Cost-effective boosting allocations in the post-Omicron era of COVID-19 management

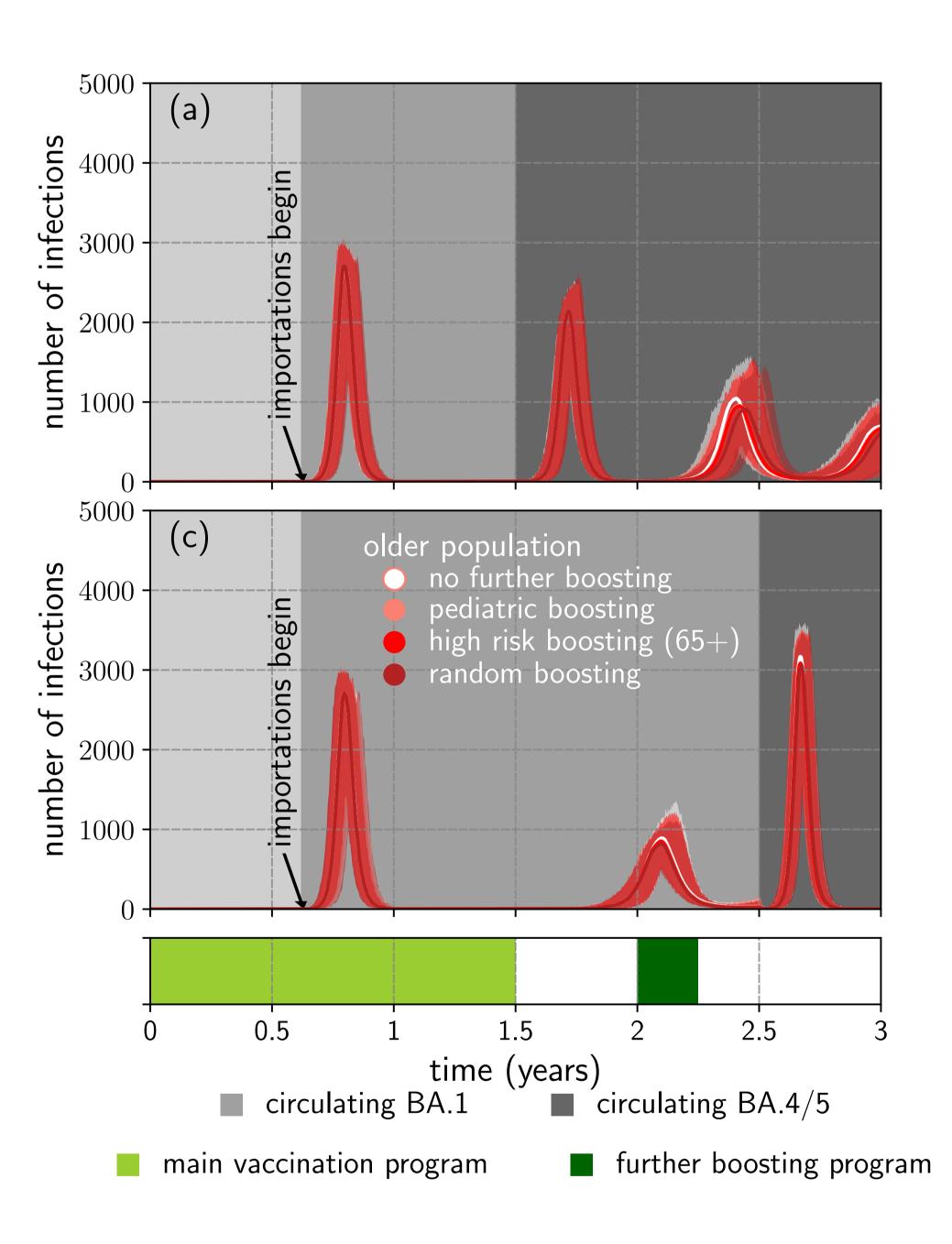
Thao Le^{1, 2, 3}, Eamon Conway⁴, Edifofon Akpan⁵, Isobel Abell^{1, 2}, Patrick Abraham⁵, Christopher Baker^{1, 2, 3}, Patricia Campbell^{5, 6}, Deborah Cromer⁷, Michael Lydeamore⁸, Yasmine McDonough⁴, Ivo Mueller^{4, 9}, Gerard Ryan^{5, 10}, Camelia Walker¹, Yingying Wang⁵, Natalie Carvalho⁵, Jodie McVernon^{6, 11}

