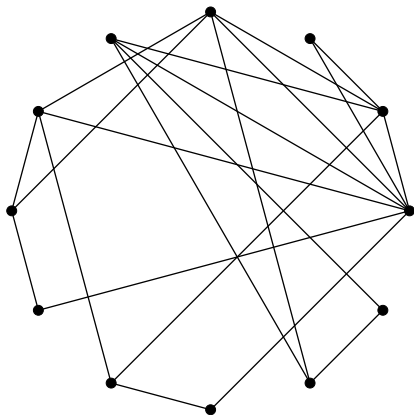
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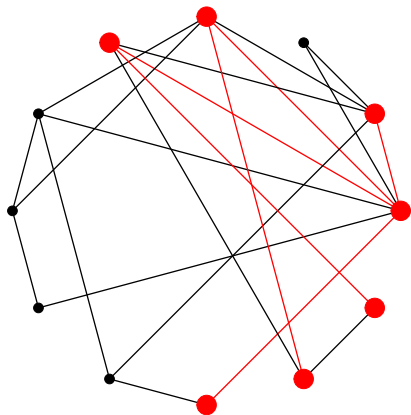
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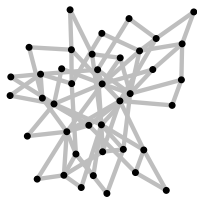


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- Attach new nodes with  $m$  edges.
- Other endpoints of degree  $d$  are chosen with probability  $\propto d^\alpha + 1$ .
- Continue until network has  $N$  nodes.
- Also consider the prevalence  $l$ .

# Do network parameters measurably affect tree shape?

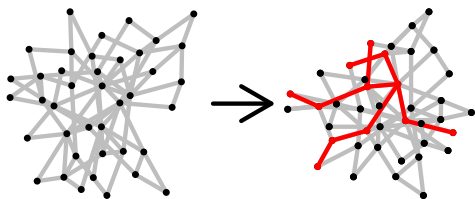


- Generate networks under different parameter values (number of nodes  $N$ , number of edges per vertex  $m$ , preferential attachment power  $\alpha$ ).

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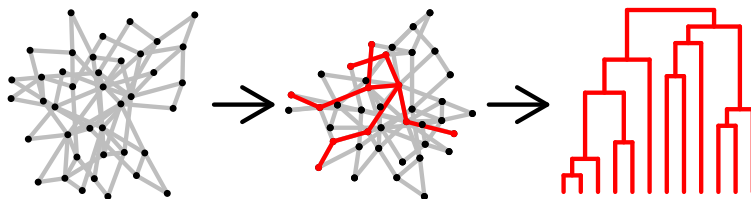


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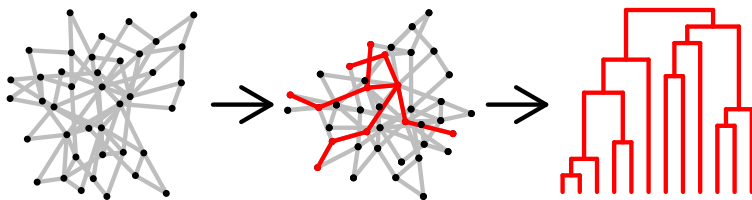


