

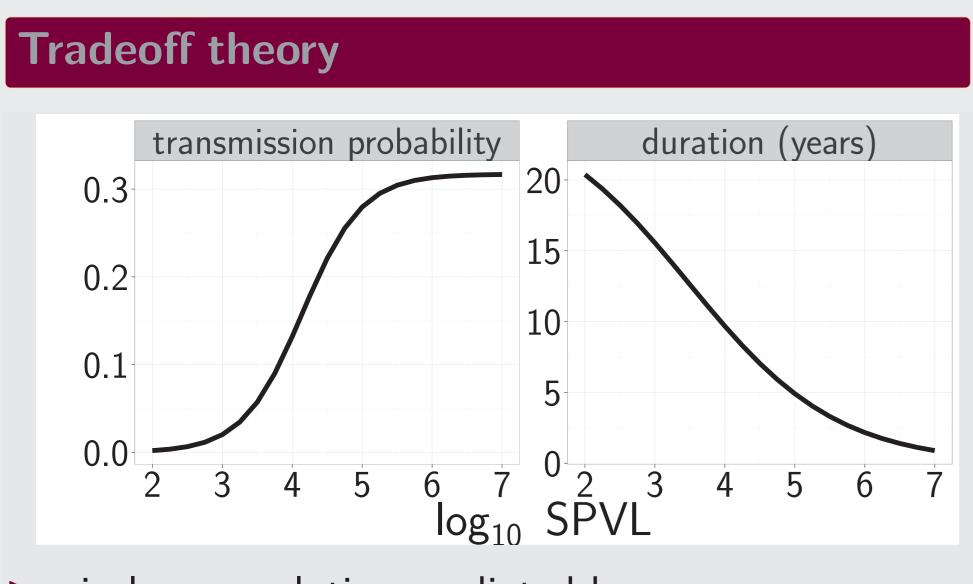
HIV virulence evolution in structured epidemic models

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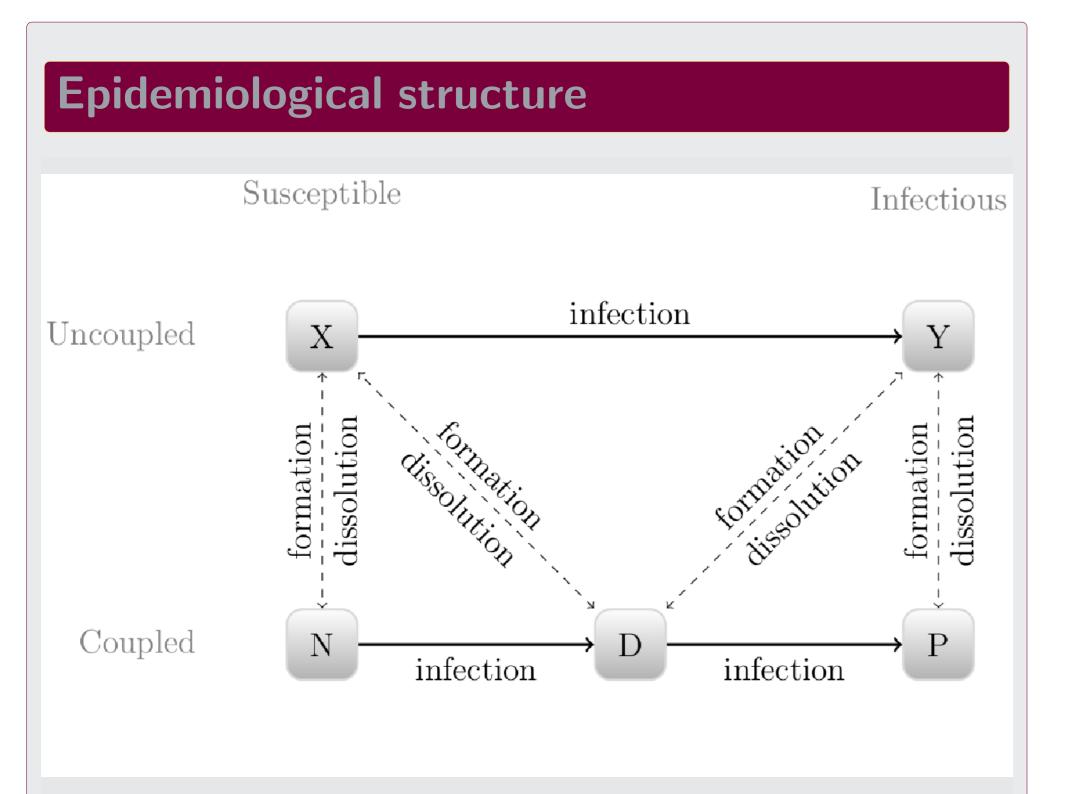


Summary

Pathogens can evolve rapidly in response to changing conditions (e.g., epidemic stage or public health interventions). Models of **eco-evolutionary dynamics** often neglect important epidemiological processes, such as the dynamics of sexual partnerships. We compared **models with a range of complexity** of partnership dynamics and extra-partnership contact.



