| 1 | Title page |
|----------|--|
| 2 3 | Full title: Superspreading potential of SARS-CoV-2 Delta variants under intensive disease control measures in China |
| 4 | |
| 5 6 | Shi Zhao, MPhil ^{1,2,+} , Zihao Guo, MSc ^{1,+} , Marc Ka Chun Chong, PhD ^{1,2,*,#} , Daihai He, PhD ^{4,*,#} , and Maggie H Wang, PhD ^{1,2,#} |
| 7 | |
| 8 9 | 1 JC School of Public Health and Primary Care, Chinese University of Hong Kong, Hong Kong, China |
| 10 | 2 CUHK Shenzhen Research Institute, Shenzhen, China |
| 11 | 3 Department of Applied Mathematics, Hong Kong Polytechnic University, Hong Kong, China |
| 12 | |
| 13 | + SZ and ZG are joint first authors. |
| 14 | * Correspondence to: marc@cuhk.edu.hk (MKCC), and daihai.he@polyu.edu.hk (DH). |
| 15 | # MKCC, DH, and MHW are joint senior authors. |
| 16 | |
| 17 | Email addresses of all authors |
| 18 | SZ: zhaoshi.cm ; ZG: guozihao9602@163.com ; DH: daihai.he@polyu.edu.hk ; MKCC: |
| 19 | <u>marc@cuhk.edu.hk;</u> MHW: <u>maggiew@cuhk.edu.hk</u> . |
| 20 | |
| 21 | |
| 22 | |
| 23 24 | |
| 25 | |
| 26 | |
| 27 | |
| 28 | |
| 29 | |
| 30 | |
| 31 | |
| 32 | |
| 33 | © The Author(s) 2022. Published by Oxford University Press on behalf of International Society |
| 34 | of Travel Medicine. All rights reserved. For permissions, please e-mail: |
| 35 | journals.permissions@oup.com |

50 Highlights

Given the heterogeneity in individual transmissibility, we estimated the superspreading
potential of SARS-CoV-2 Delta variants. Using case series of Delta variants in Guangdong, China,
we found 15% (95% CrI: 12, 19) of cases seeded 80% of offspring cases.

Keywords: COVID-19; superspreading; Delta variant; China; statistical modelling.

57 Main text

58 Characterizing the role of superspreading in infectious disease transmission is of public 59 health importance for informing control strategies. Concerning the superspreading potentials of 60 COVID-19¹, these risks from different SARS-CoV-2 genetic variants remain unassessed. Under 61 intensive disease control measures, the risk of outbreak may still not be eliminated despite self-62 limited transmission. In this work, we estimated the superspreading potential of Delta variants in 63 China using a well-traced and -sequenced case series dataset.

From May to June December 2021, SARS-CoV-2 Delta variants, which were first reported in India, carrying novel genetic mutations including L452R, T478K, and P681R were detected in Guangdong, China, and rapidly resulted in a local outbreak, where most of cases could be traced back to one initial seed case. Among a total of 167 cases identified, 126 (75.4%) of them were sequenced, and with information of transmission link between each other, which was presented in 2 . We extracted the number of infectees associated each infector and obtained 126 observations for the offspring case distribution including 74 terminal cases, 10 sporadic cases, and 42 infectors having at least one infectees, see Fig 1A. By considering the disease transmission as a stochastic branching

process, the classic negative binomial model with likelihood-based estimating framework was
 employed to characterize the superspreading potentials of Delta variants infected COVID-19 cases ³.

We estimated reproduction number R of 0.91 (95%CrI: 0.63, 1.36) and dispersion parameter 74 k of 0.26 (95% CrI: 0.16, 0.41), which is relatively smaller than the k estimates around 0.43 based on 75 76 the contact tracing data in 2020¹. When given the basic reproduction number of Delta variants at 5, the superspreading threshold can be calculated as the 99-th percentile of Poisson (rate = 5) 77 distribution 4 , i.e., > 10, and thus we identified two (1.6%) superspreading events (SSE) across all 78 126 one-generation case clusters. Although SSE plays less of a role when there is widespread 79 community transmission, SSEs could contribute to generate more source infectors during the early 80 phase of an outbreak, which is less sensitive to the settings with intense public health measures at 81 population scale. We estimated that 15% (95%CrI: 12, 19) of the most infectious cases seeded 80% 82 of all offspring cases, which suggested a higher superspreading potential than the situations of early 83 outbreaks in 2020⁴. 84