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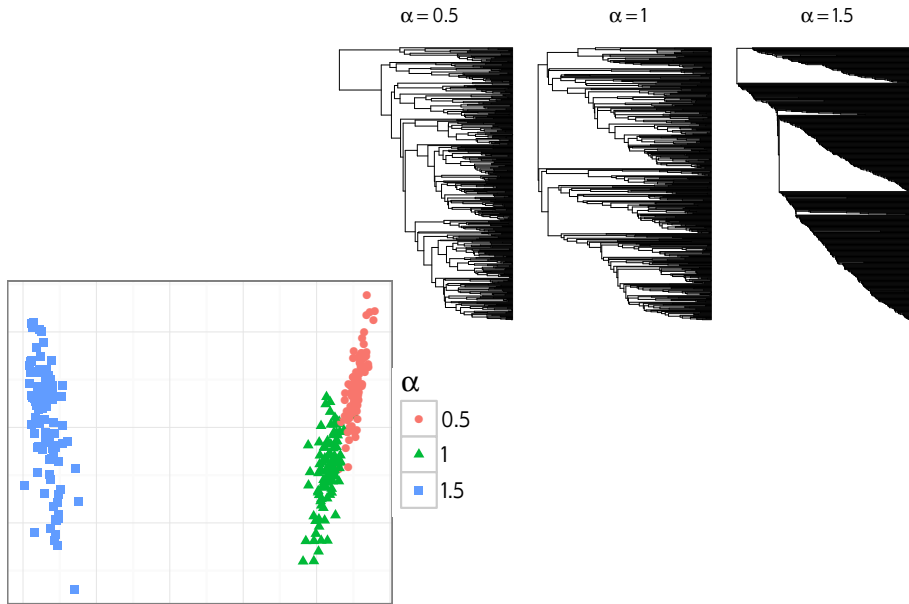
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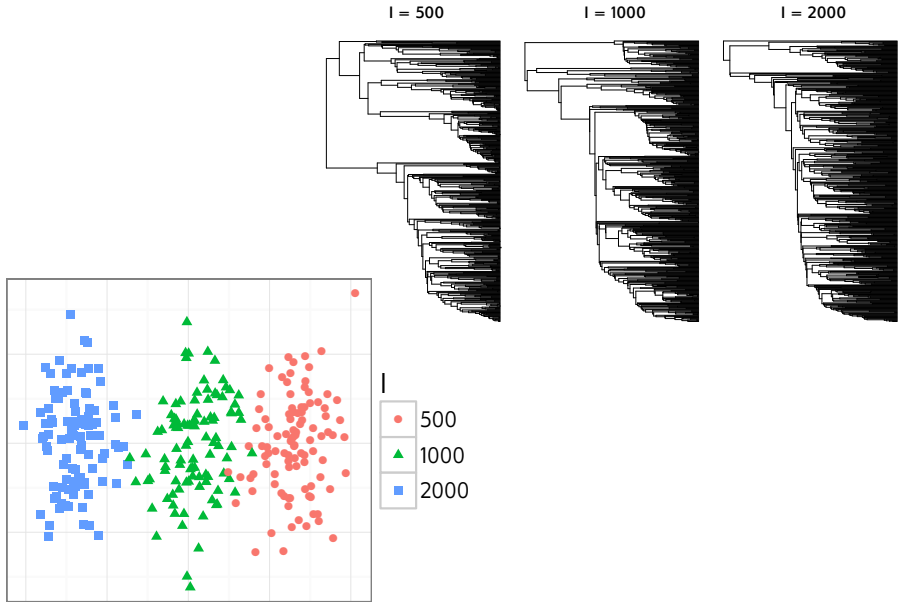
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- Simulate epidemic over each network until  $I$  nodes are infected.
- Randomly subsample to form transmission trees.
- Compare trees pairwise using tree kernel<sup>1</sup>.

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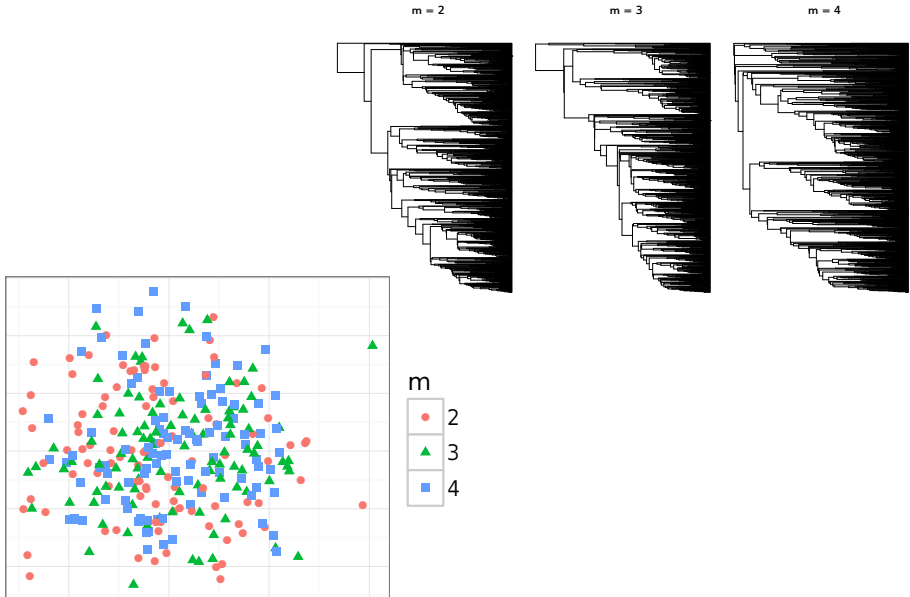
# Preferential attachment power $\alpha$ affects tree shape