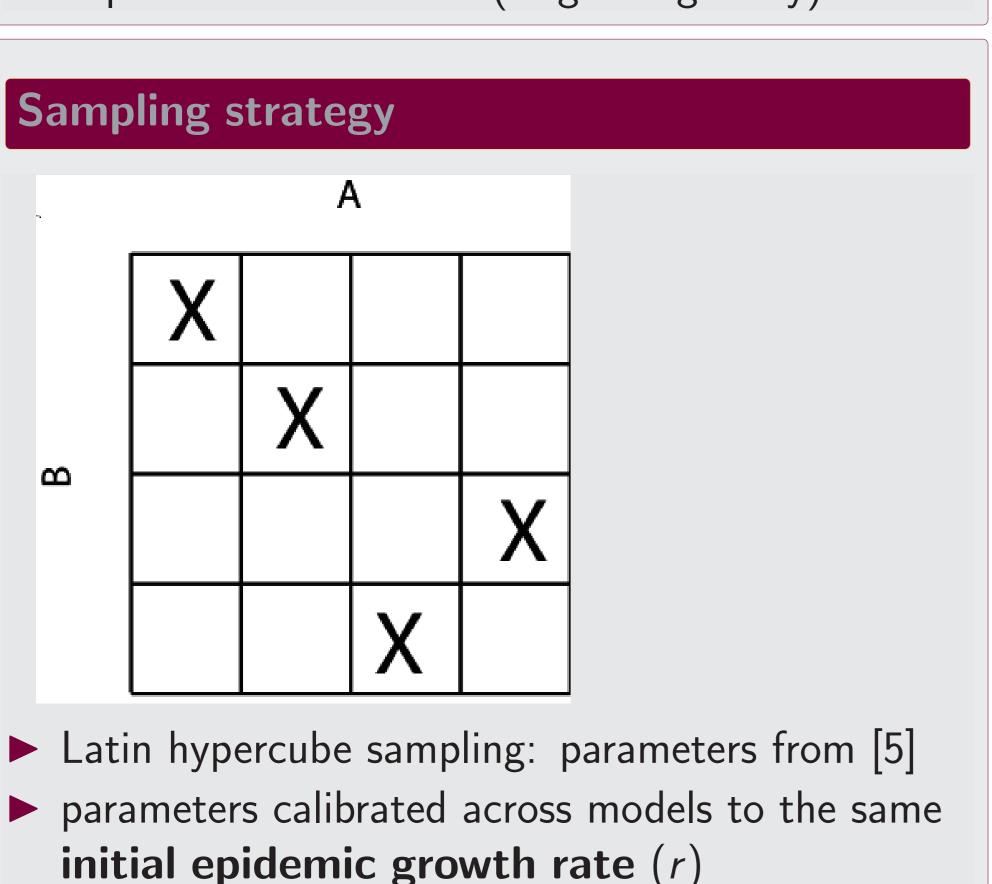
- virulence evolution mediated by transmission-vs-clearance tradeoff
- ▶ still debated [1, 2]
- ► HIV [3]: **set-point viral load** correlated with transmission probability, rate of progression to AIDS (data from Rakai, Uganda)
- eco-evolutionary virulence dynamics: [4]

