

Dr. Benjamin Bolker

Professor
Departments of Mathematics & Statistics and Biology

1280 King Street West Hamilton, ON L8S 4K1 Canada phone (905) 525-9140 x23320 fax (905) 522-0935 e-mail bolker@mcmaster.ca June 25, 2016

To the editor,

We (Daniel Park and Ben Bolker) are submitting our manuscript, "Effects of Epidemiological Structure on the Transient Evolution of HIV Virulence", for consideration for publication in *PLoS Computational Biology*. Using HIV as a model system, our study explores the transient, eco-evolutionary changes in pathogen characteristics over the course of an epidemic. Building on modeling frameworks and estimated parameters developed by Shirreff *et al.* [5] and Champredon *et al.* [1], we ask how predicted changes in virulence depend on way that contact patterns (e.g. the dynamics of sexual partnership change and contact outside of stable partnerships) are modeled.

This study is important because researchers are actively developing — and publishing in high-impact journals — studies that predict the evolutionary consequences of public-health interventions [3, 4, 2]. Our work suggests that basic epidemiological structures that are neglected in these models can lead to two- to fourfold differences in predictions of outcomes such as transmission probability or expected time to progress to AIDS.

Sincerely,

Benjamin Bolker

Beryam Baller

- [1] D. Champredon, S. Bellan, and J. Dushoff. HIV sexual transmission is predominantly driven by single individuals rather than discordant couples: A model-based approach. *PLoS ONE*, 8(12):e82906, 12 2013.
- [2] J. Herbeck, J. Mittler, G. Gottlieb, S. Goodreau, J. Murphy, A. Cori, M. Pickles, and C. Fraser. Evolution of HIV virulence in response to widespread scale up of antiretroviral therapy: a modeling study. *bioRxiv*, page 039560, 2016.
- [3] R. Payne, M. Muenchhoff, J. Mann, H. E. Roberts, P. Matthews, E. Adland, A. Hempenstall, K.-H. Huang, M. Brockman, Z. Brumme, M. Sinclair, T. Miura, J. Frater, M. Essex, R. Shapiro, B. D. Walker, T. Ndungu, A. R. McLean, J. M. Carlson, and P. J. R. Goulder. Impact of HLA-driven HIV adaptation on virulence in populations of high HIV seroprevalence. *Proceedings of the National Academy of Sciences*, 111(50):E5393–E5400, Dec. 2014.
- [4] H. E. Roberts, P. J. Goulder, and A. R. McLean. The impact of antiretroviral therapy on population-level virulence evolution of HIV-1. *Journal of The Royal Society Interface*, 12(113):20150888, 2015.
- [5] G. Shirreff, L. Pellis, O. Laeyendecker, and C. Fraser. Transmission selects for HIV-1 strains of intermediate virulence: A modelling approach. *PLoS Computational Biology*, 7(10):e1002185, Oct. 2011. WOS:000297262700019.