PASSING THE TEST: A MODEL-BASED ANALYSIS OF SAFE SCHOOL-REOPENING STRATEGIES

Alyssa Bilinski MS 1 · Joshua A. Salomon, PhD 2 · John Giardina, MA 1 , 3 Andrea Ciaranello MD, MPH 4 · Meagan C. Fitzpatrick, PhD 56

Abstract

Background. The COVID-19 pandemic has induced historic educational disruptions. In April 2021, about 40% of US public school students were not offered full-time in-person education.

Objective. To assess the risk of SARS-COV-2 transmission in schools.

Design. We developed an agent-based network model to simulate transmission in elementary and high school communities, including home, school, and inter-household interactions.

Setting. We parameterized school structure based on average US classrooms, with elementary schools of 638 students and high schools of 1,451 students. We varied daily local incidence from 1 to 100 cases per 100,000 population.

Patients (or Participants). Students, faculty/staff, and adult household members.

Interventions. Isolation of symptomatic individuals, quarantine of an infected individual's contacts, reduced class sizes, alternative schedules, staff vaccination, and weekly asymptomatic screening.

Measurements. We projected transmission among students, staff and families following a single infection in school and over an 8-week quarter, contingent on local incidence.

Results. School transmission varies according to student age and local incidence and is substantially reduced with mitigation measures. Nevertheless, when transmission occurs, it may be difficult to detect without regular testing due to the subclinical nature of most children's infections. Teacher vaccination can reduce transmission to staff, while asymptomatic screening improves understanding of local circumstances and reduces transmission.

Limitations. There is uncertainty about susceptibility and infectiousness of children and low precision regarding the effectiveness of specific countermeasures, particularly with new variants.

Conclusion. With controlled community transmission and moderate mitigation, elementary schools can open safety, while high schools require more intensive mitigation. Asymptomatic screening can facilitate reopening at higher local incidence while minimizing transmission risk.

Funding sources. The authors were supported by the Centers for Disease Control and Prevention though the Council of State and Territorial Epidemiologists (NU38OT000297-02; AB, JAS), the National Institute of Allergy and Infectious Diseases (R37AI058736-16S1; AC and K01AI141576; MCF), the National Institute on Drug Abuse (3R37DA01561217S1; JAS), and Facebook (unrestricted gift; JG, AB, JAS). The papers' contents are solely the responsibility of the authors and do not represent the official views of the funders.

¹Interfaculty Initiative in Health Policy, Harvard Graduate School of Arts and Sciences (abilinski@g.harvard.edu)

²Center for Health Policy & Center for Primary Care and Outcomes Research, Stanford University School of Medicine

³Center for Health Decision Science, Harvard T.H. Chan School of Public Health

⁴Medical Practice Evaluation Center and Division of Infectious Disease, Massachusetts General Hospital, Harvard Medical School

⁵Center for Vaccine Development and Global Health, University of Maryland School of Medicine (meagan.fitzpatrick@som.umaryland.edu)

⁶Word Count: 4331

Introduction

During the spring 2020 COVID-19 outbreak, all 50 states recommended or mandated public school closures, impacting at least 124,000 US schools and 55.1 million students (1). Reopenings have proven challenging, and as of April 2021, only about 60% of K-12 students were offered full-time in-person learning (2). While this percentage has nearly doubled since December 2020, where in-person education is offered, many families continue to opt out, including 43% in Boston Public Schools and 65% in New York City Public Schools (3,4). Nevertheless, many parents and advocates have objected strongly to school closures (5), and the Biden administration has stated a priority to facilitate safe, in-person school reopening (6).

Debates around school reopening have been heated and often invoked seemingly contradictory evidence about safety. For example, a number of well-studied cases in school settings have found minimal secondary transmission (7–9). Nevertheless, school clusters have also been documented, particularly in Israel and in parts of the United States (10–12), and some observational studies suggested that school closures may have substantially reduced transmission (13,14). Proponents of reopening schools point to evidence that COVID-19 is most often mild in children (15), while others have expressed concern about transmission to staff, families and the community. The discord plays out similarly on a macro scale: a number of Asian and European countries reopened schools with physical distancing when community transmission was low and reported negligible increases in transmission (16,17). Some, including France, the United Kingdom, and Ireland, kept schools open during the fall wave and reversed surging transmission by closing other sectors, although additional closures have occurred following the emergence of variant B.1.1.7. (18,19). Others, such as Austria, the Czech Republic, and South Korea, closed schools to address rising case burdens (18).

Nevertheless, there is little debate that the benefits of in-person education are substantial, particularly amidst reports of high levels of remote absenteeism, increased depression, anxiety, and suicidality, and parent concerns around educational quality (20–22). Beyond educational and mental health outcomes, opening schools also improves access to social services for children and labor market outcomes for working parents, especially women (23–27). In order to reopen safely, it is critical to take a comprehensive account of the full array of evidence and identify procedures to minimize risks.

We simulated SARS-CoV-2 transmission dynamics in elementary and high schools, characterizing how school transmission may occur either as isolated or sustained outbreaks. While several recent papers have discussed the impact of school reopening on local transmission under different mitigation strategies (28–30), our work focused on the risk that transmission will occur on a school campus and spread to household

members, a less-explored focus (31,32). We emphasize uncertainty in transmission risk (rather than focusing on the average) as well as the observability of infections to school and public health staff, reconciling apparent contradictions in evidence. We evaluated outcomes under varying combinations of local incidence, in-classroom mitigation efforts, testing practices, and staff vaccination.

METHODS

ANALYTIC OVERVIEW

We developed an age-specific, agent-based network model of COVID-19 transmission in elementary and high school communities (Figure 1). We incorporated interactions within schools and homes, as well as those between households outside of school. Because data are inconclusive on transmission among middle school-aged children, we did not model middle schools explicitly; accumulating evidence suggests that they may be more similar to high schoolers than elementary schoolers with respect to susceptibility and infectiousness (33,34).

STRATEGIES

We simulated different combinations of interventions, incorporating strategies that target the three axes of general infection control in schools; COVID-19-specific countermeasures; and scheduling/cohorting. Parameters for the model and strategies are detailed in Table 1.

General infection control in schools:

- 1. **Low uptake:** Schools implemented no or minimal general infection control measures, such as masking and distancing.
- 2. **Medium uptake:** Schools implemented masking and distancing, and adherence was moderate such that the risk of transmission given infectious contact was $\frac{2}{3}$ that experienced in a school with low uptake.
- 3. **High uptake:** Schools implemented masking and distancing and adherence was high, such that the risk of transmission given infectious contact was $\frac{1}{3}$ that experienced in a school with low uptake.

COVID-19-specific countermeasures:

1. Symptom-based self-isolation ("Symptomatic isolation"): Individuals in the school were screened for symptoms daily, and those who developed clinically-recognizable symptoms did not attend school. In setting guidance for symptom-based isolation, schools balance between expansive symptom definitions that lead to many unnecessary missed days and more restrictive definitions that may miss subtle presentations of COVID-19 (35,36). Our primary source for this parameter defined symptoms triggering isolation as fever, respiratory illness (e.g., cough, shortness of breath),

Table 1: Parameter values.

Parameter	Value	Source
Number of students in	638	(37)
elementary school		
Average of students per	21.3	Consistent with (37), 5 classes per grade
elementary school class		
Number of adults who are not	30	Estimate via school websites & personal communication
primary teachers in		
elementary schools		
Number of students in high	1451	(37)
school		
Average of students per high	23	Consistent with (37), 16 classes per grade
school class		
Number of adults who are not	30	Estimate via school websites & personal communication
primary teachers in		
elementary schools		
Probability of asymptomatic	0.2	(38)
disease for adults		
Probability of asymptomatic	0.4	(8,10)
disease for children		
Relative infectiousness of	0.5	(38)
asymptomatic disease		
Probability of subclinical	0.4	Set at upper bound of estimates of asymptomatic disease
disease for adults (includes		(38)
asymptomatic)		
Probability of subclinical	0.8	(39)
disease for children (includes		
asymptomatic)		
Household attack rate	20% (~4%	(40,41)
	per day)	

Parameter	Value	Source
Classroom attack rate	1/2/3%	
	per day	
Relative attack rate for	0.13	Approximately 45 minutes
random school contacts		
(vs. classroom)		
Relative attack rate for	2	Assuming staff congregate (e.g. in lunchrooms)
random staff contacts		
(vs. classroom)		
Relative attack rate for	2	Assuming fewer precautions
childcare contacts		
(vs. classroom)		
Distribution on viral load	Lognormal	Produces 20%/45% from differences in viral load (42,43)
	(.84,	
	.3)/.84	
Incubation period (from	Gamma	Estimated from (44,45)
exposure to symptoms, if	(5.8, 0.95)	
symptomatic)		
Start of infectiousness relative	Normal	Estimated as mean 1.3 days, (46) mean 1.2 days w/lower
to symptoms	(1.2, 0.4)	bound of 95% CI of 3 for start and 2 for peak (47)
Duration of infectiousness	Lognormal	(42,45,48,49)
	(5, 2)	
Duration of isolation and	10 days	(34)
quarantine		
Teacher vaccination uptake	75%	(1,50)
Vaccination effectiveness	90%	(51), assuming that vaccine blocks infections as well as
		prevents symptomatic illness (currently unknown)
Test uptake	90%	Assumed
Test sensitivity during	90%	(52–56)
infectious period		

gastrointestinal symptoms, and/or new loss of smell or taste, affecting about 20% of child and adolescent cases (39).

- 2. Diagnostic testing plus classroom notification ("Classroom quarantine"): Individuals who developed symptoms were isolated and immediately tested. The school received results within a day, and all classroom(s) associated with an individual who tested positive were notified and closed for 10 days, per Centers for Disease Control and Prevention guidelines (34).
- 3. **Teacher vaccination with an infection-blocking vaccine ("Staff vaccination"):** In addition to classroom quarantine, teacher susceptibility was reduced to 33% of baseline, roughly assuming that 75% of teachers receive an infection-blocking vaccine with 90% effectiveness (50,51,57).
- 4. Weekly asymptomatic screening ("Weekly screening"): In addition to classroom quarantine, asymptomatic students and/or staff at each school were tested weekly to reduce asymptomatic and presymptomatic transmission. We assumed that schools used a PCR test from either saliva or a swab of the anterior nares, and results were available within 24 hours. We further specified test sensitivity of 90% and 90% uptake, reflecting PCR test sensitivity, although some studies also suggest antigen tests may reach 90% sensitivity during the infectious period (with a faster turnaround time) (52–56). Upon receipt of a positive result, infected individuals isolated outside of school for 10 days based on CDC guidelines, and siblings and classroom members of an individual who tests positive were notified and quarantined for 10 days.

