

# 1 What have we learnt from mass testing for COVID-19 in universities?

2 Nicholas J Matheson, principal investigator<sup>1,2,3,\*</sup>, Ben Warne, clinical research fellow<sup>1,2</sup>,  
3 Michael P. Weekes, principal investigator<sup>1,2,4</sup>, Patrick H. Maxwell, regius professor of  
4 physic<sup>2,4,5</sup>

5

- 6 1. Cambridge Institute for Therapeutic Immunology and Infectious Disease (CITIID),  
7 Cambridge, UK
- 8 2. Cambridge University Hospitals, Cambridge, UK
- 9 3. NHS Blood and Transplant, Cambridge, UK
- 10 4. Cambridge Institute for Medical Research (CIMR), Cambridge, UK
- 11 5. University of Cambridge School of Clinical Medicine, Cambridge, UK

12

13 \*Correspondence: Nicholas J Matheson ([njm25@cam.ac.uk](mailto:njm25@cam.ac.uk))

14

15 Word count (main text, excluding references in superscript): 798

16 It is clear that SARS-CoV-2 can be incubated and transmitted in the absence of symptoms<sup>1</sup>.  
17 Nonetheless, the utility of mass testing (large-scale asymptomatic screening, to  
18 prospectively identify cases) has been contested<sup>2,3</sup>. In *principle*, isolation of individuals with  
19 presymptomatic or asymptomatic SARS-CoV-2 infection will prevent further infections as a  
20 matter of course. In *practice*, it is less clear whether enough infectious individuals can be  
21 identified to have a quantitatively important impact on transmission, and whether the direct  
22 benefits of enhanced case ascertainment may be outweighed by direct or indirect costs. The  
23 debate is complicated by an absence of randomised controlled trial data, and controversy  
24 about the suitability of lateral flow tests (LFTs)<sup>4,5</sup>.

25 Students in higher education are at increased risk of SARS-CoV-2 infection, because of their  
26 shared accommodation, abundant social contacts, low priority for vaccination, and potential  
27 for vaccine hesitancy<sup>6 7</sup>. At the same time, universities have been at the forefront of research  
28 on COVID-19. It is therefore instructive to examine how they have sought to control  
29 transmission amongst their own students. Strikingly, as well as promoting vaccination,  
30 symptomatic testing and contact tracing, many universities in the UK and North America  
31 have chosen to implement asymptomatic COVID-19 screening programmes, using weekly or  
32 twice-weekly laboratory-based PCR tests. Data from these programmes are now available  
33 from institutional websites, pre-prints and peer-reviewed manuscripts. We may therefore  
34 ask: what can they teach us about mass testing for SARS-CoV-2?

35 First, it is possible to sustain high levels of adherence to regular, voluntary asymptomatic  
36 screening using nose and throat swabs<sup>8 9</sup>. University-led testing programmes have been  
37 strongly supported by students<sup>8 10 11</sup>, providing reassurance at a time when student mental  
38 health and wellbeing has been severely impacted by the pandemic<sup>8 12</sup>.

39 Second, mass testing can markedly increase case ascertainment, including a substantial  
40 proportion of individuals with presymptomatic SARS-CoV-2 infection (before they develop  
41 symptoms)<sup>9 13-15</sup>. Remarkably, for some university communities and stages of the pandemic,  
42 more students with SARS-CoV-2 have been detected by asymptomatic screening than by  
43 symptomatic testing<sup>14-16</sup>. Provided they are supported to self-isolate, it is reasonable to infer  
44 a very substantial reduction in ongoing transmission.

45 Third, PCR testing is ideally suited to *regular screening of defined populations*, where high  
46 test sensitivity minimises the risk of “false negatives”, and samples are available for genomic  
47 sequencing<sup>17</sup>. In a university context, laboratory and logistical infrastructure can be planned  
48 in advance, turnaround time minimised, and swab or sample pooling used to reduce costs  
49 and demands on testing capacity (particularly when incidence is low)<sup>9 15 18</sup>.

50 And fourth, the impact of “false positives” on many programmes can be mitigated by a two-  
51 step testing strategy, whereby a positive screening test is followed routinely by a second,  
52 confirmatory PCR test<sup>9</sup>. Regular, frequent screening is also essential to ensure that infected  
53 individuals are detected early, whilst they are still infectious – so that self-isolation is justified.

54 What, then, are the remaining unknowns – and how can success be measured? Evidence  
55 about secondary behavioural changes, which may partially offset the benefits of enhanced  
56 case detection, remains very limited. This is a particular concern for programmes based on  
57 LFTs, because “false negatives” are more common, and clearly documented examples of  
58 sustained, high levels of adherence to twice-weekly home testing are lacking<sup>19</sup>. In addition,  
59 the impact on participation of increasing levels of vaccination remains to be determined. As a  
60 minimum, it is therefore critical for screening programmes to monitor both participation rates  
61 (the number, proportion and frequency of individuals screened) and the fraction of all cases  
62 ascertained by mass testing.

