Rapid evidence review to inform safe return to campus in the context of coronavirus disease 2019 (COVID-19) [version 1; peer review: 2 approved, 1 approved with reservations]

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Abstract

Background: Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) is transmitted predominantly through the air in crowded and unventilated indoor spaces among unvaccinated people. Universities and colleges are potential settings for its spread.

Methods: An interdisciplinary team from public health, virology, and biology used narrative methods to summarise and synthesise evidence on key control measures, taking account of mode of transmission.

Results: Evidence from a wide range of primary studies supports six measures. Vaccinate (aim for > 90% coverage and make it easy to get a jab). Require masks indoors, especially in crowded settings. If everyone wears well-fitting cloth masks, source control will be high, but for maximum self-protection, respirator masks should be worn. Masks should not be removed for speaking or singing. Space people out by physical distancing (but there is no “safe” distance because transmission risk varies with factors such as ventilation, activity levels and crowding), reducing class size (including offering blended learning), and cohorting (students remain in small groups with no cross-mixing). Clean indoor air using engineering controls—ventilation (while monitoring CO2 levels), inbuilt filtration systems, or portable air cleaners fitted with high efficiency particulate air [HEPA] filters). Test asymptomatic staff and students using lateral flow tests, with tracing and isolating infectious cases when incidence of coronavirus disease 2019 (COVID-19) is high. Support clinically vulnerable people to work remotely. There is no direct evidence to support hand sanitising, fomite controls or temperature-taking. There is evidence that freestanding plastic screens, face visors and electronic air-cleaning systems are ineffective.

Conclusions: The above six evidence-based measures should be combined into a multi-faceted strategy to maximise both student...
safety and the continuation of in-person and online education provision. Staff and students seeking to negotiate a safe working and learning environment should collect data (e.g. CO₂ levels, room occupancy) to inform conversations.

Keywords
SARS-CoV-2, aerosol transmission, hierarchy of controls, higher education policy, infection prevention and control

This article is included in the Coronavirus (COVID-19) collection.

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Introduction

Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) and university life

The United Kingdom (UK) is currently (Autumn 2021) experiencing high and rising levels of coronavirus disease 2019 (COVID-19) cases, and almost all are the highly contagious delta variant. This variant spread fastest among the 17–24 year age group in June and July 2021, likely due to a combination of low vaccination rates in this age group and suboptimal mitigation strategies in schools and colleges. Whilst young people are much less likely to develop severe acute disease from COVID-19 than older people, some will be hospitalised and a few could die. The incidence of persistent symptoms beyond the acute illness (post-acute or “long” Covid) is disputed, but a secondary analysis of Office of National Statistics data (published as a preprint) suggests that 4.7% of the 18–24 year age group have some symptoms persisting beyond 12 weeks and 1.1% have symptoms which interfere “a lot” with their daily activities. University staff and graduate students include older age groups, minority ethnic groups and those with medical conditions, all of which increase the risk of developing serious complications from COVID-19.

For all these reasons, measures to reduce transmission of the virus are needed. Most universities and colleges in the UK now have the infrastructure to implement rapid and frequent testing and support students to isolate when necessary. Since most courses were delivered online in 2020–21, they have also learnt a great deal about how to deliver effective learning online. There is, however, a considerable appetite to return to face-to-face modes of teaching as well as return to traditional levels of socialising, arts and sport.

The science of SARS-CoV-2 transmission

There is strong and consistent evidence that the main—and perhaps the only significant—mode of transmission of SARS-CoV-2 is through the air. Indeed, super-spreader events (in which one or a few people infect large numbers of others)—including choir practices, funerals, conferences, gym sessions and other mass indoor events—are likely to be the main drivers of the pandemic. Higher education includes many preconditions for such super-spreader events, including living and eating communally, lectures and seminars, sports training and competition, arts and singing performances, and socialising.

Some indoor events show no COVID-19 transmission, even when infected people are present, while others are shown in retrospect to have been super-spreader events; this phenomenon is known as heterogeneity or overdispersion of transmission dynamics, and is highly relevant to our efforts to control the virus in schools and universities. Whilst the highest risk of airborne viral transmission occurs with coughing and sneezing, speaking and singing are also high-risk activities.

A landmark paper on minimising risk of airborne transmission (written before effective vaccines had been discovered) used their own adaptation of the US Centers for Disease Control and Prevention’s “hierarchy of controls” (Figure 1).

In the sections which follow, we consider all the measures in the hierarchy of controls plus vaccination. We begin with vaccination, masking and administrative controls as these are things which individual university employees may be able to influence. We then discuss engineering controls (ventilation and air filtration).

Methods

This review, which aimed to produce a synthesis rapidly in time for the new university term, was undertaken in September 2021. We began with sources known to the authors, including a rapid review by Independent SAGE, and a search of the PubMed database using the terms “SARS-CoV-2”, “COVID-19”, “transmission”, “mitigation”, “school[s]” and “university/ies” (restricted to review articles). Using a method previously shown to be highly efficient for identifying key studies from complex and heterogeneous datasets, we initially focused on seminal papers (in high-impact journals and highly-cited for their age) and used snowball searching (tracking the article in Google Scholar and pursuing relevant sources from its reference lists) to identify further key studies from these. For specialist subsections that were beyond our own expertise, we undertook further key word searches (e.g. HEPA filters) and consulted with experts in the field. In producing our narrative synthesis of these sources, we prioritised findings that would be useful to inform actual policies in universities.
Main findings
Direct empirical evidence on mitigation strategies specifically for universities was sparse, but there was much relevant evidence on mitigation measures more generally. Our review suggested that six key findings, which we consider in turn below, are likely to be effective at reducing on-campus spread: vaccination, masking, spacing people out (physical distancing, reducing class size and cohorting), engineering controls (ventilation or filtration of air), a test/trace/isolate policy when COVID-19 incidence is high, and supporting clinically vulnerable people to work remotely. Other widely-promoted measures including sanitising hands, taking temperatures, plastic screens and face visors, were not supported by evidence.

Encourage vaccination—and make it easy
Vaccines have been a game-changer for COVID-19; they dramatically reduce the incidence of symptomatic disease and risk of transmission of the virus to others; breakthrough infections in vaccinated persons are rare and generally mild1. A recent BMJ review concluded that the most important single intervention for preventing on-campus transmission is vaccination15. These authors suggest that if 90% of staff and students are fully vaccinated, campuses may be able to reopen safely without other measures. However, this 90% cut-off is based on a single preprint modelling study16 that has yet to be peer-reviewed.