Scheduling/Cohorting:

- 1. **5-day schedule:** This scenario simulated a traditional 5-day, in-person learning schedule, allowing for classroom contacts, staff-staff interactions (10 per day), and random contacts between school members (20 per day). Related arts and special education teachers had revolving contact with 5 classrooms per day.
- 2. **Cohorting:** This scenario again assumed 5 days of in-person learning for all students, but with restricted out-of-classroom contacts, including separation of classes for lunch and recess. Students and primary teachers continued to have sustained classroom contacts. We assumed a 50% decrease in the number out-of-classroom contacts during the school day and that related arts were taught remotely.
- 3. **Half class sizes:** All students attended school five days per week, but in classes of half their typical size. To accommodate this, the number of teachers was doubled. Limited contacts as in (2) were also maintained.
- 4. Hybrid scheduling (A/B): Classes were subdivided into 2 cohorts, and students attended school for

2 days per week, either Monday/Tuesday or Thursday/Friday. Elementary school children in the same household attending the same school were sorted into the same cohort. All staff were physically present each instructional day. In sensitivity analysis, we considered alternative hybrid schedules. Limited contacts as in (2) were also maintained.

MODEL STRUCTURE

Elementary schools

We generated a set of synthetic households from US population data (58), including students, staff, and adult household members (Supplement). We simulated an average-sized elementary school of 638 pupils from 500 households, of which 114 had more than one child in the school. We set five classes per grade level, an average class size of 21 children, and one teacher. We also incorporated 30 additional staff, to reflect roles such as administrators, counselors, cafeteria staff, custodians, special education teachers, and teachers of related arts (or 'specials', e.g. music and art). These were assigned either rotating in-classroom roles (n = 15) or out-of-classroom roles (n = 15).

High schools

We simulated an average-sized high school of 1451 students in 1225 households, with 142 households having more than one student in the school (37). Students rotated among 8 class periods per day (23 students and 1 teacher), with the distribution of students to class periods chosen randomly within grade levels but repeated daily for the full simulation. The high school had 45 additional staff with out-of-classroom roles and 15 with in-classroom roles. In both elementary and high schools, teachers were modeled to have additional staff contacts per day (e.g., contacts in break rooms or offices).

Out-of-school interactions

Reflecting social interactions and out-of-school childcare, our base case assumed that each family interacted with 1 additional family on each day they did not attend school, randomly reassorted each day. We varied this number in sensitivity analysis from 0 to 9 families per day out of school. We assume that when families mix, no more than 2 adults attend at a given time, who are randomly selected.

TRANSMISSION

At each daily time-step, we modeled dyadic interactions between individuals according to household, classroom, school, and childcare relationships, drawing parameter values from the distributions specified in Table 1. A SARS-CoV-2-infected individual transmitted to susceptible individuals according to contact type (e.g. home, school), infectiousness and susceptibility (adult vs. child), symptom status, and an individual dispersion factor.

Secondary attack rate

Secondary attack rates for SARS-CoV-2, the probability that a person with SARS-CoV-2 transmitted it to an individual they contacted, vary by contact type: household, classroom, random school, and out-of-school social/childcare contacts. While there is substantial variation in household attack rates across geographic locations, corresponding to cultural norms and precautions adopted, we assumed, based on two meta-analyses, a 20% household adult-adult secondary attack rate over the full duration of an infection, translating to approximately 4% per day (40,41).

For school based transmission, we allowed adult-adult attack rates to reach up to 3% per day for scenarios with low uptake of masks and distancing, a downward adjustment from the household attack rate to account for less close contact in professional settings. We developed the scale of reductions in transmission from mitigation measures in line with observations from household settings with high prevention measures (33,59) and based on effectiveness analyses for measures like masks (60,61).

Relative susceptibility and infectiousness in children

Household contact tracing studies suggest that adults are likely more susceptible to COVID-19 and more apt to transmit it when infected than children, although data, particularly for the latter, are equivocal and limited by availability and timing of testing (62). Several sources indicate that any differences wane by the teenage years, possibly as early as age 10 (8,10,33,63,64). We assumed that elementary school children were half as susceptible as adults while high school children were equally susceptible. Data are similarly limited to inform transmission from children with COVID-19. We further specified that elementary school children are half as infectious as adults and high school students were equally as infectious as adults (33,63). We discuss the data underlying these assumptions in the Supplement and varied these assumptions in sensitivity analyses.

Asymptomatic transmission and overdispersion

We assumed that individuals with fully asymptomatic disease transmit COVID-19 at half the rate of those with any symptoms (38). Individuals with mild, subclinical symptoms but who were not fully asymptomatic transmitted at the same rate as those with clinical symptoms as defined above, although only the latter were assumed to self-isolate. While reported heterogeneity in transmission may be partially driven by differences in contact rates, we sampled individual transmissibility according to a lognormal distribution to allow for some variation by individual characteristics (e.g. viral load) in adults (42,43).

IMPLEMENTATION AND CLINICAL OUTCOMES

We first evaluated the downstream impact of a single infectious individual. We randomly designated one member of the school (student or staff) as infected, starting on a random day of the week. We assessed the spread of the virus over 30 days, as in most cases either all infections were resolved over that time horizon or the spread was sufficiently large that additional public health measures (e.g. school closures) would likely be adopted. For each scenario, we ran our model 2000 times and summarized:

- 1. Mean number of infections generated in the school over 30 days after the index case
- 2. Percentage of scenarios without transmission from the index case
- 3. Percentage of scenarios with more than 5 in-school transmissions

We also describe the composition of secondary cases (proportion occurring in students, staff, and family members of students or staff).

Second, we quantified SARS-CoV-2 infections among the school community across a typical school quarter, given a constant local incidence. On each day over 8 weeks, every susceptible individual had a probability of becoming spontaneously infected outside of school that is equivalent to an age-adjusted community per capita daily incidence (with children and adolescents at half the rate of adults), distinct from their contact-dependent risk within the school network. Local incidence is intended to reflect reported case counts, and thus we assume that cases are underreported by a factor of 3 (65). When schools were remote, we assumed, similar to the analysis of hybrid schools, that each family interacted with 1 additional family on each day they did not attend school. For each scenario, we summarized:

- 1. Cumulative incidence overall and by type
- 2. Incremental incidence compared to remote learning

Transmission control threshold: We defined a reopening strategy as controlling transmission in a group (students, educators/staff, or families) if it resulted in less than a one point increase in the percentage of the group that was infected, compared to remote learning. This threshold is consistent with thresholds used by similar papers, with the objective of minimal in-school transmission (30,66).

The model was implemented in R 4.0.2. Model code is publicly available as an R package at https://github.com/abilinski/BackToSchool2.

ROLE OF FUNDING SOURCES

Funders had no role in the design or conduct of this study or in the decision to submit this work for publication.

RESULTS

In-school transmission after the introduction of a single infection

In-school mitigation

In elementary schools with low mitigation and classroom quarantine under a 5-day schedule, we projected an average of 1.7 secondary cases over 30 days following infection of a single index case (Figure 2A, left panel). This fell to 0.9 cases with medium mitigation, and further to 0.3 cases with high mitigation. Under classroom quarantine, transmission was most reduced by replacing the 5-day schedule with an A/B schedule (to 0.1-0.4 secondary cases depending on uptake of in-school prevention). With high mitigation, there were on average <0.5 secondary transmissions per case in all scenarios.

In high schools, we found the potential for larger outbreaks after a single introduction into the school, particularly when uptake of in-school prevention was low (Figure 2B). For example, with low mitigation and classroom quarantine under a 5-day schedule, we projected 23 secondary cases in the school community over a 30-day period (in the absence of additional public health responses like school closure). High uptake of in-school mitigation reduced this to 2.0 cases.

Quarantine, teacher vaccination, and screening

In elementary schools, classroom quarantine had a modest impact on transmission, although its impact was greater if the index case was a teacher (Figure S1). In high schools, classroom quarantine reduced projected average transmissions by a factor of 0.3 under low mitigation, but had smaller impact with high mitigation. We found a low impact of teacher vaccination on overall transmission, but substantial impact on transmission to teachers. Teacher vaccination reduced secondary infections over 30 days to 91% and 97% of the average without vaccination, in elementary and high schools respectively. In both settings, it reduced staff secondary incidence to about a third of its initial rate, although the reduction was slightly greater if the index case was a teacher.

Weekly screening was projected to reduce secondary cases by a large degree compared to symptomatic isolation, classroom quarantine, or teacher vaccination: to an average of 0.3-0.8 with a 5-day schedule in elementary schools (varying by uptake of in-school prevention) and to 0.9-5.0 in high schools (Figures 2A and 2B respectively, right panel). The impact of weekly screening following a single introduced case was

greatest for settings with low mitigation: weekly screening reduced average projected secondary cases under classroom quarantine from 1.7 to 0.8 in elementary schools and from 23 to 5.0 in high schools.

Detection of school-related transmission

Across all scenarios, for elementary schools, we estimated that 73% of secondary cases would occur in students, 19% in families, and 8% in teachers and staff. For high schools, 77% would occur in students, 19% in families, and 4% in teachers and staff (Figures S2 and S3). As children are less likely than adults to have symptoms, we projected that 14% of all secondary infections in elementary school communities and 15% in high school communities would be clinically symptomatic and therefore detectable without asymptomatic testing (Figure S4). With a more expansive definition of any symptoms, these percentages increased to 28% and 29% respectively.

Stochastic variation in secondary transmission

Re-running the simulation 2000 times with each set of parameters, we observed considerable variability across possible outcomes (Figure 3). In elementary schools with high mitigation and classroom quarantine under a 5-day schedule, there was a 79% chance of zero secondary cases over a 30-day period (Figure 3A, left panel). However, there was a 0.3% chance of five or more secondary cases. The chance of >5 secondary cases occurring was higher when mitigation uptake was lower (Figure 3A, left and middle panels), reaching 52% of simulations with zero secondary cases, and 8.3% with five or more under low mitigation. In high schools with symptomatic isolation, the probability of no secondary cases similarly ranged from 18-51%, and the probability of 5 or more ranged from 61%, with long tails of outliers. However, with any interventions, the long tail of large outbreaks was substantially reduced.

Transmissions over the course of the semester

With any of the modeled scenarios (symptomatic isolation, classroom quarantine, teacher vaccination, and weekly screening), both 5-day and A/B schedules increased transmission compared to remote learning. This increase was greater when local incidence was higher, especially considering infection risk among staff (Figures 4-5). Assuming that additional measures such as school closure were not taken, Figures 4 and 5 display the values of local incidence at which our predefined threshold for "controlling transmission" was met. In elementary schools with medium mitigation under a 5-day schedule, all strategies met this threshold when local incidence fell at or below 10 cases per 100,000 per day. The control threshold for the full population was exceeded at 100/100,000/day with symptomatic isolation (cumulative incidence 18%, increment 1.5%). Evaluating the threshold specifically among teachers, the control threshold was never met when local incidence was ≥ 25 per 100,000/day (e.g., total incidence: 7.4%, increment 1.8% with classroom quarantine), unless teachers were vaccinated. With both high mitigation and teacher vaccination, strategies met the control threshold among teachers at all rates up to 100 per 100,000 cases/day.