Vaccination + Infection = Hybrid Immunity

As we transition from pandemic to endemic circulation of COVID-19 around the world, populations have varying levels of hybrid immunity resulting from previous vaccination and infection histories. In this work we assess the impact and cost-effectiveness of targeted COVID-19 vaccination strategies for various countries.

Our modelling simulates complex immune landscapes by considering a variety of factors that impact a country's infections, deaths and vaccine cost-effectiveness:

- Age demographic (older or younger)
- Vaccination coverage (high or low)
- Attack rate of previous COVID-19 waves (high or low transmission)
- Timing of immune escape of a new variant (early or late emergence)
- Current/future boosting vaccination strategy



Model Pipeline Immunological model **Population transmission Clinical pathways Cost-effectiveness analysis** Outputs: Infection parameters, **Outputs:** infection histories Outputs: clinical outcomes Outputs: Vaccine cost-effectiveness for neutralisation antibody parameters exemplar country contexts for any variant/vaccine combination Clinical progression probabilities Age distribution, vaccination Disability weights and life tables, Clinical studies and length of stay distributions coverage and schedule, contact disease management cost, by variant and age matrices, transmission potential vaccine cost, exemplar countries Scan me for more details! inputs and parameters Scenario 1:

Older population, high transmission, high vaccination coverage

We compare the impact of a variety boosting strategies on infections/deaths and vaccine cost-effectiveness given early or late emergence of an immune escape variant

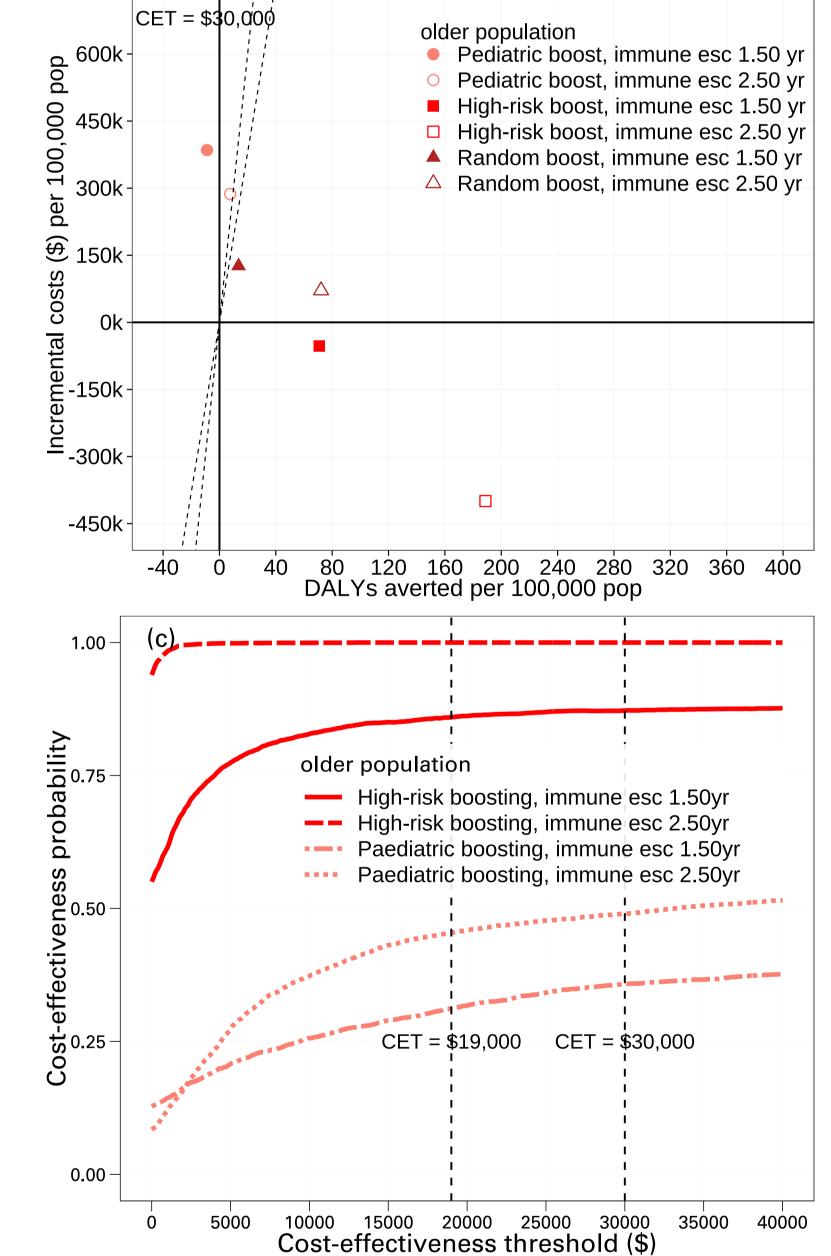
lmmune escape	Boosting strategy	Median deaths
Early	No further boosting Pediatric (ages 5-15) High-risk (65+) Random	40.0 (28.0, 53.0) 40.0 (29.0, 53.0) 34.0 (23.0, 46.0) 39.0 (27.0, 52.0)
Late	No further boosting Pediatric (ages 5-15) High-risk (65+) Random	49.0 (36.0, 62.0) 49.0 (36.0, 62.0) 34.0 (23.0, 47.0) 44.0 (32.0, 59.0)