The evidence, while sparse, supports strenuous efforts to increase the proportion of staff and students who are fully vaccinated. Most universities are a long way from meeting a 90% target. As of end September 2021, for example, only 58% of 18–24 year olds in England were fully vaccinated12. This is partly because younger age groups were the last to be invited, but also because of relatively high levels of vaccine hesitancy among student age groups, due to a combination of perceived low vulnerability and the “inconvenience” of attending for a jab13. One key measure for improving safety is to make it very easy for people to get a vaccination on campus—for example by locating vaccination hubs close to settings frequented by target groups (e.g. outside lecture theatres or dining halls) and not requiring paperwork that people are unlikely to be carrying.

Vaccination, however, is unlikely to be the sole measure to protect students and staff. Another preprint modelling study, which used the delta variant, found that even at 100% coverage, vaccination was insufficient to eliminate SARS-CoV-2 transmission in a university dormitory setting18. Some vulnerable groups are unable to receive the vaccine or mount an effective immune response to it19. An individual interacting with another group of people in a university setting has no way of knowing what proportion of them are vaccinated. For all these reasons, other measures are likely to be needed for the foreseeable future.

Everyone should wear masks
Masking has two main effects: reducing emission of the virus by the wearer (“source control”), and protecting the wearer from virus emitted by others20–22. It also has a third potential effect—reminding us that we are still in a pandemic and signalling to others that we are taking their safety seriously23.

Reviews of a wide range of evidence (including laboratory studies and natural experiments) have shown that, broadly speaking, masks are effective—but by no means perfect—for source control20–22. Masks reduce the amount of virus that gets into the air, and hence the probability that someone else in the room will be infected22. Wearing a mask reduces viral emissions from coughing and sneezing approximately 20-fold24, but around half of all people who transmit the virus have no symptoms at the time (i.e. they are not coughing or sneezing but simply exhaling the virus in aerosols)25. Different materials for cloth masks have very different filtration properties26; a well-fitting mask with no leaks round the side is crucial27. A double-layer neck gaiter (bandana) and a medical mask both reduce emission of aerosols by around 60%, but respirator (FFP2 or FFP3, N95) masks are much more effective, blocking up to 99% of aerosols26. Note that face visors reduced aerosol emission by only 5%—i.e. they are ineffective28.

There have been claims that randomised controlled trial (RCT) evidence is the only “robust” way to test the impact of masks. This is incorrect, because most such RCTs are designed only to test the hypothesis that the mask protects the wearer over a short period. Actually, masks work mainly by protecting other people, and even a non-statistically significant effect on transmission dynamics (e.g. in lectures) can lead to very large effects over time (for example, if instead of doubling every 9 days, new cases increased by only 1.9-fold, after 180 days cases would be down by 60%).

The above findings support mandating (rather than just encouraging) masking in shared spaces. If everyone is wearing a mask, source control will be high and double-layer cloth masks will be adequate for most healthy people. In one recent Centers for Disease Control and Prevention (CDC) report, US schools without mask mandates in July-August 2021 were 3.5 times more likely to have COVID-19 outbreaks than schools with mandates29. To protect the wearer effectively from airborne virus when others in the room are unmasked, a higher grade of filtration is needed, hence in the absence of near-universal use of source control masks, individuals may be left with little choice but to consider respirators30. Those who are clinically vulnerable (hence requiring masks for self-protection) should use respirators in any case.

Perhaps the most persuasive argument for masks in the university context is that if everyone wears one, there is a much lower risk that teaching will need to return to online as a result of rising case numbers.

Since speaking and singing increase emission of aerosols30,31, masks should be worn continuously indoors and not removed for these activities. The suggestion in some universities that masks should be worn only until people are seated but may be removed thereafter makes no scientific sense. Indeed, because of the airborne nature of the disease, masking is more
important when in a classroom learning setting (indoors, with others, and with some people talking) than when moving between classrooms (especially if walking alone, outdoors and in silence). Likewise, rules in gyms that masks should be worn when walking between equipment but not when exercising on the equipment are nonsensical, since heavy breathing during exercise will increase emission of viral particles.  

The benefit of mask wearing by all is not dependent on the size of a group, so suggestions that masking is needed only above a certain occupancy threshold means that unmasked smaller groups would carry a preventable risk (and also provide a false sense of security).

A major risk setting for transmission of COVID-19 is lunch and tea breaks, since masks must be removed for eating and drinking, and because people often sit at close quarters and talk. To reduce transmission, refreshment breaks should ideally be taken out of doors. If this is not possible, physical distancing should be increased and silence maintained while unmasked. Socialising in breaks could occur, for example, during the walk to the café (while masked) but not while eating.

A few people have a medical reason not to mask (e.g. neurodiverse or anxious); in some universities they may obtain a lanyard to indicate they are exempt.

Space people out (physical distancing, joining remotely, cohorting)  

Physical distancing (sometimes called social distancing) is effective at reducing droplet transmission, since droplets fall to the ground within a few feet due to gravity. Physical distancing also protects against airborne transmission, since most airborne particles are spread via close contact, especially when a person is in the direct stream of someone else’s exhaled breath (think of smelling the garlic on someone’s breath—you might be able to smell it across the room but it is much stronger at close range).

Many university guidelines stipulate a specific physical distance such as 1 or 1.5 metres to space desks apart. Whilst this is a useful rule of thumb, a “safe” distance cannot be calculated precisely, since a) airborne particles spread throughout a room within about 30 minutes (and can remain even after the room has been vacated), hence time spent indoors must also be factored in; b) if nobody is wearing a mask, viral emission is considerably greater (hence, close contact is more risky—and conversely if everyone is masked, it is less risky); c) singing or loud talking increases viral transmission (hence, again, close contact is more risky); d) even wide separation will not protect fully against the turbulent jets emitted when a symptomatic person coughs or sneezes.

Figure 2 summarises this information in a semi-quantitative way; a paper offering a quantified version of this diagram is available as a preprint.
Figure 2. Risk of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) transmission in different settings, assuming people are asymptomatic (adapted from 33 under Creative Commons licence).
class produced an approximately 2-fold decrease in cumulative dose of aerosols. In that study, opening windows during breaks only had minimal effect ($\leq 1.2$-fold decrease). The conclusion from this study is that if it is not possible to open windows more than a crack, a different way of cleaning the air is likely needed.

Whereas mechanical ventilation in domestic settings tends to occur through extractor fans (such as those in kitchens and bathrooms) or ceiling fans (often used as an alternative to air conditioning in hot climates), most mechanical ventilation in universities and colleges is through large systems which take in air through air handling units and supply and extract through a system of ducts and diffuser grilles. Lecture theatres and laboratories are generally mechanically ventilated via centralised HVAC systems. Hence it should not be assumed that if a space has no opening windows, it must be inadequately ventilated.