In high schools, stronger mitigation or prevention strategies would be required to meet the control threshold. Under medium mitigation and local incidence of 100 cases per 100,000/day, only weekly screening met this threshold for the full population with 5-day attendance. Under high mitigation, all strategies except symptomatic isolation met this threshold. For the teacher control threshold in high schools, high mitigation combined with either weekly screening or teacher vaccination kept the increase in teacher cumulative incidence below the control threshold at 50 cases per 100,000/day.

Sensitivity analyses

If children were as infectious as adults, the average number of secondary cases in elementary schools over 30 days was 1.9 times higher compared to the base case; if adolescents were half as susceptible as adults, secondary cases in high schools were reduced by a factor of 0.3 (Figures S1, S5). When we repeated our analysis of elementary schools assuming equivalent variation in individual infectiousness for children as for adults, we found more instances of no onward transmission and a lower average number of secondary infections over 30 days, but also slightly larger outbreaks when they occurred (Figures S1, S5). After a single introduction, all types of 2-day schedules (e.g. MW/ThF vs. MT/WTh) led to similar numbers of secondary infections, similar chance of any secondary infection, and similar numbers of secondary infections among teachers/staff (Figures S1, S5). Over the course of a quarter, the benefits of hybrid scheduling generally persisted across increased levels of out-of-school mixing (Figures S6-S7). In order for student and caregiver secondary infections to be greater with an A/B model than a 5-day model in any scenario, >9 families needed to interact on each day out of school.

DISCUSSION

While in-person education poses some COVID-19 transmission risk, the results of this simulation model underscore that it is possible to offset this with adequate precautions, particularly in elementary schools and when community transmission is well-controlled. In elementary schools with adherence to masking and distancing, our results demonstrate most cases introduced into a school would lead to little or no onward transmission. Nevertheless, if transmission occurs, it may be difficult to link to the school, and even modeled scenarios that commonly lead to no in-school transmission can occasionally generate larger outbreaks in a school community. The risk of this is substantially higher in high schools compared to elementary schools.

While it is difficult to determine the exact risk of these low-probability but high-consequence events, local transmission determines the number of cases that enter a school and therefore such rare but consequential outbreaks become increasingly likely when local incidence is high. The guidance from the US Centers for Disease Control and Prevention regarding K-12 education highlights population-adjusted incidence as a

core indicator (67), and our results provide additional evidence supporting use of this metric for school decisions in the absence of in-school screening or surveillance data. Nevertheless, while schools may decide that in-school transmission risk is too high above certain levels of local incidence, our results do not suggest that K-12 education with mitigation and/or modified schedules is likely to be a primary driver of sustained community transmission. In the event that community-level R(t) exceeds 1, it may be possible to lower this through targeted closure of high risk venues, and maintain R(t) below 1 while schools remain open.

Our results are compatible with global observations regarding school outbreaks, in which numerous well-studied index cases have produced no or minimal secondary cases, but larger outbreaks have also been recorded, particularly in secondary schools (10,68,69). Teachers and staff tend to be overrepresented in school outbreaks relative to their presence in a school community. At the elementary level, they often represent a third or more of diagnosed secondary cases (7,12). As a substantial fraction of staff may be at high risk of complications, it is important that schools undertake precautions specifically to prevent transmission among staff. Hospitals have similarly found patterns of staff-to-staff transmission and implemented measures such as reducing opportunities for communal food consumption (70).

We predict most in-school transmission will occur in the classroom during sustained contact, and interventions that reduce classroom transmission can be highly effective, including distancing, masking, or reducing class sizes. Reduced density may be achieved through the addition of more staff and moving into previously unused spaces, allowing families to opt out of in-person learning while maintaining current staffing levels, prioritizing a subset of vulnerable students for limited in-person slots, or implementation of hybrid scheduling. Some have raised concerns that a hybrid model could paradoxically increase SARS-CoV-2 transmission in schools by leading to greater out-of-cohort mixing on days when children are not physically in school. However, we find that the A/B schedule leads to far fewer infections in the school community than a 5-day schedule, a result that persists even with an assumption of substantial out-of-school mixing between families.

Weekly asymptomatic SARS-CoV-2 screening is particularly valuable for facilitating 5-day schedules in light of uncertainty in model parameters and outcomes. There is still debate surrounding relative susceptibility and transmissibility of children compared to adults, overdispersion of infectiousness, and, in particular, the degree to which asymptomatic children transmit infection, all of which have substantial impacts on transmission in a school setting. It may also be difficult to ascertain local behavioral parameters such as adherence to masking, distancing, and quarantine protocols, which can vary across settings and over time. Coupled with stochastic uncertainty in outbreak size and a high proportion of subclinical illnesses in school

populations, it may be difficult to detect transmission when it occurs. The low probability of clinical disease for infected children means that transmission chains in schools are likely to be only partially observed, and linked cases may be mistakenly classified as isolated introductions. While regular screening can both improve these data and prevent transmission through early detection, at a minimum, schools should be on alert for signs that an outbreak is brewing, and consider screening in response to the detection of any case without a clearly identified source. Other factors not considered in this work include changes in parent behavior and increased out-of-home work when children return to school, particularly for mothers and single parents, as well as changes in staff social interactions outside of school (71,72).

In addition, new variants of concern have recently been identified that are more transmissible than the strains that have previously dominated (73,74). One variant, B.1.1.7, has been linked to a large outbreak in a primary school in the Netherlands and triggered a new wave of school closures in Europe (19,75). In areas where these strains dominate in the United States, classroom-based efforts at infection prevention such as masking and distancing may be less effective at suppressing attack rates. For instance, schools which have achieved attack rates commensurate with "medium uptake of mitigation" may find that the same measures will result in attack rates reflecting "low uptake." This shift highlights the urgent need for data and underscores the added value of routine asymptomatic screening.

There are inevitable trade-offs between school disruption, risk of in-school transmission, and resources required to implement reopening strategies. We offer a quantitative perspective on the way that in-school transmissions are likely to occur, as well as the impact that proposed protocols could have on that transmission risk. We emphasize that particularly among young children, schools appear to be 'mirrors' of local transmission, rather than 'amplifiers' or 'brakes.' Thus, a reliable way to ensure a low infection risk in school is to reduce infectious introductions, by keeping local incidence low. However, even when introductions occur, high adherence to in-school prevention measures, complemented by regular asymptomatic testing and teacher vaccination, can also permit return to in-person education with controlled risk of in-school COVID-19 transmission. Local, state, and federal agencies should prioritize these effective interventions that permit the benefits of in-person education while protecting the safety of both students and educators.

This article was first published as a preprint on MedRxiv on January 29, 2021: https://www.medrxiv.org/content/10.1101/2021.01.27.21250388v1.

Grant Support: The authors were supported by the Centers for Disease Control and Prevention though the Council of State and Territorial Epidemiologists (NU38OT000297-02; AB, JAS), the National Institute of Allergy and Infectious Diseases (R37AI058736-16S1; AC and K01AI141576; MCF), the National Institute on Drug Abuse (3R37DA01561217S1; JAS); and Facebook (unrestricted gift; JG, AB, JAS).

Address for corresponding author: Meagan C. Fitzpatrick (meagan.fitzpatrick@som.umaryland.edu)

Current Addresses for All Authors:

Alyssa Bilinski (abilinski@g.harvard.edu)

Joshua Salomon (salomon1@stanford.edu)

Andrea Ciaranello (aciaranello@mgh.harvard.edu)

John Giardina (jgiardina@g.harvard.edu)

Meagan C. Fitzpatrick (meagan.fitzpatrick@som.umaryland.edu)

Reproducible Research Statement: Model code is publicly available as an R package at https://github.com/abilinski/BackToSchool2. Replication code is also available on GitHub.

REFERENCES

- 1. Map: Coronavirus and School Closures Education Week. Education Week [Internet]. 2020 Mar [cited 2020 Jun 11]; Available from: https://www.edweek.org/ew/section/multimedia/map-coronavirus-and-school-closures.html
- 2. K-12 School Opening Tracker [Internet]. [cited 2020 Nov 25]. Available from: https://cai.burbio.com/school-opening-tracker/
- 3. 2020-2021 Boston Public Schools Dashboard [Internet]. Google Data Studio. [cited 2021 Apr 21]. Available from: http://datastudio.google.com/reporting/77e82eef-349a-4f00-bd50-0824cd0df332/page/1vChB?feature=opengraph
- 4. Shapiro E. Over 50,000 N.Y.C. Public school students will return to classrooms, including in middle and high school. The New York Times [Internet]. 2021 Apr [cited 2021 Apr 21]; Available from: https://www.nytimes.com/2021/04/12/nyregion/nyc-public-schools-students.html
- 5. Shapiro E. How de Blasio Backed Himself Into a Corner on Closing Schools. The New York Times [Internet]. 2020 Nov [cited 2020 Nov 25]; Available from: https://www.nytimes.com/2020/11/24/nyregion/deblasio-school-reopening.html
- 6. Combating COVID-19 [Internet]. President-Elect Joe Biden. [cited 2020 Nov 25]. Available from: https://buildbackbetter.gov/priorities/covid-19/
- 7. Macartney K, Quinn HE, Pillsbury AJ, Koirala A, Deng L, Winkler N, et al. Transmission of SARS-CoV-2 in Australian educational settings: A prospective cohort study. The Lancet Child & Adolescent Health [Internet]. 2020 Aug [cited 2020 Sep 17];0(0). Available from: https://www.thelancet.com/journals/lanchi/article/PIIS2352-4642(20)30251-0/abstract
- 8. Fontanet A, Grant R, Tondeur L, Madec Y, Grzelak L, Cailleau I, et al. SARS-CoV-2 infection in primary schools in northern France: A retrospective cohort study in an area of high transmission. medRxiv [Internet]. 2020 Jun [cited 2020 Aug 1];2020.06.25.20140178. Available from: https://www.medrxiv.org/content/10.110 1/2020.06.25.20140178v2
- 9. Yung CF, Kam K-q, Nadua KD, Chong CY, Tan NWH, Li J, et al. Novel Coronavirus 2019 Transmission Risk in Educational Settings. Clinical Infectious Diseases [Internet]. [cited 2020 Nov 25]; Available from: https://academic.oup.com/cid/advance-article/doi/10.1093/cid/ciaa794/5862649

