

Reconstructing the secondary infection distribution of SARS-CoV-2 from heterogeneity in viral loads and social contacts

Billy J Quilty¹ Lloyd AC Chapman¹, Kerry LM Wong¹, Amy Gimma¹, Samuel Clifford¹, Suzanne Pickering², Stuart JD Neil², Rui Pedro Galão², W John Edmunds¹, Adam J Kucharski^{1*}, Christopher I Jarvis^{1*}

¹ CMMID Covid-19 Working Group, London School of Hygiene and Tropical Medicine
² Department of Infectious Diseases, School of Immunology & Microbial Sciences, King's College London
Contact: Billy.Quilty@lshtm.ac.uk

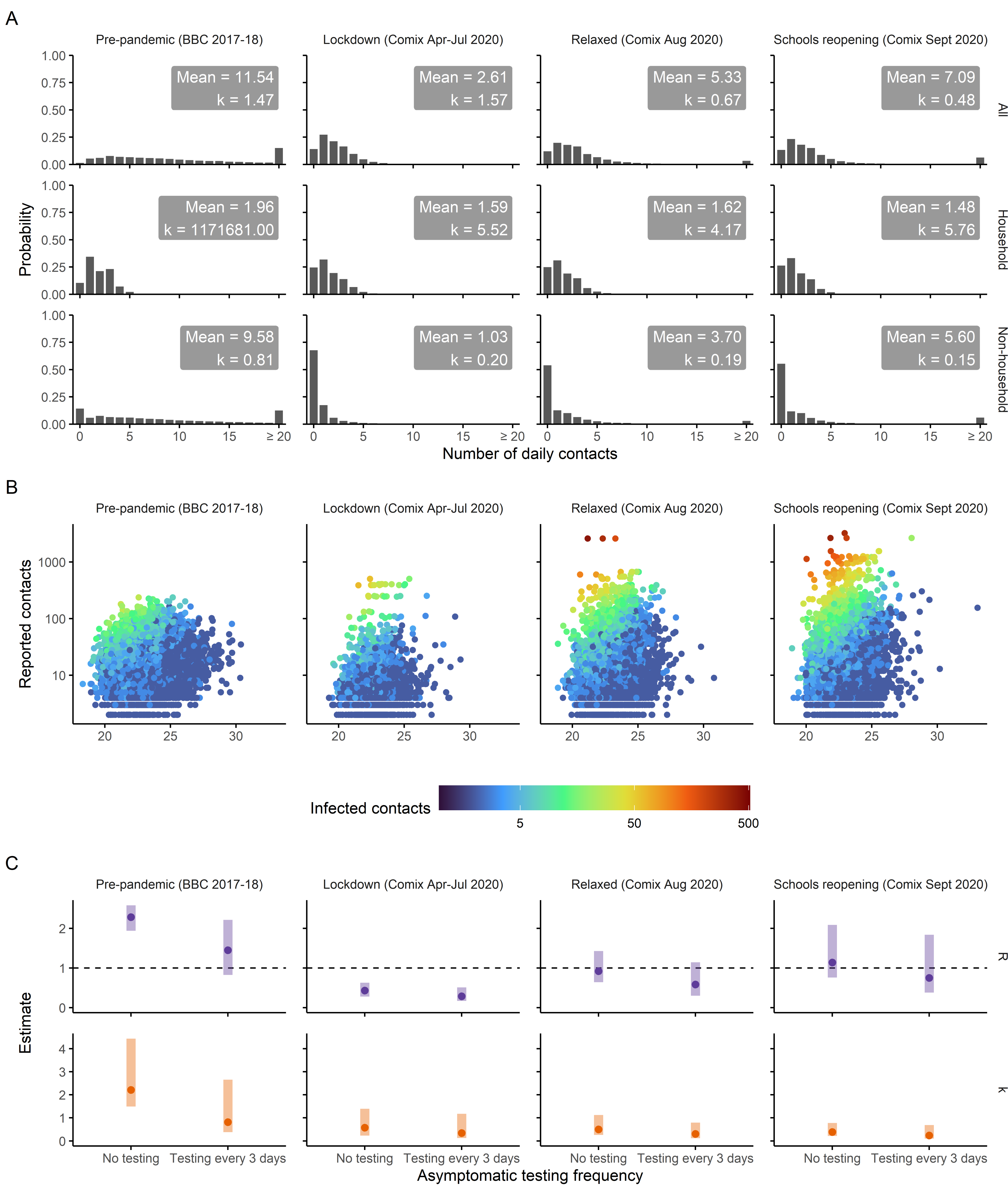
SARS-CoV-2 has been shown to spread mainly through superspreading, with ~10% of individuals responsible for ~80% of onward transmission. We use an individual-based model to combine intra- and inter-host heterogeneity in viral loads during infection with person-to-person variation in numbers of daily contacts to estimate the secondary case distribution of SARS-CoV-2. We investigate the effect of different contact rates before and during the pandemic in the UK, the effect of symptomatic self-isolation, and the impact of testing with rapid antigen tests.