If indoor spaces are fitted with air conditioning systems, it is important to ensure that air which is removed is not recycled unfiltered (or inadequately filtered) back into that space\(^1\). Air conditioning is not mechanical ventilation, though it may be linked to mechanical ventilation via large, central HVAC systems which both filter and heat (or cool) the air as needed. More problematic are isolated rooms fitted with their own local air conditioning systems, which are less likely to include any filtration and may give a false sense that the room is being ventilated.

The level of ventilation and occupancy in ventilated spaces can be approximated by measuring carbon dioxide (CO\(_2\)) levels, since this is present in higher concentrations in exhaled air than in outdoor air. The higher the CO\(_2\) level in a room, the more exhaled air (and hence, potentially, the more virus) there is. Before the pandemic, indoor air quality standards were generally set around the goal of avoiding “sick building syndrome” (with symptoms such as headaches a sense of stuffiness, due to accumulation of multiple contaminants in the air) and clearing body odours and other smells.

Whilst CO\(_2\) levels can be used to approximate the risk of COVID-19 transmission\(^9\), they are only a proxy for this risk. With that caveat, some authors have suggested that CO\(_2\) levels might be used strategically in negotiations with employers\(^40\).

Figure 3 shows some suggested cut-off levels for denoting “low risk” (below 700 ppm), “medium risk” (700–800 ppm), “high risk” (800–1000 ppm) and “very high risk” (>1000 ppm), though other publications recommend slightly different cut-offs for these categories. Measures to address moderate risk include opening classroom doors and windows, opening windows between classes, and reducing the number of students in the classroom. If levels indicate “high risk” despite these measures, infrastructure changes (such as mechanical or portable air filters) are needed.

Note that the cut-off values for unacceptable CO\(_2\) levels in Figure 3 are substantially lower than those in many official documents (e.g. from UK Health and Safety Executive, who recommend 1500 ppm\(^41\)). This is because the higher cut-off values were set historically for an entirely different purpose.

When it is not possible or desirable to use ventilation (e.g. for energy efficiency reasons) to maintain clean air, other kinds of control are needed. There are two main kinds: an inbuilt mechanical filter (for which standards are expressed as the

### Table: Risk Classification Scheme for Carbon Dioxide (CO\(_2\)) Levels in Indoor Air

<table>
<thead>
<tr>
<th>RISK LEVEL</th>
<th>LOW</th>
<th>MODERATE</th>
<th>HIGH</th>
<th>VERY HIGH</th>
</tr>
</thead>
<tbody>
<tr>
<td>CO(_2) peak concentration or class time average concentration</td>
<td>&lt; 700 ppm</td>
<td>700-800 ppm</td>
<td>800-1000 ppm</td>
<td>&gt; 1000 ppm</td>
</tr>
<tr>
<td>Recommended action</td>
<td>No action needed</td>
<td>If A active, do B If B active, do C If C active, do D</td>
<td>A and B if both active, do C then D If all active, do E</td>
<td>A and C if both active, do D If all active, do E</td>
</tr>
</tbody>
</table>

A: door always open
B: windows open for 10 min at lunch break
C: windows open for 10 min at end of each teaching hour
D: change setting (e.g. windows open longer, reduce number of students per class)
E: infrastructure changes needed (e.g. inbuilt or portable filtration system)

**Figure 3.** Risk classification scheme for carbon dioxide (CO\(_2\)) levels in indoor air (adapted under Creative Commons licence from \(^{40}\)).
minimum efficiency reporting value (MERV) in the USA\textsuperscript{27} or ISO 16890 in Europe) or a portable air cleaner fitted with a HEPA (high-efficiency particulate air) filter. Such filtration systems have been designed to remove particles of many different kinds and sizes (e.g. dust, pollen, smoke, bacteria, viruses).

The SARS-CoV-2 virus is approximately 100 nanometres (0.1 μm) in diameter (though it is unlikely to travel naked so the size of particle to be filtered will be larger than this). The system installed should be efficient in the 0.1 to 1 μm range. Of note to those in charge of supplying clean air to old-fashioned university or school buildings is this warning: “most central mechanical systems were not designed for HEPA filters. Instead, these systems use filters on a different rating scale, minimum efficiency reporting value, or MERV, and typically use a low-grade filter (eg, MERV 8) that captures only approximately 15% of 0.3–1 μm particles, 50% of 1–3 μm particles, and 74% of 3–10 μm particles. For infection control, buildings should upgrade to MERV 13 filters when possible, which could capture approximately 66%, 92%, and 98%, of these sized particles, respectively\textsuperscript{29}.”

Upgrading from MERV8 to MERV13 filters (or the ISO equivalent) is potentially a rapid, affordable and effective intervention for universities and colleges in some settings, but higher-grade filters may induce a bigger pressure drop so unless the fan speeds can be increased, the ventilation rate may be inadequate.

Portable air filtration units fitted with HEPA filters are highly effective at removing aerosols in the 0.1 to 1 μm range\textsuperscript{22–45}. In the Villiers study described above, one HEPA filter was as effective as two windows partly open all day during the winter (2.5-fold decrease in cumulative dose of aerosols) while two HEPA filters were more effective (4-fold decrease)\textsuperscript{39}. A combination of interventions (masks along with natural ventilation \textit{and} HEPA filtration) were the most effective, producing a 30-fold decrease in cumulative aerosol dose\textsuperscript{48}. Aerosol scientists have begun to develop and test home-made, low-cost box fans fitted with HEPA filters as a quick and effective solution for improving mechanical ventilation in poorly-ventilated spaces\textsuperscript{46}.

Ultraviolet (UV) light (from sunlight or radiation lamps) has been shown to destroy SARS-CoV-2 in numerous studies\textsuperscript{47}, though if this method is used it is important to select appropriate units that do not generate ozone. UV sterilization holds potential for enhancing safety in indoor spaces where risk of transmission is particularly high (e.g. hospitals, gyms). In a small before-and-after study published as a preprint, a combination of HEPA filtration \textit{and} UV sterilisation was highly effective at removing bioaerosols (including but not limited to SARS-CoV-2) in a COVID-19 surge ward and intensive care unit in one hospital, though the contribution of the UV component to the result is unknown\textsuperscript{48}.

Electronic air cleaning systems, for example those which use ozone, are of no proven efficacy in reducing COVID-19 transmission\textsuperscript{49}; they currently have no place in preventing transmission of SARS-CoV-2. Air filtration does not remove CO\textsubscript{2}, so CO\textsubscript{2} monitors cannot be used to monitor the quality of filtered air.