- 10. Stein-Zamir C, Abramson N, Shoob H, Libal E, Bitan M, Cardash T, et al. A large COVID-19 outbreak in a high school 10 days after schools' reopening, Israel, May 2020. Eurosurveillance [Internet]. 2020 Jul [cited 2020 Sep 17];25(29):2001352. Available from: https://www.eurosurveillance.org/content/10.2807/1560-7917.ES.2020.25.29.2001352
- 11. Iowa COVID-19 Tracker Please wear a mask. Please social distance. [Internet]. [cited 2020 Nov 25]. Available from: https://iowacovid19tracker.org/
- 12. Torres JP, Piñera C, De La Maza V, Lagomarcino AJ, Simian D, Torres B, et al. SARS-CoV-2 antibody prevalence in blood in a large school community subject to a Covid-19 outbreak: A cross-sectional study. Clinical Infectious Diseases [Internet]. [cited 2020 Sep 17]; Available from: https://academic.oup.com/cid/a dvance-article/doi/10.1093/cid/ciaa955/5869860
- 13. Auger KA, Shah SS, Richardson T, Hartley D, Hall M, Warniment A, et al. Association Between Statewide School Closure and COVID-19 Incidence and Mortality in the US. JAMA [Internet]. 2020 Sep [cited 2020 Nov 25];324(9):859. Available from: https://jamanetwork.com/journals/jama/fullarticle/2769034
- 14. Haug N, Geyrhofer L, Londei A, Dervic E, Desvars-Larrive A, Loreto V, et al. Ranking the effectiveness of worldwide COVID-19 government interventions. Nature Human Behaviour [Internet]. 2020 Nov [cited 2020 Nov 25];1–10. Available from: https://www.nature.com/articles/s41562-020-01009-0
- 15. Dong Y, Mo X, Hu Y, Qi X, Jiang F, Jiang Z, et al. Epidemiology of COVID-19 Among Children in China. Pediatrics [Internet]. 2020 Jun [cited 2020 Jun 11];145(6). Available from: https://pediatrics.aappublications.org/content/145/6/e20200702
- 16. Reopening schools in Denmark did not worsen outbreak, data shows. Reuters [Internet]. 2020 May [cited 2020 Jun 11]; Available from: https://www.reuters.com/article/us-health-coronavirus-denmark-reopening-idUSKBN2341N7
- 17. Pancevski B, Dandanell N. Is It Safe to Reopen Schools? These Countries Say Yes. Wall Street Journal [Internet]. 2020 May [cited 2020 Jun 11]; Available from: https://www.wsj.com/articles/is-it-safe-to-reopen-schools-these-countries-say-yes-11590928949
- 18. Vogel G, Couzin-FrankelNov. 18 J, 2020, Pm 5. As COVID-19 soars in many communities, schools attempt to find ways through the crisis [Internet]. Science AAAS. 2020 [cited 2020 Nov 25]. Available from: https://www.sciencemag.org/news/2020/11/covid-19-soars-many-communities-schools-attempt-find-ways-through-crisis

- 19. Staton B, Chazan G, Hall B, Chaffin J. European capitals follow UK with school closures as virus surges [Internet]. 2021 [cited 2021 Jan 22]. Available from: https://www.ft.com/content/8121ca0a-4d96-4cf5-b5df-a73adc16a606
- 20. Polikoff DS Anna Saavedra. Surveys show things are better for students than they were in the spring—or do they? [Internet]. Brookings. 2020 [cited 2020 Nov 20]. Available from: https://www.brookings.edu/blog/brown-center-chalkboard/2020/11/18/surveys-show-things-are-better-for-students-than-they-were-in-the-spring-or-do-they/
- 21. Gaudiano N. Missing: Millions of students [Internet]. POLITICO. [cited 2020 Nov 25]. Available from: https://politi.co/3dVrNxg
- 22. Leeb RT. Mental Health–Related Emergency Department Visits Among Children Aged 18 Years During the COVID-19 Pandemic United States, January 1–October 17, 2020. MMWR Morbidity and Mortality Weekly Report [Internet]. 2020 [cited 2020 Nov 30];69. Available from: https://www.cdc.gov/mmwr/volumes/69/wr/mm6945a3.htm
- 23. Hess A. Widespread school closures mean 30 million kids might go without meals [Internet]. CNBC. 2020 [cited 2020 Aug 1]. Available from: https://www.cnbc.com/2020/03/14/widespread-school-closuresmean-30-million-kids-might-go-without-meals.html
- 24. Baron EJ, Goldstein EG, Wallace CT. Suffering in Silence: How COVID-19 School Closures Inhibit the Reporting of Child Maltreatment [Internet]. Rochester, NY: Social Science Research Network; 2020 Jul [cited 2020 Aug 1]. Report No.: ID 3601399. Available from: https://papers.ssrn.com/abstract=3664803
- 25. Bueno C. Bricks and Mortar vs. Computers and Modems: The Impacts of Enrollment in K-12 Virtual Schools [Internet]. Rochester, NY: Social Science Research Network; 2020 Jul [cited 2020 Aug 1]. Report No.: ID 3642969. Available from: https://papers.ssrn.com/abstract=3642969
- 26. Lempel H, Hammond R, Epstein J. Costs of School Closure [Internet]. Brookings Institute; 2009 Sep [cited 2020 Aug 1]. Available from: https://www.brookings.edu/wp-content/uploads/2016/06/0930_school_closure_presentation.pdf
- 27. Soland J, Kuhfeld M, Tarasawa B, Johnson A, Ruzek E, Liu J. The impact of COVID-19 on student achievement and what it may mean for educators [Internet]. Brookings. 2020 [cited 2020 Aug 1]. Available from: https://www.brookings.edu/blog/brown-center-chalkboard/2020/05/27/the-impact-of-covid-19-on-student-achievement-and-what-it-may-mean-for-educators/

- 28. Cohen JA, Mistry D, Kerr CC, Klein DJ. Schools are not islands: Balancing COVID-19 risk and educational benefits using structural and temporal countermeasures [Internet]. Epidemiology; 2020 Sep [cited 2020 Sep 17]. Available from: http://medrxiv.org/lookup/doi/10.1101/2020.09.08.20190942
- 29. Espana G, Cavany S, Oidtman RJ, Barbera C, Costello A, Lerch A, et al. Impacts of K-12 school reopening on the COVID-19 epidemic in Indiana, USA. medRxiv [Internet]. 2020 Sep [cited 2020 Sep 17];2020.08.22.20179960. Available from: https://www.medrxiv.org/content/10.1101/2020.08.22.20179960v2
- 30. Head JR, Andrejko K, Cheng Q, Collender PA, Phillips S, Boser A, et al. The effect of school closures and reopening strategies on COVID-19 infection dynamics in the San Francisco Bay Area: A cross-sectional survey and modeling analysis. medRxiv [Internet]. 2020 Aug [cited 2020 Sep 17];2020.08.06.20169797. Available from: https://www.medrxiv.org/content/10.1101/2020.08.06.20169797v1
- 31. Bershteyn A, Kim H-Y, McGillen J, Braithwaite RS. Which policies most effectively reduce SARS-CoV-2 transmission in schools? medRxiv [Internet]. 2020 Nov [cited 2021 Apr 19];2020.11.24.20237305. Available from: https://www.medrxiv.org/content/10.1101/2020.11.24.20237305v1
- 32. McGee RS, Homburger JR, Williams HE, Bergstrom CT, Zhou AY. Model-driven mitigation measures for reopening schools during the COVID-19 pandemic [Internet]. Epidemiology; 2021 Jan [cited 2021 Apr 19]. Available from: http://medrxiv.org/lookup/doi/10.1101/2021.01.22.21250282
- 33. Park YJ, Choe YJ, Park O, Park SY, Kim Y-M, Kim J, et al. Early Release Contact Tracing during Coronavirus Disease Outbreak, South Korea, 2020 Volume 26, Number 10—October 2020 Emerging Infectious Diseases journal CDC. [cited 2020 Aug 1]; Available from: https://wwwnc.cdc.gov/eid/article/26/10/20-1315_article
- 34. CDC. COVID-19 and Your Health [Internet]. Centers for Disease Control and Prevention. 2020 [cited 2021 Jan 4]. Available from: https://www.cdc.gov/coronavirus/2019-ncov/if-you-are-sick/quarantine.html
- 35. Orscheln RC, Newland JG, Rosen DA. Practical School Algorithms for Symptomatic or SARS-CoV-2-Exposed Students Are Essential for Returning Children to In-Person Learning. The Journal of Pediatrics [Internet]. 2021 Feb [cited 2021 Apr 19];229:275–7. Available from: https://www.jpeds.com/article/S0022-3476(20)31252-X/abstract
- 36. Pediatrics: Massachusetts Chapter AA of. COVID & Kids [Internet]. [cited 2021 Apr 19]. Available from: https://mcaap.org/2018/wp-content/uploads/new-handout-copy.pages-FINAL-1.pdf?fbclid=IwAR0I HeOcdZBa_o_-Z7kWD33hSalKfk4caBgPvJ9Hgg9f7d47sFBPew0uj48

- 37. Digest of Education Statistics, 2019 [Internet]. [cited 2020 Dec 12]. Available from: https://nces.ed.gov/programs/digest/d19/tables/dt19_216.40.asp?current=yes
- 38. Byambasuren O, Cardona M, Bell K, Clark J, McLaws M-L, Glasziou P. Estimating the extent of asymptomatic COVID-19 and its potential for community transmission: Systematic review and meta-analysis. Official Journal of the Association of Medical Microbiology and Infectious Disease Canada [Internet]. 2020 Dec [cited 2021 Jan 4];5(4):223–34. Available from: https://jammi.utpjournals.press/doi/10.3138/jammi-2020-0030
- 39. Han MS, Choi EH, Chang SH, Jin B-L, Lee EJ, Kim BN, et al. Clinical Characteristics and Viral RNA Detection in Children With Coronavirus Disease 2019 in the Republic of Korea. JAMA Pediatrics [Internet]. 2020 Aug [cited 2020 Dec 13]; Available from: https://doi.org/10.1001/jamapediatrics.2020.3988
- 40. Madewell ZJ, Yang Y, Longini IM, Halloran ME, Dean NE. Household Transmission of SARS-CoV-2: A Systematic Review and Meta-analysis. JAMA Network Open [Internet]. 2020 Dec [cited 2021 Jan 3];3(12):e2031756. Available from: https://jamanetwork.com/journals/jamanetworkopen/fullarticle/277 4102
- 41. Fung HF, Martinez L, Alarid-Escudero F, Salomon JA, Studdert DM, Andrews JR, et al. The Household Secondary Attack Rate of Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2): A Rapid Review. Clinical Infectious Diseases [Internet]. 2020 Oct [cited 2021 Jan 3];ciaa1558. Available from: https://academic.oup.com/cid/advance-article/doi/10.1093/cid/ciaa1558/5921151
- 42. Kerr CC, Stuart RM, Mistry D, Abeysuriya RG, Hart G, Rosenfeld K, et al. Covasim: An agent-based model of COVID-19 dynamics and interventions. medRxiv [Internet]. 2020 May [cited 2020 Jun 2];2020.05.10.20097469. Available from: https://www.medrxiv.org/content/10.1101/2020.05.10.20097469v1
- 43. Endo A, Centre for the Mathematical Modelling of Infectious Diseases COVID-19 Working Group, Abbott S, Kucharski AJ, Funk S. Estimating the overdispersion in COVID-19 transmission using outbreak sizes outside China. Wellcome Open Research [Internet]. 2020 Apr [cited 2020 Jun 11];5:67. Available from: https://wellcomeopenresearch.org/articles/5-67/v1
- 44. Lauer SA, Grantz KH, Bi Q, Jones FK, Zheng Q, Meredith HR, et al. The Incubation Period of Coronavirus Disease 2019 (COVID-19) From Publicly Reported Confirmed Cases: Estimation and Application. Annals of Internal Medicine [Internet]. 2020 Mar [cited 2020 Jun 2];172(9):577–82. Available from: https://www.acpjournals.org/doi/10.7326/M20-0504