# Prevalence / affects tree shape

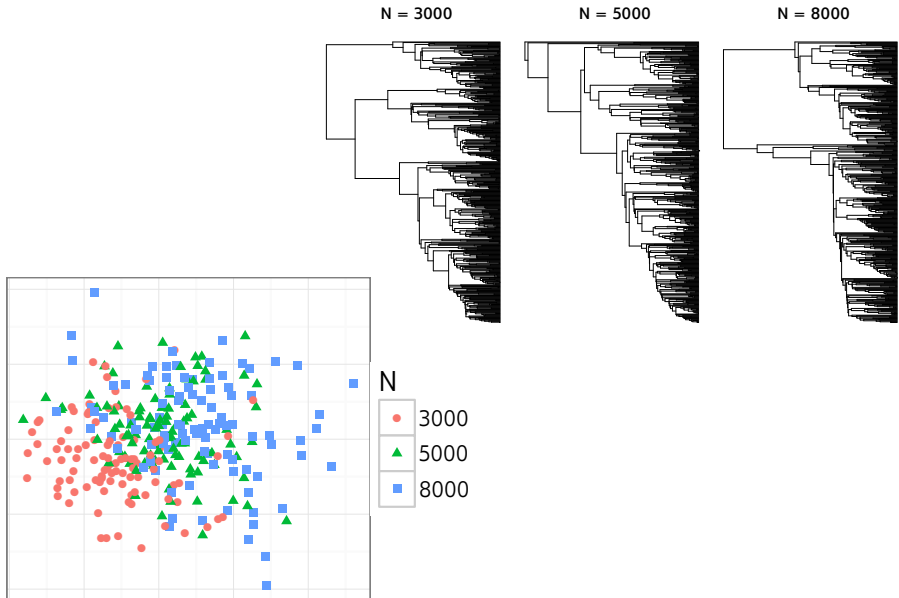


# Number of edges per vertex $m$ does not affect tree shape





# Number nodes $N$ modestly affects tree shape



# Simulated trees can be used to fit a network model

sample  
parameters

$\theta_1$

$\theta_2$

$\theta_3$

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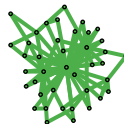
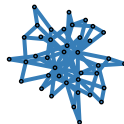
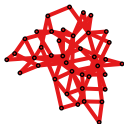
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simulate  
networks



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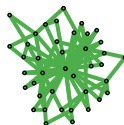
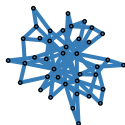
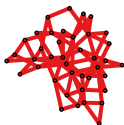
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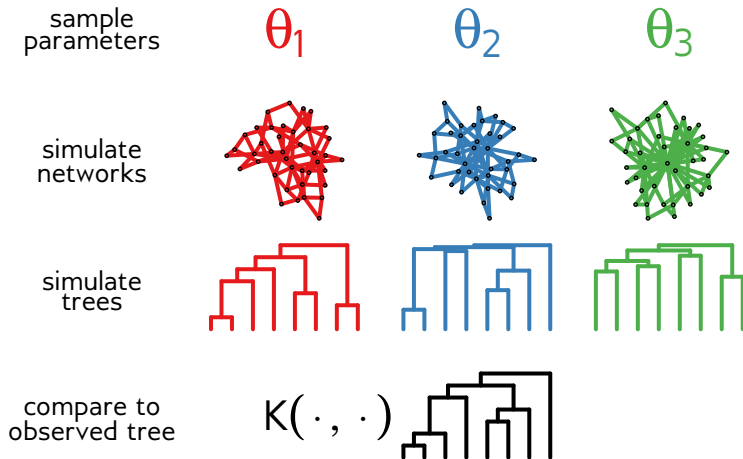
simulate  
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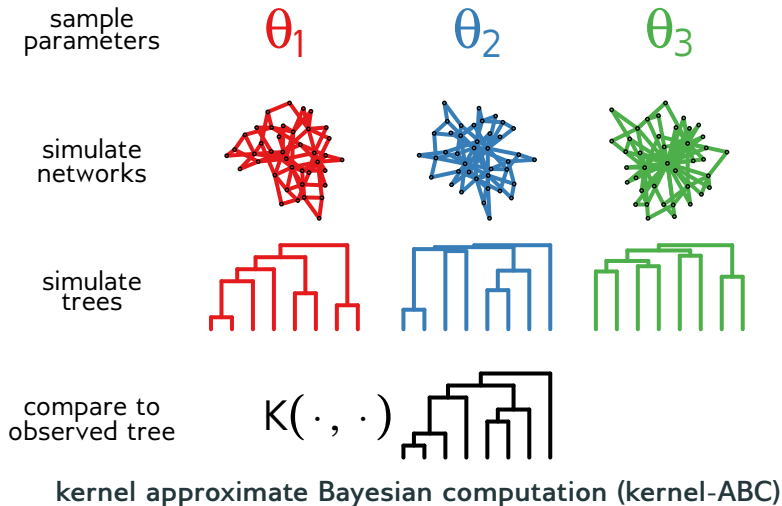
simulate  
trees