Although intensive COVID-19 control measure were implemented in mainland China as of 85 2020², the variation in individual infectious history and in the properties of SARS-CoV-2 strains 86 may still contribute to large case clusters dominated by SSEs³. With one seed case of Delta variant, 87 there were 5% (95CrI: 1, 14) or 2% (95%CrI: 0, 12) of chance to have a cluster of cases with size > 88 89 30 or 100, respectively, see Fig 1B. As such, we recommend enforcing the intervention programs to monitor and limit COVID-19 transmission, especially regarding the contexts where SSEs are likely 90 to occur, e.g., restaurants and entertainment sites. For the challenges from emerging Omicron variant 91 against the global COVID-19 pandemic control, we urge concerns on the possible risks of 92 93 superspreading when it invades places with low herd immunity in population or settings with largescale congregations ⁵. 94

95

96 **Declarations**

97 Ethics approval and consent to participate

The COVID-19 number of cases and sequencing data are collected, and thus neither ethical approval
 nor individual consent is applicable.

100 Availability of materials

101 All data used in this work are publicly available via Li *et al* 2 , and the computer codes used for 102 statistical analysis may be available based on request to the authors.

103 **Consent for publication**

- 104 Not applicable.
- 105 Funding
- 106 MKCC is partially supported by National Natural Science Foundation of China (NSFC) [71974165].
- 107 DH is partially supported by a grant from the Research Grants Council of the Hong Kong Special
- Administrative Region, China [HKU C7123- 20G]. MHW is supported by Health and Medical
- 109 Research Fund [INF-CUHK-1, COVID190103], the Food and Health Bureau, Hong Kong, China,
- 110 CUHK grant [PIEF/Ph2/COVID/06, 4054456], and partially supported by the National Natural
- 111 Science Foundation of China (NSFC) [31871340].

- 112 Acknowledgements
- 113 None.
- 114 Conflict of interests
- 115 MHW is a shareholder of Beth Bioinformatics Co., Ltd. Other authors declared no competing
- interests. The funding agencies had no role in the design and conduct of the study; collection,
- 117 management, analysis, and interpretation of the data; preparation, review, or approval of the
- 118 manuscript; or decision to submit the manuscript for publication.
- 119 Authors' contributions
- 120 Conceptualization: SZ. Methodology: SZ, ZG, and MKCC. Software: SZ, and ZG. Validation: SZ,
- 121 and ZG. Formal analysis: SZ. Investigation: SZ. Resources: SZ. Data Curation: SZ. Writing
- 122 Original Draft: SZ. Writing Review and Editing: ZG, and MKCC, DH, and MHW. Visualization:
- 123 SZ. Supervision: MKCC, and MHW. Project Administration: SZ. Funding acquisition: MKCC, and
- 124 MHW. All authors critically read the manuscript, and gave final approval for publication.
- 125
- 126

127 **References**

Wang J, Chen X, Guo Z, et al. Superspreading and heterogeneity in transmission of sars,
 mers, and covid-19: A systematic review. Computational and Structural Biotechnology Journal 2021;
 19(5039-5046.

Li B, Deng A, Li K, et al. Viral infection and transmission in a large, well-traced outbreak
caused by the sars-cov-2 delta variant. Nature Communications 2022; 13(1):460.

Lim JS, Noh E, Shim E, Ryu S. Temporal changes in the risk of superspreading events of
 coronavirus disease 2019. Open Forum Infectious Diseases 2021; 8(7):ofab350.

Adam DC, Wu P, Wong JY, et al. Clustering and superspreading potential of sars-cov-2
infections in hong kong. Nat Med 2020; 26(11):1714-1719.

Mat NFC, Edinur HA, Razab MKAA, Safuan S. A single mass gathering resulted in massive
 transmission of COVID-19 infections in Malaysia with further international spread. J Travel Med.
 2020; 27(3):taaa059.

140

142 Figures



The observed and fitted offspring cases distribution (A) and estimated probability of generating case cluster with a given size (B). Panel (A) shows observed (in pink) and fitted (in blue) offspring cases distribution. Panel (B) shows probability of observing a case cluster with a size equal to or larger than a given number that is generated by one seed case. In both panels, the light blue curves are 100 posterior MCMC samples, and dark blue curve is the posterior median.

- 150
- 151 Tables
- 152 None.

Cort

153