In the longer term, universities should consider the need for a paradigm shift in the design and ventilation of buildings, to improve air quality standards and ensure that all indoor spaces meet these through adequate ventilation, filtration or sterilisation\textsuperscript{16}.

Test, trace and isolate while COVID-19 incidence is high

In the context of high incidence of COVID-19 and an unvaccinated or partially-vaccinated student population, frequent testing of asymptomatic staff and students along with contact tracing and support to isolate has been shown to reduce on-campus transmission substantially\textsuperscript{55}. While lateral flow device (LFD) tests can detect asymptomatic cases and break chains of transmission, this measure depends on the efficacy of efforts to track and trace contacts and maintain and support the isolation of infected individuals. Anyone who is symptomatic should isolate immediately and take a gold-standard polymerase chain reaction (PCR) test, irrespective of the status of their LFD test. Universities should ensure clear and consistent communication on this matter as confusion still abounds.

Anyone with symptoms, even if they are perceived to be “just a cold”, should isolate immediately, and a negative LFD should never override the more accurate PCR test (see below). Note that the most common symptoms of delta infection (in order: headache, runny nose, sneezing, sore throat, loss of smell, fatigue) are different from the standard triad of cough, fever and shortness of breath which are still widely used to prompt PCR testing\textsuperscript{56}. Hybrid teaching options greatly facilitate immediate isolation, and students and staff with symptoms that may be due to COVID-19 should be supported to engage remotely if they are well enough to do so. Track and trace efforts are constrained by the specifics of the system. Universities may have additional information that can be harnessed to provide a further layer of safety. In the UK for example, individuals sharing a confined space for extended periods of time, for example, may not be contacted by the official Track and Trace system but could be identified via attendance lists.

Some authors have questioned the validity and expense of mass asymptomatic testing in populations where incidence of COVID-19 is low, due to the very large number of tests required to detect small numbers of positive cases\textsuperscript{52}. A recent modelling study suggests that as vaccination rates rise and the incidence of COVID-19 falls, the cost-benefit balance of frequent testing becomes less favourable\textsuperscript{56}. However, at the
time of writing the UK is a long way from a low-incidence state and we strongly recommend maintaining asymptomatic testing. Below, we explain some of the science behind the tests.

Lateral flow devices (LFDs), which detect the presence of virus antigen in the nose and throat using a swab sample tested in a flow device (like a pregnancy test)\(^1\). Multiple types of LFD test are available, and they are designed to test people (perhaps repeatedly) who are not displaying overt COVID symptoms. LFD tests are all highly specific i.e. they are very unlikely to give a positive result if the person is not infected. But LFD tests are not particularly sensitive (i.e. less able to detect very small quantities of the virus) compared with the gold standard PCR (polymerase chain reaction) tests. This means that testing negative on an LFD is not a “green light” i.e. it does NOT guarantee that the individual is not infected with SARS-CoV-2, so they should continue to practice mitigations as advised. On the other hand, testing positive on an LFD means it is highly likely the person is infected (it is a “red light”, indicating that they are potentially infectious). Such individuals should self-isolate immediately, report the positive test, and order a confirmatory PCR test as soon as possible. A positive LFD test should trigger a call from the Track and Trace service.

Whilst LFD tests are used mainly in people without symptoms, they are actually more likely to be positive if the infected person is symptomatic (probably because such people have higher levels of the virus)\(^2\). However, people with a positive LFD may well be infectious despite lack of symptoms—hence the value of these tests in identifying infectious cases (who should then isolate) and reducing the chance of a super-spreader event on campus. LFD tests also tend to reflect past infection (they are more likely to be positive 2 weeks after the onset of symptoms than on the day symptoms appear)\(^3\).

In sum, the on-site LFD testing established at many UK university sites appears to be evidence-based (though not scientifically perfect) and its regular, frequent use is recommended while the incidence of COVID-19 remains high. Those with symptoms also need a PCR test.

Clinically vulnerable staff and students

Universities and colleges have a duty of care to their staff and students. They must provide a safe environment for learning, teaching, and working. If a person has a condition or risk state which makes them vulnerable to COVID-19 and its complications, the institution must take account of this. Increased vulnerability to COVID-19 occurs in people who are immunosuppressed (including those on medication which suppresses the immune system, and pregnant women), those with certain long-term conditions, older age groups, some minority ethnic groups and those who are overweight. These risk groups were considered in detail in the Independent SAGE report\(^4\).

The evidence supports a policy of vulnerable groups (whether staff or students) being supported to work from home if possible while the incidence of COVID-19 is high. If they must enter indoor spaces they should be advised to wear a respirator mask for self-protection, and it is particularly important for others in the room to wear a mask to maximise source control. If clinically vulnerable people are required to enter indoor spaces, those spaces should be adequately ventilated (confirmed using CO\(_2\) levels) or have high-quality air filtration systems (MERV13 or HEPA) installed.

Interventions for which there is no evidence (“hygiene theatre”)

We found no scientific evidence to support taking temperatures, sanitising hands before entering the classroom (though washing hands when they are dirty and after going to the lavatory is of course a general hygiene measure), restricting the sharing or exchange of fomites (i.e. potentially contaminated objects such as pens, paper, books or other study materials), wearing face visors, or separating desks with plastic screens. Such “hygiene theatre”, which links to a discredited hypothesis that the virus is spread mainly or exclusively by droplets\(^5\), could potentially distract staff and students from measures which do work.

In relation to sanitising, hand hygiene is recognised good practice for the prevention of many infectious diseases, so it should not be dismissed or discouraged (but equally, should not be over-emphasised). In relation to fomite transmission, a large Brazilian study detected no SARS-CoV-2 virus on over 400 samples of mask fronts, cell phones, paper money or card machines during a wave of the pandemic\(^6\). In other words, there is some evidence against the importance of fomite transmission. However, since the mode of transmission remains contested, it would seem sensible to discourage widespread sharing of pencils, books and other objects among students.

Conclusion

The key to effective prevention of COVID-19 is acknowledge- ment of its predominantly airborne mode of transmission. Many widely-promoted measures—hand sanitising, strict 1- or 2-metre distancing, fomite precautions—wrongly assume an exclusively droplet mode of transmission assume and are there- fore ineffective. Such thinking also dominates the thinking of senior management and many staff and students.