63 Countries with high levels of vaccination are generally rolling back non-pharmaceutical  
64 interventions designed to limit case numbers, such as social distancing and face masks. At  
65 the same time, the relative benefit of identification and isolation of *contacts* has been  
66 reduced, because secondary attack rates are lower when index cases and/or contacts have  
67 been vaccinated<sup>20 21</sup>. Nonetheless, the development of novel SARS-CoV-2 variants means  
68 that large outbreaks may still occur in vaccinated populations<sup>22</sup>. Compared with other non-  
69 pharmaceutical interventions, asymptomatic screening offers a number of advantages.  
70 Critically, it is focused on the identification and isolation of *cases*, rather than *contacts*;  
71 provided testing is informed and voluntary, there need be no impact on the freedom of  
72 individuals; and finally, the costs are direct and quantifiable on a per programme basis, with  
73 few indirect economic consequences.

74 Presuming *any* measures to control SARS-CoV-2 transmission are required, there is  
75 therefore a strong argument for mass testing of populations at high risk of infection – such as  
76 students in higher education. Accordingly, faced with spread of the delta variant, many  
77 universities have committed to continue their programmes of regular PCR-based  
78 asymptomatic screening. When prevalence declines, surveillance testing (regular screening  
79 of a proportion of the population) and genomic sequencing (for new variants of concern) may  
80 be a proportionate response – and universities will again be ideal laboratories to test the  
81 coherence and effectiveness of these approaches.

## 82 Acknowledgements

83 NJM receives funding from the MRC (CSF MR/P008801/1) and NHS Blood and Transplant  
84 (WPA15-02). MPW receives funding from the Wellcome Trust (SCF 108070/Z/15/Z). BW  
85 receives funding from the NIHR Cambridge Biomedical Research Centre at the Cambridge  
86 University Hospitals NHS Foundation Trust. The views expressed are those of the authors  
87 and not necessarily those of the NHS, the NIHR or the Department of Health and Social  
88 Care.

## 89 Competing interests

90 We have read and understood the BMJ Group policy on declaration of interests and declare  
91 the following interests: none.

## 92 Copyright statement

93 The Corresponding Author has the right to grant on behalf of all authors and does grant on  
94 behalf of all authors, a worldwide licence to the Publishers and its licensees in perpetuity, in  
95 all forms, formats and media (whether known now or created in the future), to i) publish,  
96 reproduce, distribute, display and store the Contribution, ii) translate the Contribution into

97 other languages, create adaptations, reprints, include within collections and create  
98 summaries, extracts and/or, abstracts of the Contribution, iii) create any other derivative  
99 work(s) based on the Contribution, iv) to exploit all subsidiary rights in the Contribution, v)  
100 the inclusion of electronic links from the Contribution to third party material where-ever it may  
101 be located; and, vi) licence any third party to do any or all of the above.

## 102 References

- 103 1. Buitrago-Garcia D, Egli-Gany D, Counotte MJ, et al. Occurrence and transmission  
104 potential of asymptomatic and presymptomatic SARS-CoV-2 infections: A living  
105 systematic review and meta-analysis. *PLoS Med* 2020;17(9):e1003346. doi:  
106 10.1371/journal.pmed.1003346 [published Online First: 2020/09/23]
- 107 2. Deeks JJ, Brookes AJ, Pollock AM. Operation Moonshot proposals are scientifically  
108 unsound. *BMJ* 2020;370:m3699. doi: 10.1136/bmj.m3699 [published Online First:  
109 2020/09/24]
- 110 3. Mina MJ, Peto TE, Garcia-Finana M, et al. Clarifying the evidence on SARS-CoV-2  
111 antigen rapid tests in public health responses to COVID-19. *Lancet*  
112 2021;397(10283):1425-27. doi: 10.1016/S0140-6736(21)00425-6 [published Online  
113 First: 2021/02/21]
- 114 4. Deeks JJ, Raffle AE. Lateral flow tests cannot rule out SARS-CoV-2 infection. *BMJ*  
115 2020;371:m4787. doi: 10.1136/bmj.m4787 [published Online First: 2020/12/15]
- 116 5. Taylor-Phillips S, Dinnes J. Asymptomatic rapid testing for SARS-CoV-2. *BMJ*  
117 2021;374:n1733. doi: 10.1136/bmj.n1733 [published Online First: 2021/07/09]
- 118 6. Hill EM, Atkins BD, Keeling MJ, et al. Modelling SARS-CoV-2 transmission in a UK  
119 university setting. *medRxiv* 2021:2020.10.15.20208454. doi:  
120 10.1101/2020.10.15.20208454

