

Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active. of National Health Service (NHS) staff during the epidemic. The NHS staff testing policy was only to test symptomatic staff, precisely to reduce absenteeism by encouraging staff with negative results back to work, thus intentionally reducing their time in self-isolation. The Secretary of State for Health and Social Care, Matt Hancock, himself stated that "we want to get [NHS staff absences] down, and the way to do that is to get the amount of testing up".<sup>2</sup> This testing approach was then also applied to other groups of public sector workers.<sup>3</sup>

The UK Government's approach of using SARS-CoV-2 testing as a strategy to reduce absenteeism rather than to increase the detection of otherwise asymptomatic spreaders was surely symptomatic of flawed analysis and misunderstanding of the utility of the SARS-CoV-2 pharyngeal swab RT-PCR test. WHO expressly advises against using this test as a rule-out in the event of negative results.4 Sensitivity of the test might be as low as 83%,<sup>5</sup> and in our practice many colleagues believe it to be lower still. Overzealous redirection of self-isolating staff back to work before they had completed sufficient self-isolation to exclude infectivity was therefore likely to increase spread of the virus to other staff and to patients or care-receivers in a substantial number of cases, especially given the high prevalence and likelihood of SARS-CoV-2 infection among exposed health-care workers during the epidemic. Surely the only defensible policy would have been national opportunistic and frequent testing of NHS and social care sector staff regardless of symptomology, and the test should be used exclusively as a rule-in and not a rule-out test as per existing WHO guidance.<sup>4</sup>

I declare no competing interests.

Bernard Freudenthal b.freudenthal@imperial.ac.uk Clinical Research Fellow, Imperial College London, London, W12 ONN, UK

- Black JRM, Bailey C, Przewrocka J, Dijkstra KK, Swanton C. COVID-19: the case for health-care worker screening to prevent hospital transmission. Lancet 2020; 395: 1418–20.
- Booth R. Number of NHS doctors off sick 'may be nearly triple the official estimate'. April 5, 2020. https://www.theguardian.com/ world/2020/apr/05/number-nhs-doctors-offsick-may-be-three-times-more-than-thought (accessed April 19, 2020).
- 3 The Guardian. UK Covid-19 testing expanded to police, fire service and judiciary. April 17, 2020. https://www.theguardian.com/ world/2020/apr/17/uk-covid-19-testingexpanded-to-police-fire-service-and-judiciary. (accessed April 19, 2020).
- 4 WHO. Laboratory testing for coronavirus disease (COVID-19) in suspected human cases. March 19, 2020. https://www.who.int/ publications-detail/laboratory-testing-for-2019-novel-coronavirus-in-suspected-humancases-20200117 (accessed April 19, 2020).
- Long C, Xu H, Shen Q, et al. Diagnosis of the Coronavirus disease (COVID-19): rRT-PCR or CT? Eur J Radiol 2020; **126:** 108961.

## **Authors' reply**

We thank Bernard Freudenthal for his response to our previous Correspondence.<sup>1</sup> We agree that use of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) testing among health-care workers (HCWs) solely to reduce absenteeism is inappropriate. Freudenthal correctly outlines the risks, posed by falsenegative results, of advising potentially infectious HCWs to return to work. Moreover, staffing levels are currently far less problematic within UK health-care settings than during the peak of the pandemic.

HCW testing should aim to identify infectious cases and reduce nosocomial transmission of SARS-CoV-2: testing only selfreported symptomatic cases risks missing many infectious cases. For instance, HCWs might unwittingly attend work with mild or non-specific symptoms. Furthermore, although the relationship between RT-PCR cycle threshold (Ct) values and infectivity requires further elucidation, evidence suggests that Ct values among asymptomatic and symptomatic cases are similar.<sup>2</sup> Crucially, viable virus has been isolated up to 6 days before symptom onset.<sup>3</sup>