Acknowledging the importance of airborne transmission should lead to policies such as: a) masking at all times while indoors, with encouragement to wear higher-grade respirators for best protection (especially if clinically vulnerable); b) continuing attention to physical distancing but in a way that does not assume that a particular interval between desks makes the space “safe”, and using additional measures (joining remotely, cohorting, frequent breaks) to reduce crowding and time spent indoors; c) a greater focus on engineering controls (ventilation and/or filtration of air). In addition, university and
college staff should encourage and facilitate vaccination, attend to testing and tracing, and be ready to instigate tighter controls (e.g. return to online teaching) if case numbers rise.

These measures should be implemented and evaluated. Monitoring of metrics such as CO2 levels and room occupancy rates may provide staff with hard data with which to negotiate with management.

**Data availability**

No data are associated with this article.

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**References**

15. Williams SN, Yamey G: How universities can make reopening safer this autumn. BMJ. 2021; 374: n2019. Published Abstract | Publisher Full Text


Open Peer Review

Current Peer Review Status: 💡 ✔️ ✔️

Version 1

Reviewer Report 03 November 2021

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Michelle Ananda-Rajah
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Excellent and timely review with no major concerns. The following are suggestions that may enhance the manuscript:

- Worth including a statement on mandating vaccination which some universities in the US (Yale I think) and Australia have done. This intervention has greatly increased uptake of vaccination in younger people and staff.

- Please further clarify page 4 para 2 "new cases increased by only 1.9 fold... 60%". It is difficult to understand, please revise.

- To remain silent while unmasked is not pragmatic nor feasible in a University setting. The credibility of the paper could be called into question if the measures proposed are not implementable. Consider making this a suggestion or placing a higher emphasis on HVAC improvements.

- Please elaborate as to why natural ventilation with the windows fully open in spring and summer was less effective accounting for a 2 fold decrease only. What is the likely basis for this result?

- A CO₂ ppm upper limit of 1500 ppm was set for another purpose, please mention this other purpose because it may be a barrier otherwise to the adoption of a lower threshold.

- Re LFD testing in asymptomatic people: how often should this be done and would it vary depending on community transmission rates.

- Consider a box summarising key interventions.

Is the topic of the review discussed comprehensively in the context of the current
Are all factual statements correct and adequately supported by citations?  
Yes

Are the review written in accessible language?  
Yes

Are the conclusions drawn appropriate in the context of the current research literature?  
Yes

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: health services research, artificial intelligence, clinical leadership

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.

Author Response 08 Dec 2021

Trish Greenhalgh, University of Oxford, Oxford, UK

REVIEWER 3

Excellent and timely review with no major concerns. The following are suggestions that may enhance the manuscript:

○ Worth including a statement on mandating vaccination which some universities in the US (Yale I think) and Australia have done. This intervention has greatly increased uptake of vaccination in younger people and staff.

Yes, this is an important point. New text and reference(s) added:  


○ Please further clarify page 4 para 2 “new cases increased by only 1.9 fold... 60%”. It is difficult to understand, please revise. We have changed this text to “(for example, if instead of doubling every 9 days, new cases increased by only 1.9-fold (i.e. almost but not quite doubling), after 180 days cases would be down by 60%)”

○ To remain silent while unmasked is not pragmatic nor feasible in a University setting. The credibility of the paper could be called into question if the measures proposed are not implementable. Consider making this a suggestion or placing a higher emphasis on HVAC improvements. Point taken. We've adapted the paragraph to read as follows:

“A major risk setting for transmission of COVID-19 is lunch and tea breaks, since masks must be removed for eating and drinking, and because people often sit at close quarters and talk. To reduce transmission, refreshment breaks should ideally be taken out of doors. If this is not possible, physical distancing should be increased and silence maintained while unavoidably unmasked (i.e. when eating or drinking). Chatting during breaks could occur, for example, during the walk to the café (while masked) but not while eating - though this is likely to prove unimplementable in most campus settings.”

○ Please elaborate as to why natural ventilation with the windows fully open in spring and summer was less effective accounting for a 2 fold decrease only. What is the likely basis for this result?

Added:

), because the effectiveness of natural ventilation depends on the difference between indoor and outdoor temperature (which is less in spring and summer 38 )

○ A CO2 ppm upper limit of 1500 ppm was set for another purpose, please mention this other purpose because it may be a barrier otherwise to the adoption of a lower threshold.

We have softened this recommendation since SAGE have recommended 1500 as the cutoff for Covid controls and we don't want to lock horns over it. We've now made clear that the levels recommended by Di Gilio et al are a counsel of perfection but the priority is to address settings where levels are above 1500.

○ Re LFD testing in asymptomatic people: how often should this be done and would it vary depending on community transmission rates.

We have acknowledged that there is uncertainty about this and said that optimum interval will depend on transmission rates and size of population. We have also given a recommended interval of twice weekly from a recent BMJ editorial and said this appears based on convention and an attempt to balance costs/hassle (and possible
This is an excellent and comprehensive review of the evidence relevant to improve the safety of return of students to universities and schools in the context of the continuing transmission of SARS-CoV-2. The review should be augmented to consider more explicitly a number of issues:

**Vaccination:**
- The review is written specifically for a United Kingdom audience, and it is perhaps for that reason that it does not mention universities or schools requiring vaccination of students or staff. In Australia, for example, in some states, government regulations effectively require that only vaccinated people are able to return to campus. A number of universities have also introduced mandatory vaccination of the staff and students who are to be on campus. This obviously requires consultation with industrial organisations, but some reference to this issue would be useful to ensure a comprehensive review of options.

**Physical distancing:**
- The review appropriately refers to the droplet induced theories about physical distancing. This section should be broadened to include stronger consideration of issues relating to remote and hybrid working, cohorting, and workplace bubbles. There may be limited published evidence on these strategies, and for this reason they may have been excluded, but these are strategies that universities and schools are considering and have a logical basis.
- Separation of cohorts is an effective way of reducing the likelihood of spread from one group to another. For example, if a university or school has multiple offerings of the same
subject, it may introduce rules to require students not to swap between offerings in a term/semester. Similarly, it might encourage staff work on campus only on designated days to minimise overlap between staff cohorts.

- Risk of transmission is maximised when people are congregating in small areas such as waiting to use lifts/elevators. In administrative workplaces, it may be useful to consider encouraging staggered start times to minimise congregation in lift/elevator waiting areas. Similarly, consideration should be given to dispersal strategies for students leaving large lecture theatres, including encouraging use of stairs. The emphasis on masks is appropriate but could be augmented by other strategies.

**Testing:**

- The paper discusses the importance of rapid testing but does not discuss how to encourage people to respond if they test positive. This includes for staff, ensuring that they are paid while isolating. Special provisions may be required for casual and temporary contract staff.

- Some staff will be able to continue to work remotely, and that should be encouraged, and arrangements should already be in place for that to occur.