- 121 7. Troiano G, Nardi A. Vaccine hesitancy in the era of COVID-19. *Public Health*  
122 2021;194:245-51. doi: 10.1016/j.puhe.2021.02.025 [published Online First:  
123 2021/05/10]
- 124 8. Blake H, Corner J, Cirelli C, et al. Perceptions and Experiences of the University of  
125 Nottingham Pilot SARS-CoV-2 Asymptomatic Testing Service: A Mixed-Methods  
126 Study. *Int J Environ Res Public Health* 2020;18(1) doi: 10.3390/ijerph18010188  
127 [published Online First: 2021/01/02]
- 128 9. Warne B, Metaxaki M, Fuller S, et al. Feasibility and efficacy of mass testing for SARS-  
129 CoV-2 in a UK university using swab pooling and PCR. *Research Square* 2021 doi:  
130 10.21203/rs.3.rs-520626/v1
- 131 10. Cox C, Ansari A, McLaughlin M, et al. Developing an ethical framework for asymptomatic  
132 COVID-19 testing programmes in higher education institutions: Wellcome Open Res,  
133 2021.
- 134 11. Gillam TB, Cole J, Gharbi K, et al. Norwich COVID-19 testing initiative pilot: evaluating  
135 the feasibility of asymptomatic testing on a university campus. *J Public Health (Oxf)*  
136 2021;43(1):82-88. doi: 10.1093/pubmed/fdaa194 [published Online First: 2020/10/31]
- 137 12. National Union of Students: Coronavirus Student Survey phase III November 2020 -  
138 Mental health and wellbeing [Available from:  
139 [https://www.nusconnect.org.uk/resources/coronavirus-and-students-phase-3-study-  
140 mental-health-with-demographics-nov-2020](https://www.nusconnect.org.uk/resources/coronavirus-and-students-phase-3-study-mental-health-with-demographics-nov-2020).
- 141 13. Hamer DH, White LF, Jenkins HE, et al. Assessment of a COVID-19 Control Plan on an  
142 Urban University Campus During a Second Wave of the Pandemic. *JAMA Netw*  
143 *Open* 2021;4(6):e2116425. doi: 10.1001/jamanetworkopen.2021.16425 [published  
144 Online First: 2021/06/26]
- 145 14. Rennert L, McMahan C, Kalbaugh CA, et al. Surveillance-based informative testing for  
146 detection and containment of SARS-CoV-2 outbreaks on a public university campus:  
147 an observational and modelling study. *Lancet Child Adolesc Health* 2021 doi:  
148 10.1016/S2352-4642(21)00060-2 [published Online First: 2021/03/23]

- 149 15. Denny TN, Andrews L, Bonsignori M, et al. Implementation of a Pooled Surveillance  
150 Testing Program for Asymptomatic SARS-CoV-2 Infections on a College Campus -  
151 Duke University, Durham, North Carolina, August 2-October 11, 2020. *MMWR Morb*  
152 *Mortal Wkly Rep* 2020;69(46):1743-47. doi: 10.15585/mmwr.mm6946e1 [published  
153 Online First: 2020/11/20]
- 154 16. University of Cambridge: Data from the COVID-19 testing service [Available from:  
155 [https://www.cam.ac.uk/coronavirus/stay-safe-cambridge-uni/data-from-covid-19-](https://www.cam.ac.uk/coronavirus/stay-safe-cambridge-uni/data-from-covid-19-testing-service)  
156 [testing-service](https://www.cam.ac.uk/coronavirus/stay-safe-cambridge-uni/data-from-covid-19-testing-service).
- 157 17. Aggarwal D, Warne B, Jahun AS, et al. Genomic epidemiology of SARS-CoV-2 in a UK  
158 university identifies dynamics of transmission. *Research Square* 2021 doi:  
159 10.21203/rs.3.rs-520627/v1
- 160 18. University of Edinburgh: TestEd [Available from: <https://www.ed.ac.uk/tested-covid>.
- 161 19. Iacobucci G. Covid-19: Mass testing at UK universities is haphazard and unscientific,  
162 finds BMJ investigation. *BMJ* 2021;372:n848. doi: 10.1136/bmj.n848 [published  
163 Online First: 2021/04/02]
- 164 20. Harder T, Koch J, Vygen-Bonnet S, et al. Efficacy and effectiveness of COVID-19  
165 vaccines against SARS-CoV-2 infection: interim results of a living systematic review,  
166 1 January to 14 May 2021. *Euro Surveill* 2021;26(28) doi: 10.2807/1560-  
167 7917.ES.2021.26.28.2100563 [published Online First: 2021/07/17]
- 168 21. Harris RJ, Hall JA, Zaidi A, et al. Effect of Vaccination on Household Transmission of  
169 SARS-CoV-2 in England. *N Engl J Med* 2021 doi: 10.1056/NEJMc2107717  
170 [published Online First: 2021/06/24]
- 171 22. Brown CM, Vostok J, Johnsho H, et al. Outbreak of SARS-CoV-2 Infections, Including  
172 COVID-19 Vaccine Breakthrough Infections, Associated with Large Public  
173 Gatherings — Barnstable County, Massachusetts, July 2021. *MMWR Morb Mortal*  
174 *Wkly Rep* 2021 doi: <http://dx.doi.org/10.15585/mmwr.mm7031e2> [published Online  
175 First: 30 July 2021]