Introduction

- Transmission of SARS-CoV-2 is characterised by substantial variation in the number of secondary infections generated by an infected individual, i.e., “superspreading”.
- This variability is believed to stem from variation in viral loads and social contacts between individuals and over the course of infection.
- Understanding the relative contribution of each to transmission may enable estimation of individual-level variation in the reproduction number with changes in social contacts during the SARS-CoV-2 pandemic, and the design of more finely-targeted control measures.

Methods

- Individual-based simulation model combining published data on wild-type viral load trajectories with daily social contacts reported via surveys before (BBC Pandemic, 2017-2018) and during (CoMix, 2020) the pandemic in the UK.
- Primary infectors were allocated a set of household contacts and daily out-of-household contacts.
- Infection of contacts was modelled as a stochastic process with probability of infection determined by primary case's infectious viral load on day of contact and duration of contact.
- Individuals assumed to self-isolate at home upon



symptom onset or following a positive lateral flow antigen test taken every 3 days when asymptomatic, preventing non-household contact from that day forward.

- Mean, R , and degree of overdispersion, k , in reproduction number estimated by fitting Negative Binomial distribution to resulting secondary infection distribution.

Findings

- Proportion of individuals reporting >20 contacts decreased from 14% pre-pandemic to 0.5% in the first lockdown, then increased to 3% after restrictions were relaxed and 6% when schools reopened.
- Estimated R_0 from model of 2.3 (95% UI: 1.9, 2.6) for pre-pandemic contact levels, and R_c (control reproduction number) of 0.4 (0.3, 0.6) in the first lockdown, 0.92 (0.6, 1.4) after restrictions were relaxed, and 1.1 (0.8, 2.1) when schools reopened, closely matching other estimates.
- High numbers of secondary infections were likely to occur when individuals had many contacts and a high viral load on a given day. For example when individuals had >100 reported contacts and a viral load higher than the average peak (<22.4 Ct) there was a 49% chance that >20 of those contacts would be infected.
- Rapid testing every 3 days has a greater relative impact when individuals have more contacts.

Conclusions

- SARS-CoV-2 superspreading can be reconstructed from a model of individual-level variation in viral load and daily contacts.
- Superspreading events occur when high viral loads coincide with high numbers of contacts, suggesting that testing prior to events may reduce their occurrence.
- Explicitly accounting for variation in both infectivity and contact rates may allow for targeted infection control measures with lower societal costs than blanket interventions.

Key references:

Jarvis CI, Van Zandvoort K, Gimma A, et al. Quantifying the impact of physical distance measures on the transmission of COVID-19 in the UK. *BMC Med* 2020; 18: 124.

Pickering S, Batra R, Snell LB, et al. Comparative performance of SARS CoV-2 lateral flow antigen tests demonstrates their utility for high sensitivity detection of infectious virus in clinical specimens. *medRxiv* 2021; : 2021.02.27.21252427

Endo A, Centre for the Mathematical Modelling of Infectious Diseases COVID-19 Working Group, Abbott S, Kucharski AJ, Funk S. Estimating the overdispersion in COVID-19 transmission using outbreak sizes outside China. *Wellcome Open Res* 2020; 5: 67

Chen PZ, Bobrovitz N, Premji Z, Koopmans M, Fisman DN, Gu FX. Heterogeneity in transmissibility and shedding SARS-CoV-2 via droplets and aerosols. *eLife* 2021; 10: e65774

Kucharski AJ, Klepac P, Conlan AJK, et al. Effectiveness of isolation, testing, contact tracing, and physical distancing on reducing transmission of SARS-CoV-2 in different settings: a mathematical modelling study. *Lancet Infect Dis* 2020; : S1473309920304576

Quilty BJ, Clifford S, Hellewell J, et al. Quarantine and testing strategies in contact tracing for SARS-CoV-2: a modelling study. *Lancet Public Health* 2021; published online Jan 20. DOI:10.1016/S2468-2667(20)30308-X.

Goyal A, Reeves DB, Cardozo-Ojeda EF, Schiffer JT, Mayer BT. Viral load and contact heterogeneity predict SARS-CoV-2 transmission and super-spreading events. *Walczak AM, Childs L, Forde J, editors. eLife*. 2021 Feb 23;10:e63537.