Robust epidemiological studies help detail asymptomatic spread. Results have been heterogeneous; assumptions vary between studies which might be subject to recall bias, definitions of symptoms are inconsistent, and some studies do not account for the critical presymptomatic phase of infection. Nonetheless, most such studies find evidence of asymptomatic SARS-CoV-2 transmission.<sup>4</sup>

False-positive results can also limit HCW screening utility. They can be biological, with dead virus detected in non-infectious cases, and technical, where a test is positive in the absence of viral RNA. Regular screening risks identification of biological false positives; however, more research is required to understand the biology of persistent viral RNA shedding. Technical false positives might be reduced to manageable levels by testing in duplicate.<sup>5</sup>

We believe a symptom-agnostic testing approach for SARS-CoV-2 among HCWs is an effective measure of reducing viral transmission. This approach is advocated on a population level<sup>6</sup> and might be particularly beneficial among HCWs given reports of hospitals acting as hotbeds of COVID-19.

Arguments against mass testing approaches previously have suggested a lack of resources might make this ineffective. However, UK daily testing capacity has increased tenfold since the publication of our Correspondence,<sup>1</sup> while rapid point-of-care antigen tests facilitate early intervention to limit transmission.<sup>6</sup>

Screening for SARS-CoV-2 in asymptomatic HCWs could be a vital weapon in the fight against COVID-19 now and over the winter months. This will help the National Health Service to maintain the capacity to treat other diseases in the face of a second wave. We must act to prevent further virus spread, economic disruption, and unnecessary death.



CS reports grants from BMS, Ono-Pharmaceuticals, and Archer Dx (collaboration in minimal residual disease sequencing technologies), outside the submitted work; personal fees from Bristol Myers Squibb, Roche-Ventana, Ono Pharmaceutical, GlaxoSmithKline, Novartis, Celgene, Illumina, MSD, Sarah Canon Research Institute, Genentech, Bicycle Therapeutics, and Medicixi, outside the submitted work; personal fees and stock options from GRAIL and Achilles Therapeutics, outside the submitted work; and stock options from Epic Biosciences and Apogen Biotechnologies, outside the submitted work. All other authors declare no competing interests.

James R M Black, Chris Bailey, Joanna Przewrocka, Krijn K Dijkstra, Sonia Gandhi, Steve Gamblin, Sam Barrell, \*Charles Swanton charles.swanton@crick.ac.uk

Cancer Genome Evolution Research Group (JRMB) and Cancer Research UK Lung Cancer Centre of Excellence (CS), University College London, London, UK; Cancer Evolution and Genome Instability Laboratory, Francis Crick Institute, London NW1 1AT, UK (CB, JP, KKD, CS); Francis Crick Institute, London, UK (SGan, SGam, SB); and University College London Hospitals NHS Trust, London, UK (CS)

- 1 Black JRM, Bailey C, Przewrocka J, Dijkstra KK, Swanton C. COVID-19: the case for health-care worker screening to prevent hospital transmission. *Lancet* 2020; **395**: 1418–20.
- 2 Long Q-X, Tang X-J, Shi Q-L, et al. Clinical and immunological assessment of asymptomatic SARS-CoV-2 infections. *Nat Med* 2020; 26: 1200–04.
- 3 Arons MM, Hatfield KM, Reddy SC, et al. Presymptomatic SARS-CoV-2 infections and transmission in a skilled nursing facility. N Engl J Med 2020; 382: 2081–90.
- 4 Buitrago-Garcia DC, Egli-Gany D, Counotte MJ, et al. Asymptomatic SARS-CoV-2 infections: a living systematic review and meta-analysis. medRxiv 2020; published online July 28. https://doi.org/10.1101/2020.04.25.20079103 (preprint).
- 5 Aitken J, Ambrose K, Barrell S, et al. Scalable and robust SARS-CoV-2 testing in an academic center. Nat Biotechnol 2020; 38: 927–31.
- 6 Wain R, Sleat D. Taking the UK's testing strategy to the next level. Aug 17, 2020. https://institute.global/sites/default/files/ articles/Taking-the-UK-s-Testing-Strategy-tothe-Next-Level.pdf (accessed Aug 19, 2020).