- The review does not discuss the frequency of testing — whether people should be tested every day before attending school/university, every second day, etc. There may be evidence on optimal testing frequency that should be included. On a related matter, there is no mention of managing the risk of staff who are employed in multiple organisations because of casualisation of the workplace. Multiple employment increases the risk of transmission from one workplace to another and universities and schools may wish to attempt to limit the number of separate workplaces that casual staff physically attend.

**Is the topic of the review discussed comprehensively in the context of the current literature?**

Yes

**Are all factual statements correct and adequately supported by citations?**

Yes

**Is the review written in accessible language?**

Yes

**Are the conclusions drawn appropriate in the context of the current research literature?**

Yes

**Competing Interests:** No competing interests were disclosed.

**Reviewer Expertise:** Health policy, including policies about SARS-CoV-2

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.
This is an excellent and comprehensive review of the evidence relevant to improve the safety of return of students to universities and schools in the context of the continuing transmission of SARS-CoV-2. The review should be augmented to consider more explicitly a number of issues:

**Vaccination:**
- The review is written specifically for a United Kingdom audience, and it is perhaps for that reason that it does not mention universities or schools requiring vaccination of students or staff. In Australia, for example, in some states, government regulations effectively require that only vaccinated people are able to return to campus. A number of universities have also introduced mandatory vaccination of the staff and students who are to be on campus. This obviously requires consultation with industrial organisations, but some reference to this issue would be useful to ensure a comprehensive review of options.

Yes, this is an important point. New text and reference(s) added:


**Physical distancing:**
- The review appropriately refers to the droplet induced theories about physical distancing. This section should be broadened to include stronger consideration of issues relating to remote and hybrid working, cohorting, and workplace bubbles. There may be limited published evidence on these strategies, and for this reason they may have been excluded, but these are strategies that universities and schools are considering and have a logical basis.

Agree, we have added some text to address all these points. In relation to ‘bubbles’, we've included a new reference (Thompson DA, Abbasizanjani H, Fry R, et al Staff–pupil SARS-CoV-2 infection pathways in schools in Wales: a population-level linked data approach. BMJ Paediatrics Open 2021;5:e001049. doi: 10.1136/bmjpo-2021-001049) which provides statistically significant evidence that year-level bubbles have a positive impact, since cases are correlated within year cohorts but not between years.
Separation of cohorts is an effective way of reducing the likelihood of spread from one group to another. For example, if a university or school has multiple offerings of the same subject, it may introduce rules to require students not to swap between offerings in a term/semester. Similarly, it might encourage staff work on campus only on designated days to minimise overlap between staff cohorts.

Added - thanks. We’ve emphasised the important contribution of hybrid learning (with some people learning remotely).
  ○ Risk of transmission is maximised when people are congregating in small areas such as waiting to use lifts/elevators. In administrate workplaces. it may be useful to consider encouraging staggered start times to minimise congregation in lift/elevator waiting areas. Similarly, consideration should be given to dispersal strategies for students leaving large lecture theatres, including encouraging use of stairs. The emphasis on masks is appropriate but could be augmented by other strategies.

Added - thanks

Testing:
  ○ The paper discusses the importance of rapid testing but does not discuss how to encourage people to respond if they test positive. This includes for staff, ensuring that they are paid while isolating. Special provisions may be required for casual and temporary contract staff.

Thanks - we’ve added some text on this.
  ○ Some staff will be able to continue to work remotely, and that should be encouraged, and arrangements should already be in place for that to occur.

We completely agree. Staff should be enabled to work from home wherever possible to do so.
  ○ The review does not discuss the frequency of testing — whether people should be tested every day before attending school/university, every second day, etc. There may be evidence on optimal testing frequency that should be included. On a related matter, there is no mention of managing the risk of staff who are employed in multiple organisations because of casualisation of the workplace. Multiple employment increases the risk of transmission from one workplace to another and universities and schools may wish to attempt to limit the number of separate workplaces that casual staff physically attend.

We’ve added some comments about this. The only published recommendations we could find are based on convention (twice weekly). We’ve added that this interval appears to seek to balance cost, the potential reduction in compliance if the interval is very short, and the need to pick up new cases quickly.

**Competing Interests:** no coi
This review aims to provide a timely synthesis of the literature relevant to SARS-CoV-2 transmission in an educational setting and of the available mitigation strategies that might be employed to facilitate a return of in-person university teaching.

This is an ambitious piece and will be of value to those involved in delivering higher education during the pandemic. Helpfully, the authors critically evaluate a wide variety of potential mitigation strategies, highlighting those supported by published evidence, while importantly identifying those for which the evidence does not support continuation. In the former “effective” category, the authors include vaccination, mask wearing, physical distancing, engineering solutions (room ventilation and air filtration), test/trace/isolation, and blended learning. Approaches that the authors conclude to be “ineffective” include strict hand hygiene, restricting spread via fomites, the wearing of face shields, and the separation of individuals by plastic screens.

The article is clear, well-written, and makes useful evidence-based recommendations. The authors may wish to consider the following critical comments, which might help to improve the article further. In particular, in their effort to provide a useful synthesis of the evidence, at times the distinction between published literature and their interpretation becomes unclear. It might be helpful, and would strengthen the review, if this distinction were to be made more obvious.

Methods section:
- The methods section describes a search of PubMed in September 2021 for the terms “SARS-CoV-2”, “COVID-19”, “transmission”, “mitigation”, “school[s]” and “university/ies”, restricted initially to reviews. These were examined for relevant papers by “snowball searching”. It would be helpful to know the total number of articles assessed and to be provided with more detail about the processes and metrics used to evaluate their quality. Would a PRISMA diagram be valuable in this regard? The authors refer to the screening of articles, in part, by considering the impact factor of the journals in which they were published. While impact factors have some value as a heuristic for quality, this approach also has limitations. It would be reassuring to learn how the threshold for impact factor was set and whether this was corrected for journal type, for example, clinical vs non-clinical journals. It would also be valuable to provide more detail on the methodologies employed to avoid the introduction of any bias. This might be provided in the Methods section or in a dedicated “limitations” section.

Main findings section:
- General comments:
  - This section provides a narrative synthesis of the above search. It is clearly written and generally well-referenced. In places, however, the referencing is less extensive and it is not
always clear if this reflects a deficiency in the published literature. More clarity on this would be helpful in allowing readers to evaluate the validity of the recommendations. It might also be helpful to provide a quantification of the strength of the recommendations, perhaps along the lines of the GRADE (Grades of Recommendation Assessment, Development and Evaluation) approach used in clinical guidelines.¹

Specific comments:

- It is stated that “.. this 90% cut-off is based on a single preprint modelling study [16] that has yet to be peer-reviewed.” Would the authors double check that this statement is true? The article appears to have been published on 31 August 2021 in *Annals of Internal Medicine*, although at the time of writing (24/10/2021), its status has not been updated on medRxiv.