- 45. Li Q, Guan X, Wu P, Wang X, Zhou L, Tong Y, et al. Early Transmission Dynamics in Wuhan, China, of Novel Coronavirus–Infected Pneumonia. New England Journal of Medicine [Internet]. 2020 Jan [cited 2020 Mar 14];0(0):null. Available from: https://doi.org/10.1056/NEJMoa2001316
- 46. Gatto M, Bertuzzo E, Mari L, Miccoli S, Carraro L, Casagrandi R, et al. Spread and dynamics of the COVID-19 epidemic in Italy: Effects of emergency containment measures. Proceedings of the National Academy of Sciences [Internet]. 2020 May [cited 2020 Jun 2];117(19):10484–91. Available from: http://www.pnas.org/lookup/doi/10.1073/pnas.2004978117
- 47. He X, Lau EHY, Wu P, Deng X, Wang J, Hao X, et al. Temporal dynamics in viral shedding and transmissibility of COVID-19. Nature Medicine [Internet]. 2020 Apr [cited 2020 Apr 29];1–4. Available from: https://www.nature.com/articles/s41591-020-0869-5
- 48. He D, Zhao S, Lin Q, Zhuang Z, Cao P, Wang MH, et al. The relative transmissibility of asymptomatic COVID-19 infections among close contacts. International Journal of Infectious Diseases [Internet]. 2020 May [cited 2020 Jun 2];94:145–7. Available from: http://www.sciencedirect.com/science/article/pii/S120197122 0302502
- 49. Firth JA, Hellewell J, Klepac P, Kissler SM, CMMID COVID-19 working group, Kucharski AJ, et al. Combining fine-scale social contact data with epidemic modelling reveals interactions between contact tracing, quarantine, testing and physical distancing for controlling COVID-19 [Internet]. Epidemiology; 2020 May [cited 2020 Jun 2]. Available from: http://medrxiv.org/lookup/doi/10.1101/2020.05.26.20113720
- 50. Gallup I. Willingness to Get COVID-19 Vaccine Ticks Up to 63% in U.S. [Internet]. Gallup.com. 2020 [cited 2021 Jan 4]. Available from: https://news.gallup.com/poll/327425/willingness-covid-vaccine-ticks.aspx
- 51. Polack FP, Thomas SJ, Kitchin N, Absalon J, Gurtman A, Lockhart S, et al. Safety and Efficacy of the BNT162b2 mRNA Covid-19 Vaccine. New England Journal of Medicine [Internet]. 2020 Dec [cited 2021 Jan 4];383(27):2603–15. Available from: https://doi.org/10.1056/NEJMoa2034577
- 52. Atkeson A, Droste M, Mina MJ, Stock JH. Economic Benefits of COVID-19 Screening Tests with a Vaccine Rollout [Internet]. Health Economics; 2021 Mar [cited 2021 Mar 16]. Available from: http://medrxiv.org/lookup/doi/10.1101/2021.03.03.21252815
- 53. Larremore DB, Wilder B, Lester E, Shehata S, Burke JM, Hay JA, et al. Test sensitivity is secondary to frequency and turnaround time for COVID-19 screening. Science Advances [Internet]. 2021 Jan [cited 2021 Mar 16];7(1). Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC7775777/

- 54. Cevik M, Tate M, Lloyd O, Maraolo AE, Schafers J, Ho A. SARS-CoV-2, SARS-CoV, and MERS-CoV viral load dynamics, duration of viral shedding, and infectiousness: A systematic review and meta-analysis. The Lancet Microbe [Internet]. 2021 Jan [cited 2021 Mar 16];2(1):e13–22. Available from: https://www.sciencedirect.com/science/article/pii/S2666524720301725
- 55. Wyllie AL, Fournier J, Casanovas-Massana A, Campbell M, Tokuyama M, Vijayakumar P, et al. Saliva or Nasopharyngeal Swab Specimens for Detection of SARS-CoV-2. New England Journal of Medicine [Internet]. 2020 Aug [cited 2021 Apr 19]; Available from: https://www.nejm.org/doi/10.1056/NEJMc2016359
- 56. Kojima N, Turner F, Slepnev V, Bacelar A, Deming L, Kodeboyina S, et al. Self-Collected Oral Fluid and Nasal Swabs Demonstrate Comparable Sensitivity to Clinician Collected Nasopharyngeal Swabs for Coronavirus Disease 2019 Detection. Clinical Infectious Diseases [Internet]. 2020 Oct [cited 2021 Apr 19];(ciaa1589). Available from: https://doi.org/10.1093/cid/ciaa1589
- 57. Will M. School Workers May Get Early Shot at COVID-19 Vaccine. Will They Take It? [Internet]. Education Week. 2020 [cited 2021 Jan 4]. Available from: https://www.edweek.org/leadership/school-workers-may-get-early-shot-at-covid-19-vaccine-will-they-take-it/2020/12
- 58. Wheaton WD. U.S. Synthetic Population 2010 Version 1.0 Quick Start Guide, RTI International. 2014 May.
- 59. Cheng H-Y, Jian S-W, Liu D-P, Ng T-C, Huang W-T, Lin H-H. Contact Tracing Assessment of COVID-19 Transmission Dynamics in Taiwan and Risk at Different Exposure Periods Before and After Symptom Onset. JAMA Internal Medicine [Internet]. 2020 May [cited 2020 Jun 2]; Available from: https://jamanetwork.com/journals/jamainternalmedicine/fullarticle/2765641
- 60. Clapp PW, Sickbert-Bennett EE, Samet JM, Berntsen J, Zeman KL, Anderson DJ, et al. Evaluation of Cloth Masks and Modified Procedure Masks as Personal Protective Equipment for the Public During the COVID-19 Pandemic. JAMA Internal Medicine [Internet]. 2020 Dec [cited 2021 Jan 4]; Available from: https://jamanetwork.com/journals/jamainternalmedicine/fullarticle/2774266
- 61. Chu DK, Akl EA, Duda S, Solo K, Yaacoub S, Schünemann HJ, et al. Physical distancing, face masks, and eye protection to prevent person-to-person transmission of SARS-CoV-2 and COVID-19: A systematic review and meta-analysis. The Lancet [Internet]. 2020 Jun [cited 2021 Jan 4];395(10242):1973–87. Available from: https://linkinghub.elsevier.com/retrieve/pii/S0140673620311429
- 62. Viner RM, Mytton OT, Bonell C, Melendez-Torres GJ, Ward J, Hudson L, et al. Susceptibility to SARS-CoV-2 Infection Among Children and Adolescents Compared With Adults: A Systematic Review and

Meta-analysis. JAMA Pediatrics [Internet]. 2020 Sep [cited 2021 Jan 6]; Available from: https://jamanetwork.com/journals/jamapediatrics/fullarticle/2771181

- 63. Dattner I, Goldberg Y, Katriel G, Yaari R, Gal N, Miron Y, et al. The role of children in the spread of COVID-19: Using household data from Bnei Brak, Israel, to estimate the relative susceptibility and infectivity of children [Internet]. Infectious Diseases (except HIV/AIDS); 2020 Jun [cited 2020 Jun 19]. Available from: http://medrxiv.org/lookup/doi/10.1101/2020.06.03.20121145
- 64. Fontanet A, Tondeur L, Madec Y, Grant R, Besombes C, Jolly N, et al. Cluster of COVID-19 in northern France: A retrospective closed cohort study. medRxiv [Internet]. 2020 Apr [cited 2020 Jul 9];2020.04.18.20071134. Available from: https://www.medrxiv.org/content/10.1101/2020.04.18.20071134v1
- 65. COVID-19 Projections Using Machine Learning [Internet]. COVID-19 Projections Using Machine Learning. [cited 2021 Apr 21]. Available from: https://covid19-projections.com/
- 66. Schools and the Path to Zero Pandemics Explained [Internet]. [cited 2021 Jan 20]. Available from: https://globalepidemics.org/2020/12/18/schools-and-the-path-to-zero/
- 67. CDC. Communities, Schools, Workplaces, & Events [Internet]. Centers for Disease Control and Prevention. 2020 [cited 2020 Nov 25]. Available from: https://www.cdc.gov/coronavirus/2019-ncov/community/schools-childcare/indicators.html
- 68. Link-Gelles R. Limited Secondary Transmission of SARS-CoV-2 in Child Care Programs Rhode Island, June 1–July 31, 2020. MMWR Morbidity and Mortality Weekly Report [Internet]. 2020 [cited 2020 Sep 17];69. Available from: https://www.cdc.gov/mmwr/volumes/69/wr/mm6934e2.htm
- 69. Ladhani S, Ahmad S, Garstang J, Brent AJ, Brent B. Prospective active national surveillance of preschools and primary schools for SARS-CoV-2 infection and transmission in England, June 2020.:23.
- 70. BWH Press Release Brigham and Women's Hospital [Internet]. [cited 2020 Nov 25]. Available from: https://www.brighamandwomens.org/about-bwh/newsroom/press-releases-detail?id=3684
- 71. Beauregard P-L, Connolly M, Haeck C, Molnar TL. Primary School Reopenings and Parental Work [Internet]. Research Group on Human Capital, University of Quebec in Montreal's School of Management; 2020 Aug [cited 2021 Mar 16]. Report No.: 20-06. Available from: https://ideas.repec.org/p/grc/wpaper/20-06.html
- 72. Collins C, Ruppanner L, Christin Landivar L, Scarborough WJ. The Gendered Consequences of a