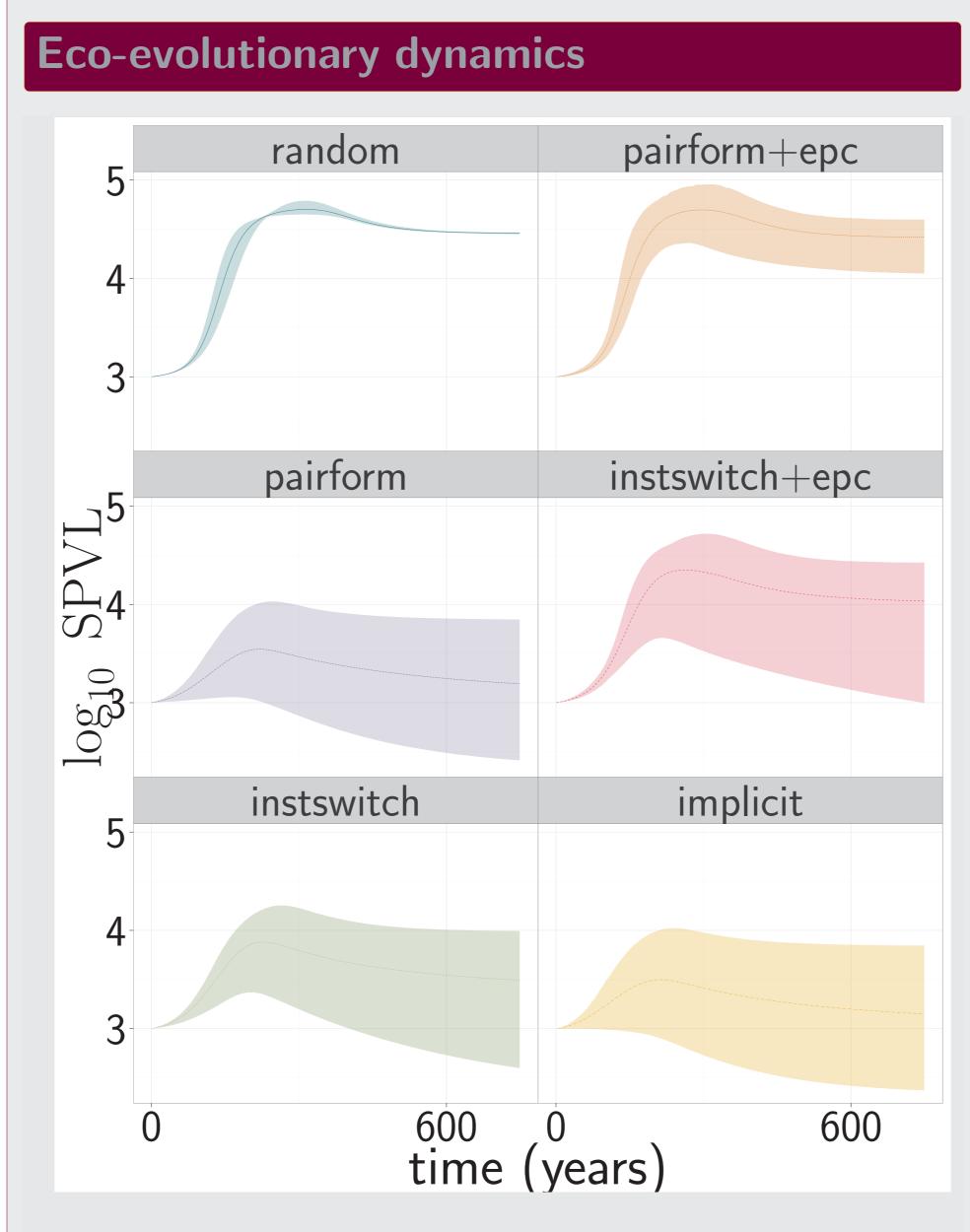


Champredon et al. 2013 [5]