## Accelerated surgery for hip fractures—the HIP ATTACK results discussed

The HIP ATTACK investigators<sup>1</sup> found that accelerated surgery (median 6 h) did not lower the risk of mortality or a composite of major complications compared with standard care (median 24 h) but reduced the risk of delirium, urinary tract infection, pain, and length of hospital stay. We have two concerns.

First, substantial changes in practice would be required to implement the recommendation. We have previously suggested that costs incurred by waiting might provide a financial incentive to mitigate delays for surgery.<sup>2</sup> However, hip fracture numbers are increasing, and in Ontario, Canada (where 898 patients or 30% of participants in this trial were enrolled) less than 5% of patients with hip fracture have surgery within 6 h, and less than 1% have surgery overnight.<sup>3</sup> That those who presented outside regular working hours (16 082 [58%] of 27 701 participants) had to be excluded from a highly funded trial might also be evidence that surgery within 6 h for most patients might not be achievable.

Scheduling must also be balanced with the needs of patients waiting for other operations, which might be delayed if hip fractures are singularly prioritised.

Another unexpected consequence might be the increasing provision of surgery overnight. Whereas surgery on evenings and weekends appears to be safe, quality of overnight hip fracture surgery has not been assessed.<sup>4</sup>

Second, the investigators acknowledge "results primarily inform the effects for patients who went to surgery a median of 6 h versus 24 h,"1 but conclusions are not framed within the existing literature to help inform clinical decision making and policy. Population-based studies attempting to triangulate a timethreshold for surgery found no risk difference between 6 h and 24 h because mortality and complications only began to increase 24 h after hip fracture.<sup>5,6</sup> Did the HIP ATTACK investigators consider delays much longer than 24 h in the standard care-high troponin subgroup as an explanation of the interaction between

troponin measurement and the effect of accelerated surgery? The median time-to-surgery for hip fracture is 28 h in Ontario, Canada, compared to 24 h in the trial: did the investigators consider a Hawthorne effect that improved standard of care for trial participants as an explanation for the null findings?

We caution against dismissing observational studies as confounded when it is impractical (and possibly unethical) to allocate patients to increasing thresholds of delayed surgery in a trial. Perhaps those who care for hip fracture patients might "weigh the potential reduction in delirium and length of hospital stay against organising an accelerated [within 6 h] surgery pathway<sup>"1</sup> in addition to the potential reduction in mortality and major medical complications against organising a (within 24 h) surgery pathway.

We declare no competing interests.

## \*Daniel Pincus, Bheeshma Ravi daniel.pincus@sunnybrook.ca

University of Toronto Faculty of Medicine, Toronto, ON M5S 1A8, Canada

- The HIP ATTACK Investigators. Accelerated surgery versus standard care in hip fracture (HIP ATTACK): an international, randomised, controlled trial. Lancet 2020; 395: 698–708.
- 2 Pincus D, Wasserstein D, Ravi B, et al. Medical costs of delayed hip fracture surgery. JBJS 2018; 100: 1387–96.
- 3 Pincus D, Wasserstein D, Ravi B, et al. Reporting and evaluating wait times for urgent hip fracture surgery in Ontario, Canada. CMAJ 2018; 190: 702–09.
- 4 Pincus D, Desai SJ, Wasserstein D, et al. Outcomes of after-hours hip fracture surgery. JBJS 2017; **99:** 914–22.
- 5 Pincus D, Ravi B, Wasserstein D, et al. Association between wait time and 30-day mortality in adults undergoing hip fracture surgery. JAMA 2017; **318**: 1994–2003.
- 6 Sobolev B, Guy P, Sheehan KJ, et al. Mortality effects of timing alternatives for hip fracture surgery. *CMAJ* 2018; **190**: 923–32.

We would like to make several points regarding the HIP ATTACK trial results.<sup>1</sup>

The study shows the need to base health policy and structural organisation of medical care on randomised controlled trials. Retrospective, observational data are