Weak Infrastructure of Care: School Reopening Plans and Parents' Employment During the COVID-19 Pandemic. Gender & Society [Internet]. 2021 Mar [cited 2021 Mar 16];08912432211001300. Available from: https://doi.org/10.1177/08912432211001300

73. Volz E, Mishra S, Chand M, Barrett JC, Johnson R, Geidelberg L, et al. Transmission of SARS-CoV-2 Lineage B.1.1.7 in England: Insights from linking epidemiological and genetic data. medRxiv [Internet]. 2021 Jan [cited 2021 Jan 6];2020.12.30.20249034. Available from: https://www.medrxiv.org/content/10.1101/2020 .12.30.20249034v2

74. Walensky RP, Walke HT, Fauci AS. SARS-CoV-2 Variants of Concern in the United States—Challenges and Opportunities. JAMA [Internet]. 2021 Mar [cited 2021 Apr 23];325(11):1037. Available from: https://jamanetwork.com/journals/jama/fullarticle/2776739

75. VogelJan. 15 G, 2021, Pm 4. School risk calculations scrambled by fast-spreading virus strains [Internet]. Science AAAS. 2021 [cited 2021 Jan 22]. Available from: https://www.sciencemag.org/news/2021/01/new-coronavirus-variant-scrambles-school-risk-calculations

76. Davies NG, Klepac P, Liu Y, Prem K, Jit M, Eggo RM. Age-dependent effects in the transmission and control of COVID-19 epidemics. Nature Medicine [Internet]. 2020 Aug [cited 2021 Jan 24];26(8):1205–11. Available from: https://www.nature.com/articles/s41591-020-0962-9

77. Zhu Y, Bloxham CJ, Hulme KD, Sinclair JE, Tong ZWM, Steele LE, et al. A meta-analysis on the role of children in SARS-CoV-2 in household transmission clusters. medRxiv [Internet]. 2020 Dec [cited 2021 Jan 24];2020.03.26.20044826. Available from: https://www.medrxiv.org/content/10.1101/2020.03.26.20044826v2

78. Exclusive: Kids catch and spread coronavirus half as much as adults, Iceland study confirms [Internet]. Science. 2020 [cited 2021 Jan 24]. Available from: https://www.nationalgeographic.com/science/2020/12/we-now-know-how-much-children-spread-coronavirus/

79. Kim J, Choe YJ, Lee J, Park YJ, Park O, Han MS, et al. Role of children in household transmission of COVID-19. Archives of Disease in Childhood [Internet]. 2020 Aug [cited 2021 Jan 24]; Available from: https://adc.bmj.com/content/early/2020/08/06/archdischild-2020-319910

80. Lee EJ, Kim DH, Chang SH, Suh SB, Lee J, Lee H, et al. Absence of SARS-CoV-2 Transmission from Children in Isolation to Guardians, South Korea - Volume 27, Number 1—January 2021 - Emerging Infectious Diseases journal - CDC. [cited 2021 Jan 24]; Available from: https://wwwnc.cdc.gov/eid/article/27/1/20-3450_article

- 81. Goldstein E, Lipsitch M, Cevik M. On the Effect of Age on the Transmission of SARS-CoV-2 in Households, Schools, and the Community. The Journal of Infectious Diseases [Internet]. 2020 Oct [cited 2021 Jan 24];(jiaa691). Available from: https://doi.org/10.1093/infdis/jiaa691
- 82. Carcione D, Giele CM, Goggin LS, Kwan KS, Smith DW, Dowse GK, et al. Secondary attack rate of pandemic influenza A(H1N1)2009 in Western Australian households, 29 May–7 August 2009. Eurosurveillance [Internet]. 2011 Jan [cited 2020 Jun 19];16(3):19765. Available from: https://www.eurosurveillance.org/content/10.2807/ese.16.03.19765-en
- 83. Odaira F, Takahashi H, Toyokawa T, Tsuchihashi Y, Kodama T, Yahata Y, et al. Assessment of secondary attack rate and effectiveness of antiviral prophylaxis among household contacts in an influenza A(H1N1)v outbreak in Kobe, Japan, May–June 2009. Eurosurveillance [Internet]. 2009 Sep [cited 2020 Jun 19];14(35):19320. Available from: https://www.eurosurveillance.org/content/10.2807/ese.14.35.19320-en
- 84. Sikora C, Fan S, Golonka R, Sturtevant D, Gratrix J, Lee BE, et al. Transmission of pandemic influenza A (H1N1) 2009 within households: Edmonton, Canada. Journal of Clinical Virology [Internet]. 2010 Oct [cited 2020 Jun 19];49(2):90–3. Available from: http://www.sciencedirect.com/science/article/pii/S13866532100 02593
- 85. Sugimoto JD, Borse NN, Ta ML, Stockman LJ, Fischer GE, Yang Y, et al. The Effect of Age on Transmission of 2009 Pandemic Influenza A (H1N1) in a Camp and Associated Households. Epidemiology (Cambridge, Mass) [Internet]. 2011 Mar [cited 2020 Jun 19];22(2). Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3755879/
- 86. Klick B, Nishiura H, Ng S, Fang VJ, Leung GM, Malik Peiris JS, et al. Transmissibility of seasonal and pandemic influenza in a cohort of households in Hong Kong in 2009. Epidemiology (Cambridge, Mass) [Internet]. 2011 Nov [cited 2020 Jun 19];22(6):793–6. Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3206962/
- 87. Wu JT, Ma ESK, Lee CK, Chu DKW, Ho P-L, Shen AL, et al. The Infection Attack Rate and Severity of 2009 Pandemic H1N1 Influenza in Hong Kong. Clinical Infectious Diseases [Internet]. 2010 Nov [cited 2020 Jun 19];51(10):1184–91. Available from: https://academic.oup.com/cid/article/51/10/1184/393386
- 88. Bi Q, Wu Y, Mei S, Ye C, Zou X, Zhang Z, et al. Epidemiology and transmission of COVID-19 in 391 cases and 1286 of their close contacts in Shenzhen, China: A retrospective cohort study. The Lancet Infectious Diseases [Internet]. 2020 Aug [cited 2021 Jan 24];20(8):911–9. Available from: https://www.thelancet.com/jo

urnals/laninf/article/PIIS1473-3099(20)30287-5/abstract

- 89. Fateh-Moghadam P, Battisti L, Molinaro S, Fontanari S, Dallago G, Binkin N, et al. Contact tracing during Phase I of the COVID-19 pandemic in the Province of Trento, Italy: Key findings and recommendations. medRxiv [Internet]. 2020 Jul [cited 2021 Jan 24];2020.07.16.20127357. Available from: https://www.medrxiv.org/content/10.1101/2020.07.16.20127357v1
- 90. Li F, Li Y-Y, Liu M-J, Fang L-Q, Dean NE, Wong GWK, et al. Household transmission of SARS-CoV-2 and risk factors for susceptibility and infectivity in Wuhan: A retrospective observational study. The Lancet Infectious Diseases [Internet]. 2021 Jan [cited 2021 Jan 24];0(0). Available from: https://www.thelancet.com/journals/laninf/article/PIIS1473-3099(20)30981-6/abstract
- 91. Grijalva CG. Transmission of SARS-COV-2 Infections in Households Tennessee and Wisconsin, April–September 2020. MMWR Morbidity and Mortality Weekly Report [Internet]. 2020 [cited 2021 Jan 24];69. Available from: https://www.cdc.gov/mmwr/volumes/69/wr/mm6944e1.htm
- 92. Lopez Bernal J, Panagiotopoulos N, Byers C, Garcia Vilaplana T, Boddington NL, Zhang X, et al. Transmission dynamics of COVID-19 in household and community settings in the United Kingdom [Internet]. Epidemiology; 2020 Aug [cited 2021 Jan 24]. Available from: http://medrxiv.org/lookup/doi/10.1101/2020. 08.19.20177188
- 93. Leclerc QJ, Fuller NM, Knight LE, CMMID COVID-19 Working Group, Funk S, Knight GM. What settings have been linked to SARS-CoV-2 transmission clusters? Wellcome Open Research [Internet]. 2020 Jun [cited 2021 Jan 24];5:83. Available from: https://wellcomeopenresearch.org/articles/5-83/v2
- 94. Jones RD. COVID-19 Trends in Florida K-12 Schools, August 10 November 14, 2020. medRxiv [Internet]. 2020 Dec [cited 2021 Jan 24];2020.11.30.20241224. Available from: https://www.medrxiv.org/content/10.110 1/2020.11.30.20241224v1
- 95. National COVID-19 School Response Dashboard [Internet]. [cited 2021 Jan 24]. Available from: https://statsiq.co1.qualtrics.com/public-dashboard/v0/dashboard/5f78e5d4de521a001036f78e#/dashboard/5f78e5d4de521a001036f78e?pageId=Page_f6071bf7-7db4-4a61-942f-ade4cce464de
- 96. Covid-19: Sparkletots Preschool cluster increases to 20 cases, forming 3rd largest local cluster [Internet]. [cited 2021 Jan 24]. Available from: https://mothership.sg/2020/03/covid-19-sparkletots-preschool-coi/
- 97. Heavey L, Casey G, Kelly C, Kelly D, McDarby G. No evidence of secondary transmission of COVID-19

from children attending school in Ireland, 2020. Eurosurveillance [Internet]. 2020 May [cited 2021 Jan 24];25(21). Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC7268273/