- It is stated, “One key measure for improving safety is to make it very easy for people to get a vaccination on campus—for example by locating vaccination hubs close to settings frequented by target groups (e.g. outside lecture theatres or dining halls) and not requiring paperwork that people are unlikely to be carrying.” This is an important point, but is this an opinion of the authors or are there empirical studies to support it? Is it possible to make clearer the distinction between available data, knowledge gaps, and opinion?

- It is stated, “There have been claims that randomised controlled trial (RCT) evidence is the only “robust” way to test the impact of masks. This is incorrect, because most such RCTs are designed only to test the hypothesis that the mask protects the wearer over a short period.” Again, this is very important. Is it possible to provide citations for the relevant articles? Similarly, elsewhere in this section, comments are made such as “Perhaps the most persuasive argument for masks in the university context is that if everyone wears one, there is a much lower risk that teaching will need to return to online as a result of rising case numbers”. These comments are important and would be strengthened if published evidence could be cited. However, if this is interpretation, would it be valuable to separate the literature review from the interpretation?

- It is stated, “The benefit of mask wearing by all is not dependent on the size of a group, so suggestions that masking is needed only above a certain occupancy threshold means that unmasked smaller groups would carry a preventable risk (and also provide a false sense of security).” Are there published studies available to support this?

- Figure 2 encodes “risk of transmission” graphically by using a red/amber/green colour scheme. Would it be possible to amend the figure to be more accessible for those with deficiencies in colour vision? I suspect that in its current form it would be challenging for readers with deuteranopia or protanopia.

- In the section on testing, the authors might wish to consider a discussion of mass asymptomatic PCR testing of undergraduates, which has been shown both to be possible and effective in the university setting, for example, through the testing of pooled screening-samples from groups of 10 to 12 students followed by individual testing of positive groups. This has recently been reviewed.²

- I do not feel able to comment usefully on the engineering solutions section, which falls outside my areas of knowledge.
References

Is the topic of the review discussed comprehensively in the context of the current literature?
Yes

Are all factual statements correct and adequately supported by citations?
Partly

Is the review written in accessible language?
Yes

Are the conclusions drawn appropriate in the context of the current research literature?
Yes

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: Respiratory medicine, including COVID-19; cell biology, ER stress

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Author Response 08 Dec 2021

Trish Greenhalgh, University of Oxford, Oxford, UK

Reviewer Report 1
25 Oct 2021 | for Version 1
Stefan J. Marciniak, Cambridge Institute for Medical Research, University of Cambridge, Cambridge, UK

APPROVED WITH RESERVATIONS
This review aims to provide a timely synthesis of the literature relevant to SARS-CoV-2 transmission in an educational setting and of the available mitigation strategies that might be employed to facilitate a return of in-person university teaching.

This is an ambitious piece and will be of value to those involved in delivering higher education during the pandemic. Helpfully, the authors critically evaluate a wide variety of potential mitigation strategies, highlighting those supported by published evidence, while
importantly identifying those for which the evidence does not support continuation. In the former “effective” category, the authors include vaccination, mask wearing, physical distancing, engineering solutions (room ventilation and air filtration), test/trace/isolation, and blended learning. Approaches that the authors conclude to be “ineffective” include strict hand hygiene, restricting spread via fomites, the wearing of face shields, and the separation of individuals by plastic screens.

The article is clear, well-written, and makes useful evidence-based recommendations. The authors may wish to consider the following critical comments, which might help to improve the article further. In particular, in their effort to provide a useful synthesis of the evidence, at times the distinction between published literature and their interpretation becomes unclear. It might be helpful, and would strengthen the review, if this distinction were to be made more obvious.

Methods section:
- The methods section describes a search of PubMed in September 2021 for the terms “SARS-CoV-2”, “COVID-19”, “transmission”, “mitigation”, “school[s]” and “university/ies”, restricted initially to reviews. These were examined for relevant papers by “snowball searching”. It would be helpful to know the total number of articles assessed and to be provided with more detail about the processes and metrics used to evaluate their quality. Would a PRISMA diagram be valuable in this regard? The authors refer to the screening of articles, in part, by considering the impact factor of the journals in which they were published. While impact factors have some value as a heuristic for quality, this approach also has limitations. It would be reassuring to learn how the threshold for impact factor was set and whether this was corrected for journal type, for example, clinical vs non-clinical journals. It would also be valuable to provide more detail on the methodologies employed to avoid the introduction of any bias. This might be provided in the Methods section or in a dedicated “limitations” section.
- [response]

We agree the methods section was too short and didn't sufficiently describe the search strategy or its rationale. We've now included a longer methods section and also a new paragraph at the start of the findings section describing the dataset. Note however that we deliberately did not use a 'technocratic' approach to searching and critical appraisal for reasons set out in the revised paper. In short, we weren't resourced to do so and the sheer breadth of the review (along with the trade-off against time pressures) precluded anything approaching a Cochrane-style approach or formal application of GRADE criteria. There is also the argument that GRADE was designed for clinical epidemiology type studies and is less useful for assessing studies from the engineering literature for example. We've flagged that this is a preliminary review and that the literature would benefit from others undertaking more focused reviews with formal critical appraisal of studies in all the topic areas covered.

We have removed the phrase “impact factor of the journal”. Agree it’s a red herring, and whilst MOST seminal sources were published in high-impact factor journals, that’s not what makes them seminal. What makes them seminal is the fact that others in the same sub-field judge them to be authoritative. We've amended the text accordingly. The use of seminal sources is discussed in more detail in the Greenhalgh and Peacock 2004 paper we
Main findings section:
General comments:

- This section provides a narrative synthesis of the above search. It is clearly written and generally well-referenced. In places, however, the referencing is less extensive and it is not always clear if this reflects a deficiency in the published literature. More clarity on this would be helpful in allowing readers to evaluate the validity of the recommendations. It might also be helpful to provide a quantification of the strength of the recommendations, perhaps along the lines of the GRADE (Grades of Recommendation Assessment, Development and Evaluation) approach used in clinical guidelines.