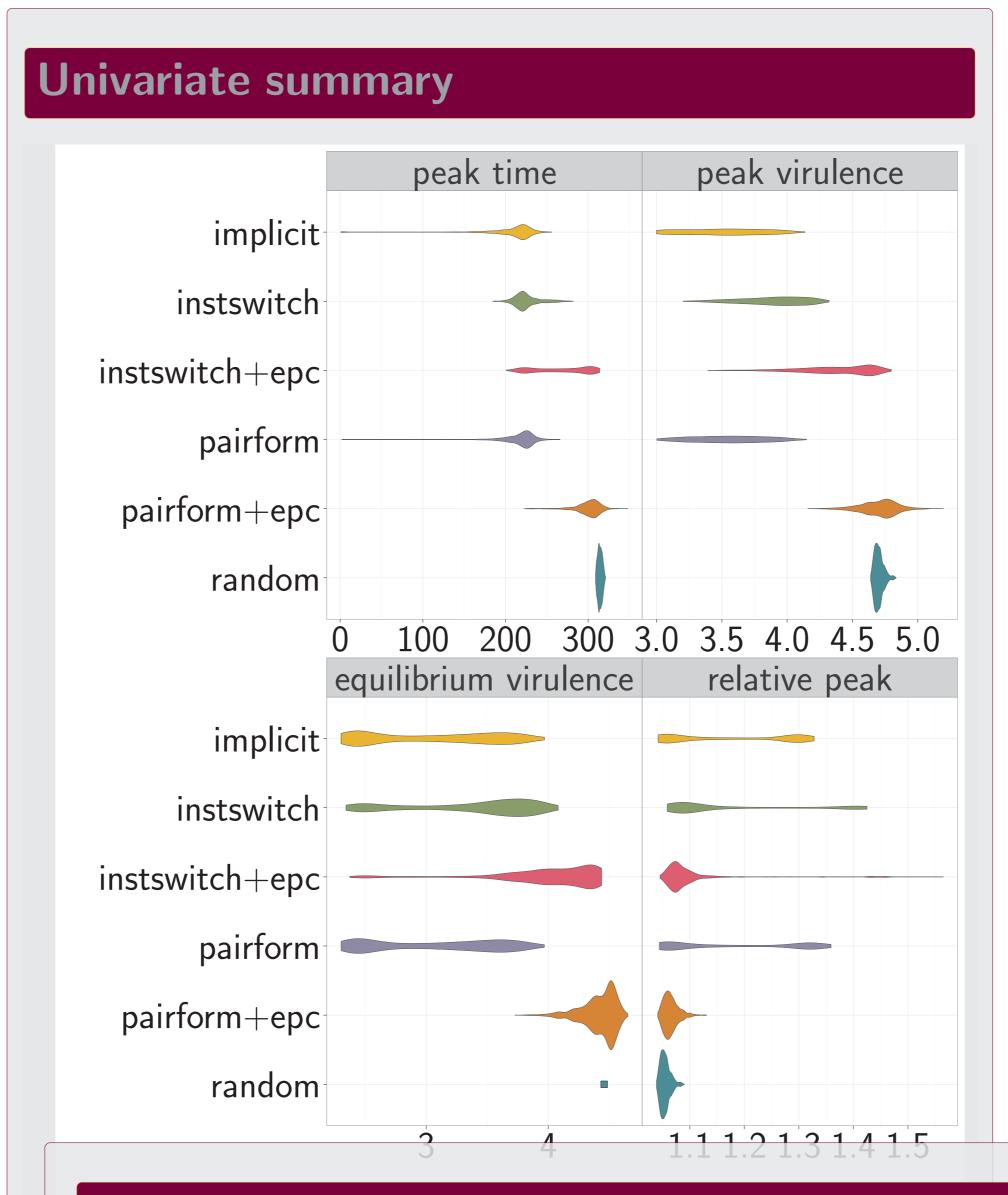
Spectrum of model complexity:

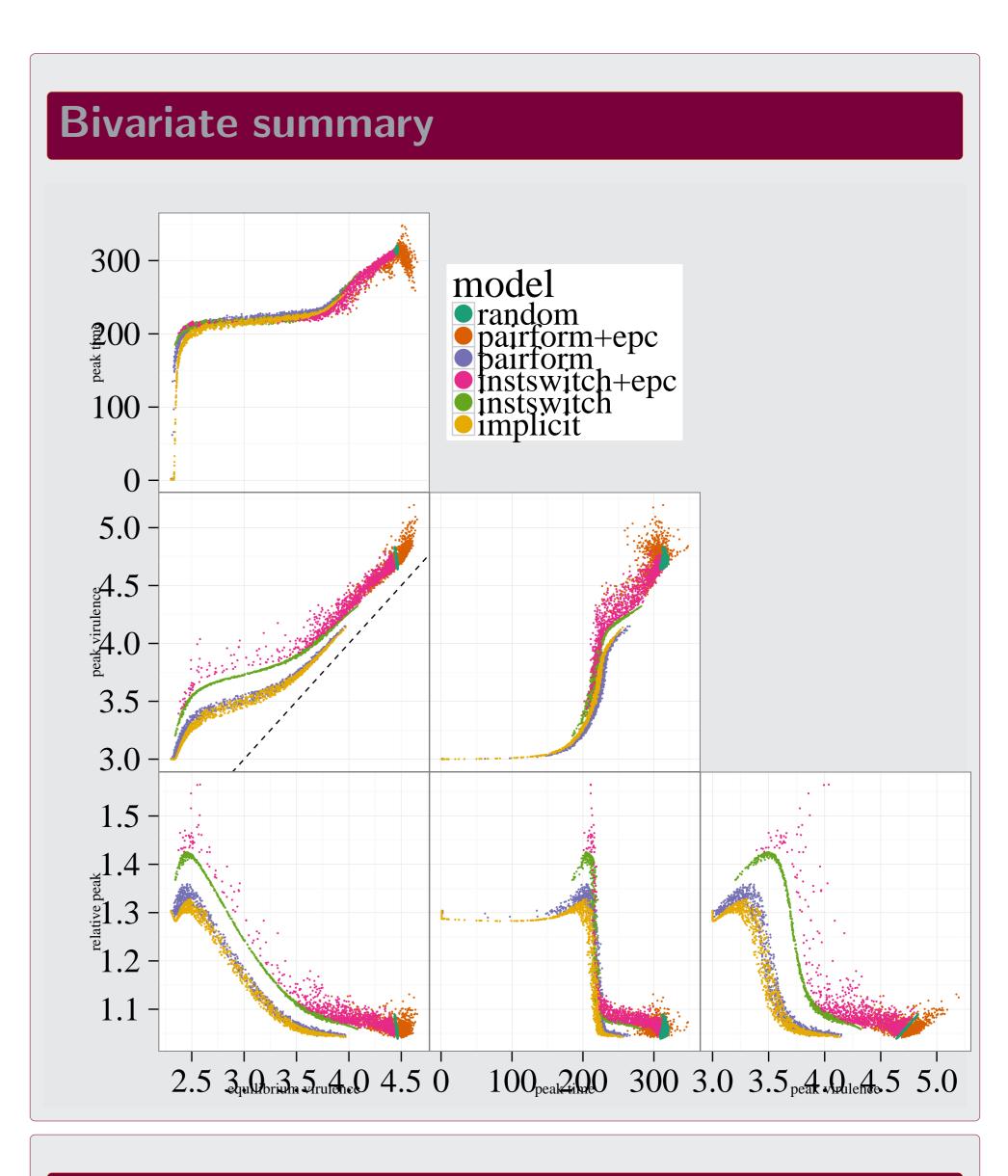
- **pair formation**: instantaneous or delayed?
- extra-pair contact (epc): present or absent?
- ▶ implicit model: no explicit partnerships, force of infection expression derived from \mathcal{R}_0 of pair-formation model (without epc)
- ► random-mixing model: standard SIR model Simplified disease model (single stage only)





- significant variation across model structures
- ▶ least (random) and most (pairform+epc) models most similar: single individuals and extra-pair contact wash out effects of structure
- implicit model is **most different**
- ▶ random-mixing model underestimates (?) variability





Conclusions and open questions

- ► Random-mixing models gave the closest match to the most realistic models; extra-pair contact washed out the effects of epidemiological structure
- ightharpoonup Variation among models (model structure) pprox variation within models (parameter uncertainty)
- Large differences in evolutionary dynamics among different epidemiological models suggest caution in predicting evolutionary responses

References

- [1] Ebert D, Bull JJ. Challenging the trade-off model for the evolution of virulence: is virulence management feasible? Trends Microbiol. 2003;11(1):15–20.
- [2] Alizon S, Michalakis Y. Adaptive virulence evolution: the good old fitness-based approach. Trends in Ecology & Evolution. 2015 Jan;30(5):248-254. Available from: http://www.cell.com/article/S016953471500049X/abstract.
- [3] Fraser C, Lythgoe K, Leventhal GE, Shirreff G, Hollingsworth TD, Alizon S, et al. Virulence and Pathogenesis of HIV-1 Infection: An Evolutionary Perspective. Science. 2014 Mar;343(6177):1243727. Available from: http://www.sciencemag.org/content/343/6177/1243727.
- [4] Shirreff G, Pellis L, Laeyendecker O, Fraser C. Transmission Selects for HIV-1 Strains of Intermediate Virulence: A Modelling Approach. PLoS Computational Biology. 2011 Oct;7(10):e1002185. WOS:000297262700019.
- [5] Champredon D, Bellan S, Dushoff J. HIV Sexual Transmission Is Predominantly Driven by Single Individuals Rather than Discordant Couples: A Model-Based Approach. PLoS ONE. 2013 12;8(12):e82906.

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