- 98. Zimmerman KO, Akinboyo IC, Brookhart MA, Boutzoukas AE, McGann KA, Smith MJ, et al. Incidence and Secondary Transmission of SARS-CoV-2 Infections in Schools. Pediatrics [Internet]. 2021 Apr [cited 2021 Apr 21];147(4):e2020048090. Available from: http://pediatrics.aappublications.org/lookup/doi/10.1542/pedia.2020-048090
- 99. Falk A. COVID-19 Cases and Transmission in 17 K–12 Schools Wood County, Wisconsin, August 31–November 29, 2020. MMWR Morbidity and Mortality Weekly Report [Internet]. 2021 [cited 2021 Apr 21];70. Available from: https://www.cdc.gov/mmwr/volumes/70/wr/mm7004e3.htm
- 100. Gillespie DL, Meyers LA, Lachmann M, Redd SC, Zenilman JM. The Experience of Two Independent Schools with In-Person Learning During the COVID-19 Pandemic [Internet]. Public; Global Health; 2021 Jan [cited 2021 Feb 6]. Available from: http://medrxiv.org/lookup/doi/10.1101/2021.01.26.21250065
- 101. Brandal LT, Ofitserova TS, Meijerink H, Rykkvin R, Lund HM, Hungnes O, et al. Minimal transmission of SARS-CoV-2 from paediatric COVID-19 cases in primary schools, Norway, August to November 2020. Eurosurveillance [Internet]. 2021 Jan [cited 2021 May 14];26(1):2002011. Available from: https://www.eurosurveillance.org/content/10.2807/1560-7917.ES.2020.26.1.2002011
- 102. Fong MW, Cowling BJ, Leung GM, Wu P. Letter to the editor: COVID-19 cases among school-aged children and school-based measures in Hong Kong, July 2020. Eurosurveillance [Internet]. 2020 Sep [cited 2021 May 14];25(37):2001671. Available from: https://www.eurosurveillance.org/content/10.2807/1560-7917.ES.2020.25.37.2001671?crawler=true
- 103. Doyle T. COVID-19 in Primary and Secondary School Settings During the First Semester of School Reopening Florida, August–December 2020. MMWR Morbidity and Mortality Weekly Report [Internet]. 2021 [cited 2021 May 14];70. Available from: https://www.cdc.gov/mmwr/volumes/70/wr/mm7012e2.ht m
- 104. Ismail SA, Saliba V, Bernal JL, Ramsay ME, Ladhani SN. SARS-CoV-2 infection and transmission in educational settings: A prospective, cross-sectional analysis of infection clusters and outbreaks in England. The Lancet Infectious Diseases [Internet]. 2020 Dec [cited 2021 Jan 14];0(0). Available from: https://www.thelancet.com/journals/laninf/article/PIIS1473-3099(20)30882-3/abstract
- 105. COVID-19 Schools Infection Survey Round 1, England Office for National Statistics [Internet]. [cited

2021 Jan 24]. Available from: https://www.ons.gov.uk/peoplepopulationandcommunity/healthandsocialcare/conditionsanddiseases/bulletins/covid19schoolsinfectionsurveyround1england/november2020

106. Oster E. Opinion Schools are not spreading covid-19. This new data makes the case. Washington Post [Internet]. 2020 Nov [cited 2021 Jan 24]; Available from: https://www.washingtonpost.com/opinions/2020/11/20/covid-19-schools-data-reopening-safety/

107. COVID Testing Results [Internet]. web. [cited 2021 Jan 24]. Available from: https://www.schools.nyc.gov/school-year-20-21/return-to-school-2020/health-and-safety/covid-19-testing/covid-testing-results

108. Second school in Thorncliffe neighbourhood closed over COVID; third York Catholic school shuts down [Internet]. thestar.com. 2020 [cited 2021 Jan 24]. Available from: https://www.thestar.com/news/gta/2020/12/07/tdsb-closes-thorncliffe-park-elementary-school-amid-rising-number-of-students-testing-positive-for-covid-19.html

109. Monod M, Blenkinsop A, Xi X, Hebert D, Bershan S, Bradley VC, et al. Age groups that sustain resurging COVID-19 epidemics in the United States. medRxiv [Internet]. 2020 Sep [cited 2021 Jan 24];2020.09.18.20197376. Available from: https://www.medrxiv.org/content/10.1101/2020.09.18.20197376v1

110. Harris D, Ziedan E, Hassig S. The Effects of School Reopenings on COVID-19 Hospitalizations [Internet]. [cited 2021 Jan 24]. Available from: https://www.reachcentered.org/publications/the-effects-of-school-reopenings-on-covid-19-hospitalizations

111. Goldhaber D, Imberman S, Strunk KO, Hopkins B, Brown N, Harbatkin E, et al. To What Extent Does In-Person Schooling Contribute to the Spread of COVID-19? Evidence from Michigan and Washington [Internet]. 2020 Dec [cited 2021 Jan 24]. Available from: https://epicedpolicy.org/does-in-person-schooling-contribute-to-the-spread-of-covid-19/

112. Berg P van den, Schechter-Perkins EM, Jack RS, Epshtein I, Nelson R, Oster E, et al. Effectiveness of three versus six feet of physical distancing for controlling spread of COVID-19 among primary and secondary students and staff: A retrospective, state-wide cohort study. Clinical Infectious Diseases [Internet]. 2021 Mar [cited 2021 Apr 19];(ciab230). Available from: https://doi.org/10.1093/cid/ciab230

113. Liang M, Gao L, Cheng C, Zhou Q, Uy JP, Heiner K, et al. Efficacy of face mask in preventing respiratory virus transmission: A systematic review and meta-analysis. Travel Medicine and Infectious Disease [Internet]. 2020 [cited 2021 Apr 23];36:101751. Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC725 3999/

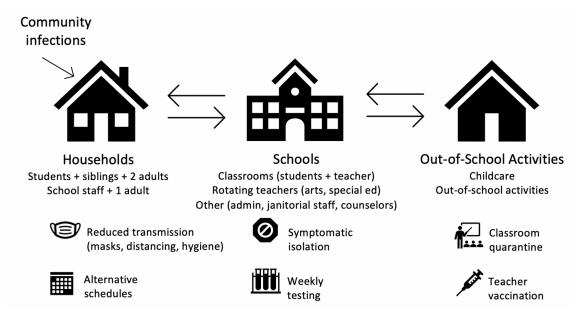


Figure 1: Model diagram. The model includes 3 primary domains: households, schools, and out-of-school social/childcare mixing and incorporates a range of interventions to prevent or reduce transmission.

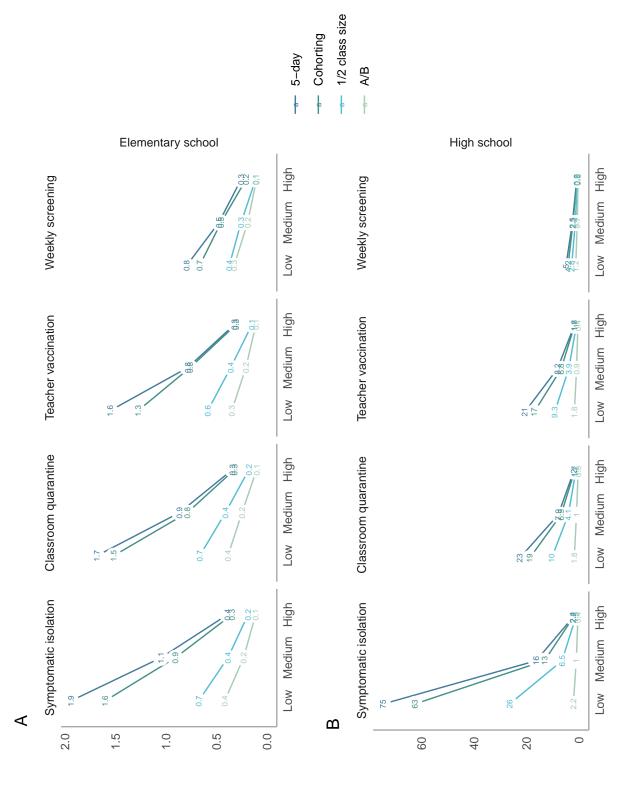


Figure 2: Average number of total secondary transmissions over 30 days (outside of the index case's household) following a single introduction into a school. (This is not an estimate of R, the effective reproduction number, which is displayed in Figure S10.) These include both transmission directly from the index case, as well as from secondary and tertiary cases. The top panel shows elementary schools, where children are assumed to be less susceptible and less infectious, while the bottom panel shows high schools. Note that axes differ across rows. The x-axes vary the level of prevention measure uptake, with low uptake assuming minimal interventions and high uptake assuming intensive interventions. Line colors correspond to scheduling strategies.

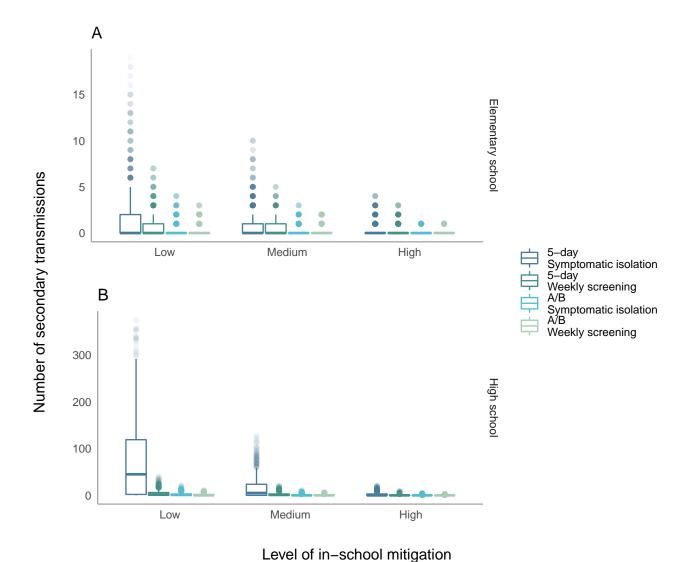


Figure 3: Distribution of secondary transmissions when a single case is introduced. The x-axis varies the level of prevention measure uptake, with low uptake assuming minimal interventions and high uptake assuming intensive interventions. The y-axis displays the number of secondary transmissions (outside of the index case's household) when a case is introduced. Transmissions include both

those directly from the index case, as well as those from secondary and tertiary cases. Distributions are truncated at the 99.5th quantile, i.e. all outcomes occur with at least probability 1/200.

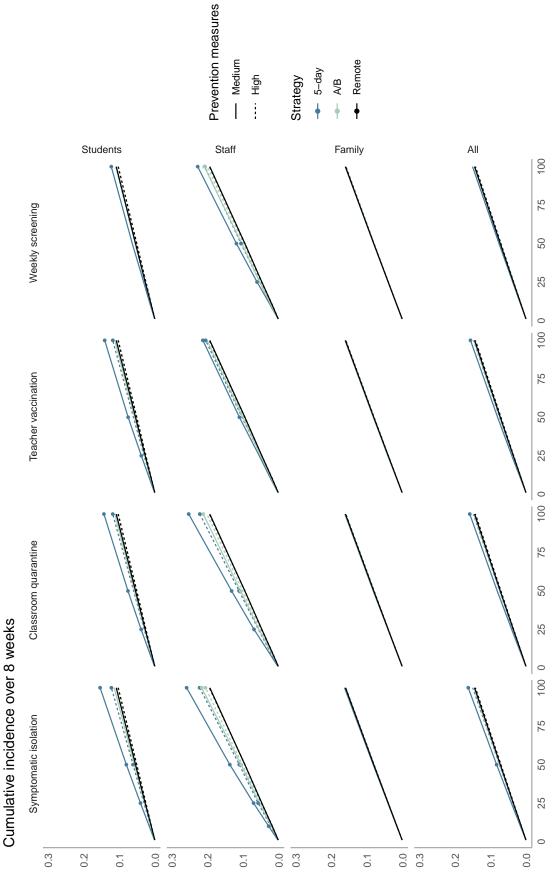


Figure 4: Cumulative incidence over 8 weeks in elementary schools. The x-axis shows the average daily local incidence per 100,000 population. The y-axis shows cumulative incidence over 8 weeks in the elementary school community. Columns denote different isolation, quarantine, vaccination, and detection strategies, while rows show different population subgroups. We mark a strategy with a point if it fails to meet the transmission control threshold for the subgroup, i.e., if there is more than a one point increase over remote learning in the percentage of the group infected during 8 weeks.