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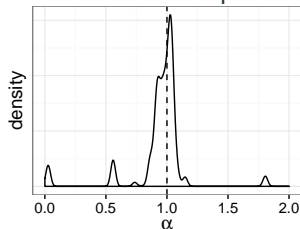


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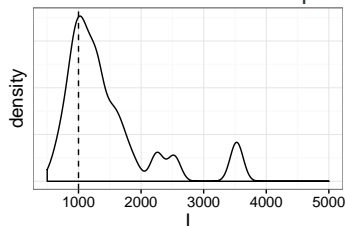


# Example simulation shows $\alpha$ and $I$ can be reconstructed

preferential attachment power

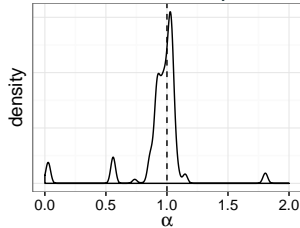


prevalence

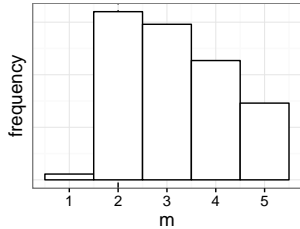
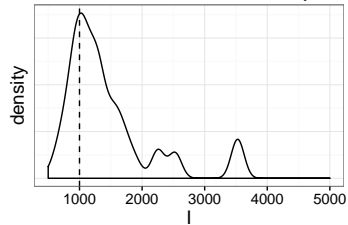


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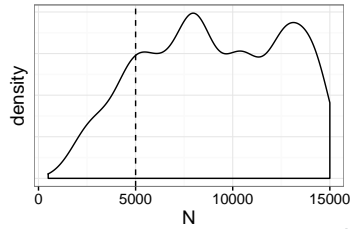
preferential attachment power



prevalence



edges per vertex

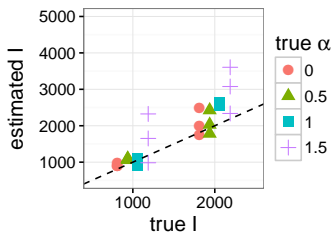
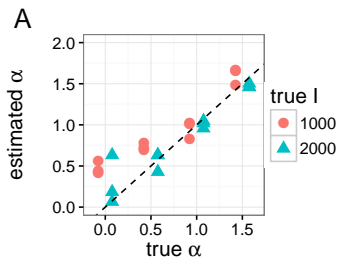


total nodes



# Maximum *a posteriori* estimates are accurate for $\alpha$ and $I$

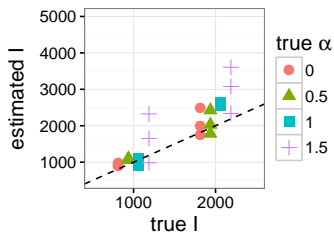
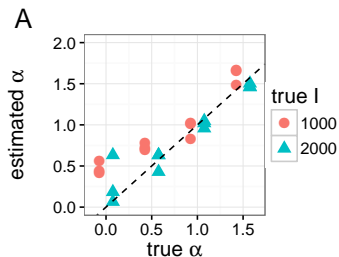
preferential attachment power