Infections and deaths:

- We see little difference in infection dynamics between vaccination strategies
- Late emergence of an immune escape variant shifts epidemic peaks to the right
- We do see a difference in severe outcomes between strategies
- High-risk boosting averts the most severe disease

Vaccine cost-effectiveness

- Boosting is more cost-effective when it occurs prior to immune escape
- High-risk boosting is likely to be cost-effective or cost-saving
- Pediatric boosting does not appear to be cost-effective
- The cost-effectiveness of high-risk boosting is driven primarily by vaccine program (delivery and dose) costs, followed by disease management costs in general ward



CET = \$19,000

Elder-targeted strategies are most likely to be cost-effective and could even be cost-saving The flexibility of our modelling pipeline allowed us to assess the impact and costeffectiveness of booster strategies across a variety of populations. Across all scenarios we found:

750k-

- Pediatric programs (primary of boosting) were not cost-effective
- Absolute harms averted by vaccination are influenced by: age and risk profile of population, prior hybrid immunity, and timing of emergence of an immune escape variant in relation to booster delivery
- Half-yearly 'high risk' booster programs are more expensive, but may be cost effective in older, high income populations

This work was presented to the Advisory Committee on Immunization and Vaccines-related Implementation Research (IVIRAC) and was subsequently cited as part of the WHO updated COVID-19 vaccination guidance for March 2023.

Author affiliations:

- ¹ School of Mathematics and Statistics, The University of Melbourne, Victoria, Australia
- ² Melbourne Centre for Data Science, The University of Melbourne, Victoria, Australia ³ Centre of Excellence for Biosecurity Risk Analysis, The University of Melbourne, Victoria, ⁸ Department of Econometrics and Business Statistics, Monash University, Victoria, Australia
- ⁴ Population Health & Immunity Division, Walter and Eliza Hall Institute of Medical Research, Victoria, Australia
- ⁵ Melbourne School of Population and Global Health, The University of Melbourne, Victoria, Australia
- ⁶ Department of Infectious Diseases at the Peter Doherty Institute for Infection and Immunity, The University of Melbourne, Victoria, Australia
- ⁷ Kirby Institute, University of New South Wales, New South Wales, Australia Australia
- ⁹ Department of Medical Biology, The University of Melbourne, Victoria, Australia ¹⁰ Telethon Kids Institute, Western Australia, Australia
- ¹¹ Victorian Infectious Diseases Reference Laboratory Epidemiology Unit at the Peter Doherty Institute for Infection and Immunity, The University of Melbourne, Victoria, Australia