See above. We're concerned that GRADE isn't the right approach here. The lead author has just written the following paragraph for a different paper: “Evidence hierarchies such as GRADE (Grading of Recommendations, Assessment, Development and Evaluations) have a respectable provenance within clinical epidemiology (Guyatt et al., 2008) but in recent years they have come to be used excessively, naively and in increasingly politicized ways. Such hierarchies are used by journal editors, reviewers and grant-giving panels—usually in well-intentioned efforts to maintain scientific quality—as a technocratic substitute for deeper engagement with quality of scholarship. The effect of such hierarchies is to vastly restrict the range and diversity of evidence that is generated, published, acknowledged as good research, and fed into policy deliberations.”

We do agree that ideally, this entire study would be repeated with each section subjected to a more exhaustive literature search AND a formal assessment of the evidence base using quality criteria appropriate to the sub-discipline (e.g. modelling studies assessed according to what good looks like in modelling science etc). However, the present paper is not that more exhaustive output (which would take an order of magnitude more hours). We’ve amended the paper to make it very clear that we agree this piece of work needs doing and we invite others to contribute to the collective effort.

Specific comments:

- It is stated that “...this 90% cut-off is based on a single preprint modelling study [16] that has yet to be peer-reviewed.” Would the authors double check that this statement is true? The article appears to have been published on 31 August 2021 in Annals of Internal Medicine, although at the time of writing (24/10/2021), its status has not been updated on medRxiv.


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a vaccination on campus—for example by locating vaccination hubs close to settings frequented by target groups (e.g. outside lecture theatres or dining halls) and not requiring paperwork that people are unlikely to be carrying.” This is an important point, but is this an opinion of the authors or are there empirical studies to support it? Is it possible to make clearer the distinction between available data, knowledge gaps, and opinion?

We agree this was a comment rather than an empirical finding and have now flagged it as such.

- It is stated, “There have been claims that randomised controlled trial (RCT) evidence is the only “robust” way to test the impact of masks. This is incorrect, because most such RCTs are designed only to test the hypothesis that the mask protects the wearer over a short period.” Again, this is very important. Is it possible to provide citations for the relevant articles?


- Similarly, elsewhere in this section, comments are made such as “Perhaps the most persuasive argument for masks in the university context is that if everyone wears one, there is a much lower risk that teaching will need to return to online as a result of rising case numbers”. These comments are important and would be strengthened if published evidence could be cited. However, if this is interpretation, would it be valuable to separate the literature review from the interpretation?

  We understand that this request reflects how conventional medical reviews are presented. However, this review is constructed with a view to being a very quick reference guide (it's likely to be skim-read by busy academics and administrators). In the social sciences, it is standard practice for the ‘findings’ from a review to be presented alongside an interpretation of those findings. Indeed, one can think of a continuum with ‘pure findings’ at one end and ‘pure interpretation’ at the other end but quite a bit in between. If this were a RCT with an effect size and a confidence interval that directly addressed the research question, we'd be operating at the poles of the continuum--with a clear ‘results’ section and then a ‘discussion’ section. But what we have here is a bunch of marginally-relevant studies with findings that don't stand independently and it's impossible to present them without also adding a rider about how relevant or useful they are. This means that SOME of the interpretation is inevitably going to be in the findings section. We don't want to double-handle the studies so would prefer to keep the interpretation where it is.

- It is stated, “The benefit of mask wearing by all is not dependent on the size of a group, so suggestions that masking is needed only above a certain occupancy threshold means that unmasked smaller groups would carry a preventable risk (and also provide a false sense of security).” Are there published studies available to support this?

  We agree this was a misleading sentence. We've changed it to “whilst the odds of becoming infected increases with the number of people in any indoor space, there is no occupancy threshold below which those odds reach zero. Since asymptomatic
transmission is common in this disease, groups of any size should wear masks to maximise protection of everyone.”

- Figure 2 encodes “risk of transmission“ graphically by using a red/amber/green colour scheme. Would it be possible to amend the figure to be more accessible for those with deficiencies in colour vision? I suspect that in its current form it would be challenging for readers with deuteranopia or protanopia.

This is an important point (one of the authors has colourblindness and another is a carrier of the condition and has brothers and a son with it). We have replaced Fig 2 with a colourblind friendly version for which we thank the charity Colourblind Awareness for the revised graphic which has been tested on three commonest forms of colourblindness (deut, trit and prot).

- In the section on testing, the authors might wish to consider a discussion of mass asymptomatic PCR testing of undergraduates, which has been shown both to be possible and effective in the university setting, for example, thought the testing of pooled screening-samples from groups of 10 to 12 students followed by individual testing of positive groups. This has recently been reviewed.2

We have amended the text to acknowledge that this is a potential option (and included the helpful reference suggested by the reviewer). However, there are also some counter-arguments which we have set out in the revised paper. We summarised those below:

We agree that PCR could represent a gold-standard means of conducting mass testing, and sample pooling is an eminently sensible strategy for the University setting. However, there may also be disadvantages compared with the use of lateral flow devices (LFDs) and a cost/benefit analysis may be pertinent.
First, the costs and work necessary to establish testing facilities on a large scale will be substantially increased in terms of required equipment, reagents and laboratory safety; sample processing for CL3 materials will require additional staff training, appropriate PPE and microbial safety cabinets. This compares to a self-administered LFD, where both swabbing and disposal are conducted primarily at the individual level in adults.
Second, the speed with which results are delivered by PCR will clearly be slower than LFD, and the information will require a form of confidential electronic dissemination; presumably relevant apps exist, but will further increase cost. This could potentially mean that infected individuals attend classes or other interactions during the interim (unless symptomatic), increasing transmission risk. Furthermore, pooled individuals will require re-testing and, depending upon the timescale in which this is achieved, could mean that peak virus load has passed, especially for vaccinated individuals in which the kinetics of declining virus titre in the upper respiratory tract are more rapid. Whilst LFDs lack the fine sensitivity of PCR, they retain impressive specificity and are particularly suited to detecting the higher end of virus loads present within more infectious individuals.
Lastly, convenience and immediacy are likely to appeal to students and staff alike as, whilst...
LFDs ought not to be used as a green light test, regular asymptomatic testing amongst fixed populations is likely to detect the majority of cases and, combined with other mitigations, limit transmission. However, the value of asymptomatic LFD tests is likely to be reduced where SARS-CoV2 prevalence is low amongst populations.

- I do not feel able to comment usefully on the engineering solutions section, which falls outside my areas of knowledge.
- Is the topic of the review discussed comprehensively in the context of the current literature?
  - Yes
- Are all factual statements correct and adequately supported by citations?
  - Partly
- Is the review written in accessible language?
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- Are the conclusions drawn appropriate in the context of the current research literature?
  - Yes

References

Competing Interests
*No competing interests were disclosed.*

Reviewer Expertise
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*Competing Interests:* none