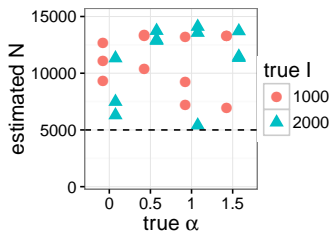
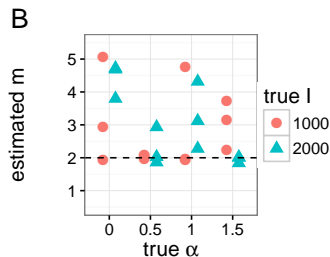
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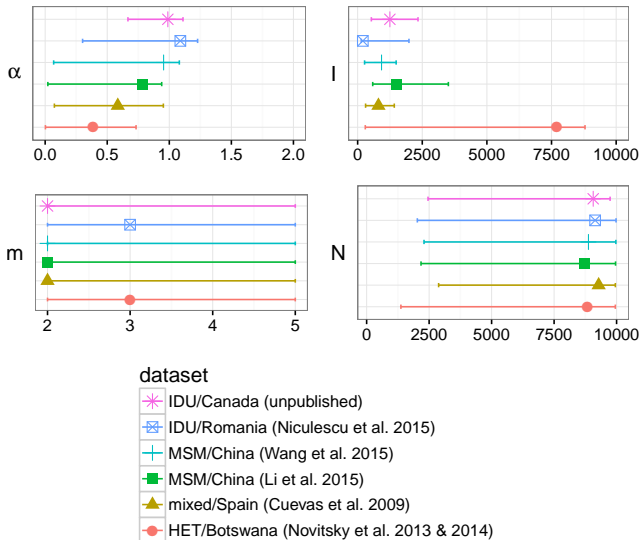


prevalence

total nodes

# Real world HIV datasets exhibit network heterogeneity

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[github.com/rmcclosk/netabc](https://github.com/rmcclosk/netabc)



# Acknowledgements

## BC Centre for Excellence in HIV/AIDS

Art Poon

Jeff Joy

Richard Liang

Thuy Nguyen

P. Richard Harrigan



BRITISH COLUMBIA  
CENTRE for EXCELLENCE  
in HIV/AIDS



CIHR IRSC  
Canadian Institutes of Health Research  
Instituts de recherche en santé du Canada



Genome  
British Columbia

10 a decade of innovation

## University of British Columbia

Sarah Otto

Alexandre Bouchard-Côté



GenomeCanada



GenomeQuébec

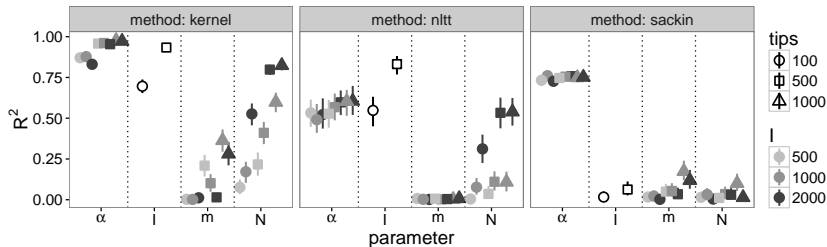
BILL & MELINDA  
GATES foundation



CIHR Strategic Training Program in  
BIOINFORMATICS

# Classifiers for network parameters

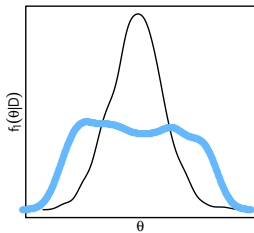
- kSVR using tree kernel
- SVR using normalized lineages-through-time<sup>2</sup>
- linear regression using Sackin's index<sup>3</sup>



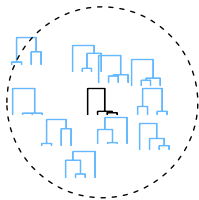
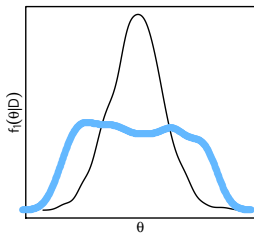
<sup>2</sup>Thijs Janzen, Sebastian Höhna, and Rampal S Etienne (2015). "Approximate Bayesian computation of diversification rates from molecular phylogenies: introducing a new efficient summary statistic, the nlTT". In: *Methods in Ecology and Evolution* 6.5, pp. 566–575.

<sup>3</sup>Kwang-Tsao Shao (1990). "Tree balance". In: *Systematic Biology* 39.3, pp. 266–276.

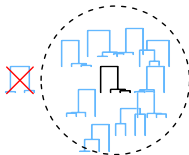
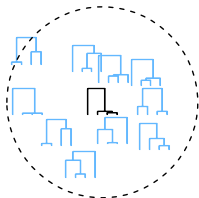
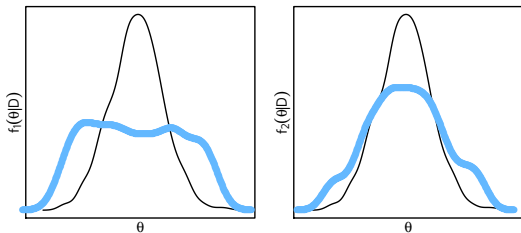
# Sequential Monte Carlo for ABC



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