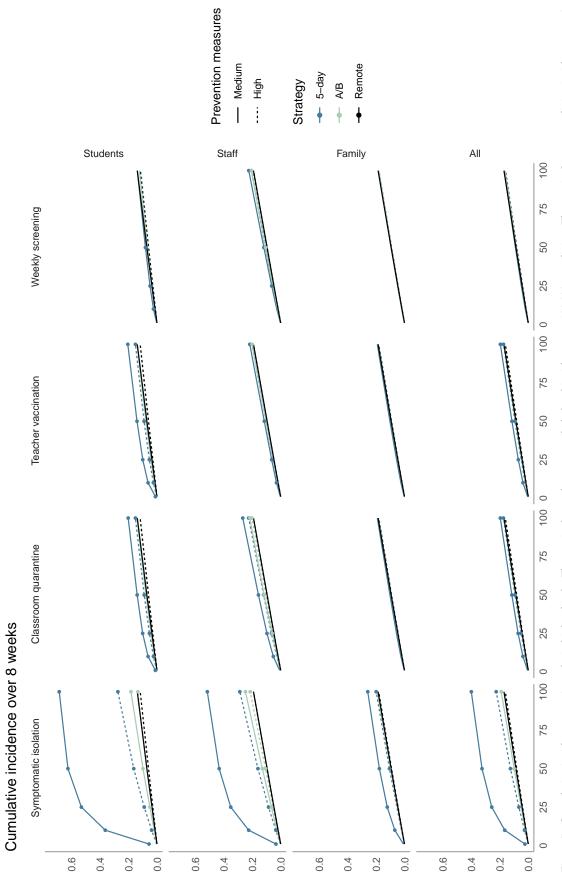


Figure 5: Cumulative incidence over 8 weeks in high schools. The x-axis shows the average daily local incidence per 100,000 population. The y-axis shows cumulative incidence over 8 weeks in the high school community. Columns denote different isolation, quarantine, vaccination, and detection strategies, while rows show different population subgroups. We mark a strategy with a point if it fails to meet the transmission control threshold for the subgroup, i.e., if there is more than a one point increase over remote learning in the percentage of the group infected during 8 weeks.

SUPPLEMENT FOR PASSING THE TEST: A MODEL-BASED ANALYSIS OF SAFE SCHOOL-REOPENING STRATEGIES

Alyssa Bilinski $MS^7 \cdot$ Joshua A. Salomon, $PhD^8 \cdot$ John Giardina, $MA^{1,9}$ Andrea Ciaranello MD, $MPH^{10} \cdot$ Meagan C. Fitzpatrick, PhD^{11}

⁷Interfaculty Initiative in Health Policy, Harvard Graduate School of Arts and Sciences (abilinski@g.harvard.edu)

⁸Center for Health Policy & Center for Primary Care and Outcomes Research, Stanford University School of Medicine

⁹Center for Health Decision Science, Harvard T.H. Chan School of Public Health

 $^{^{10}} Medical\ Practice\ Evaluation\ Center\ and\ Division\ of\ Infectious\ Disease,\ Massachusetts\ General\ Hospital,\ Harvard\ Medical\ School$

¹¹Center for Vaccine Development and Global Health, University of Maryland School of Medicine (meagan.fitzpatrick@som.umaryland.edu)

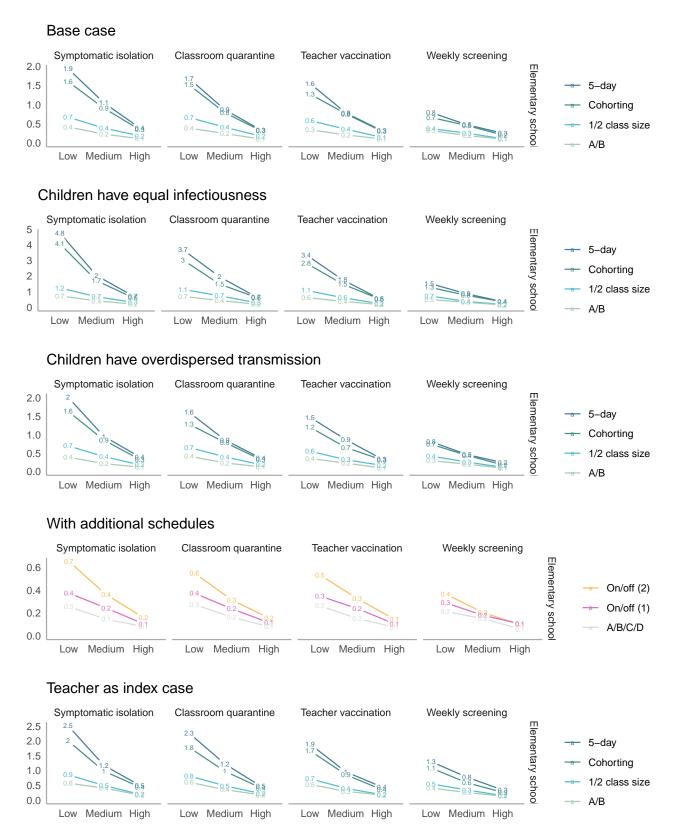


Figure S1: Sensitivity analyses (elementary schools) – average number of total secondary transmissions over 30 days (outside of the index case's household) following a single introduction into a school community. The columns vary the level of prevention measure uptake, with low uptake assuming minimal interventions and high uptake assuming intensive interventions. Line colors correspond to scheduling strategies.

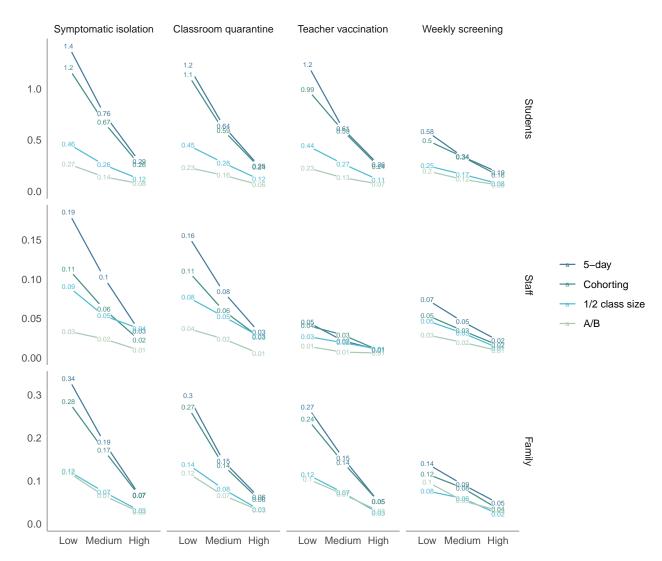


Figure S2: Sensitivity analysis - elementary school base case broken down by case type. Average number of total secondary transmissions over 30 days (outside of the index case's household) following a single introduction into an elementary school community. These include both transmission directly from the index case, as well as from secondary and tertiary cases. The x-axes varies the level of prevention measure uptake, with low uptake assuming minimal interventions and high uptake assuming intensive interventions. Line colors correspond to scheduling strategies.

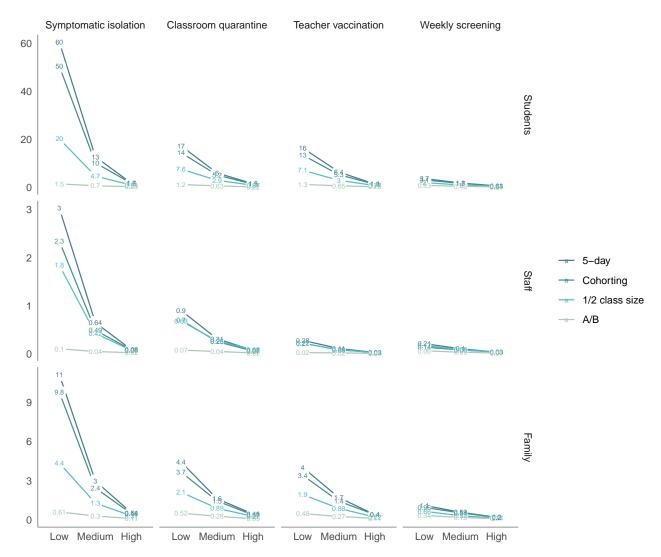


Figure S3: Sensitivity analysis - high school base case broken down by case type. Average number of total secondary transmissions over 30 days (outside of the index case's household) following a single introduction into a high school community. These include both transmission directly from the index case, as well as from secondary and tertiary cases. The x-axes varies the level of prevention measure uptake, with low uptake assuming minimal interventions and high uptake assuming intensive interventions. Line colors correspond to scheduling strategies.

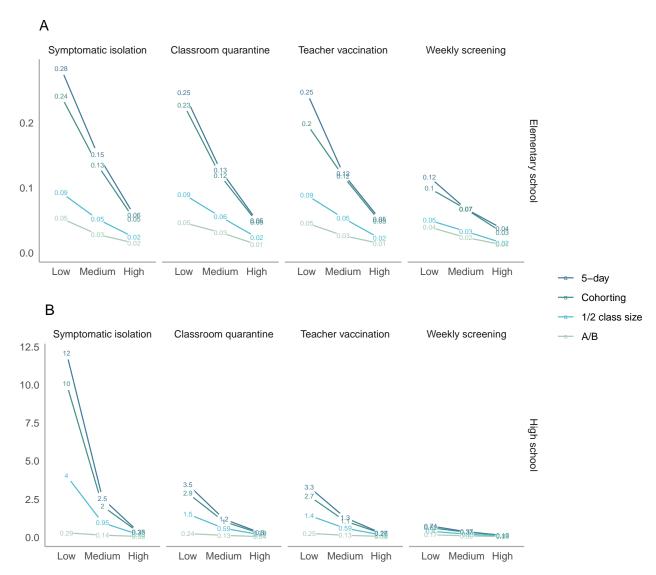


Figure S4: Average number of clinically symptomatic cases in staff and students over 30 days following a single introduction into a school community. These include both transmission directly from the index case, as well as from secondary and tertiary cases. The top panel shows elementary schools, where children are assumed to be less susceptible and less infectious, while the bottom panel shows high schools. Note that axes differ across rows. The x-axes vary the level of prevention measure uptake, with low uptake assuming minimal interventions and high uptake assuming intensive interventions. Line colors correspond to scheduling strategies.

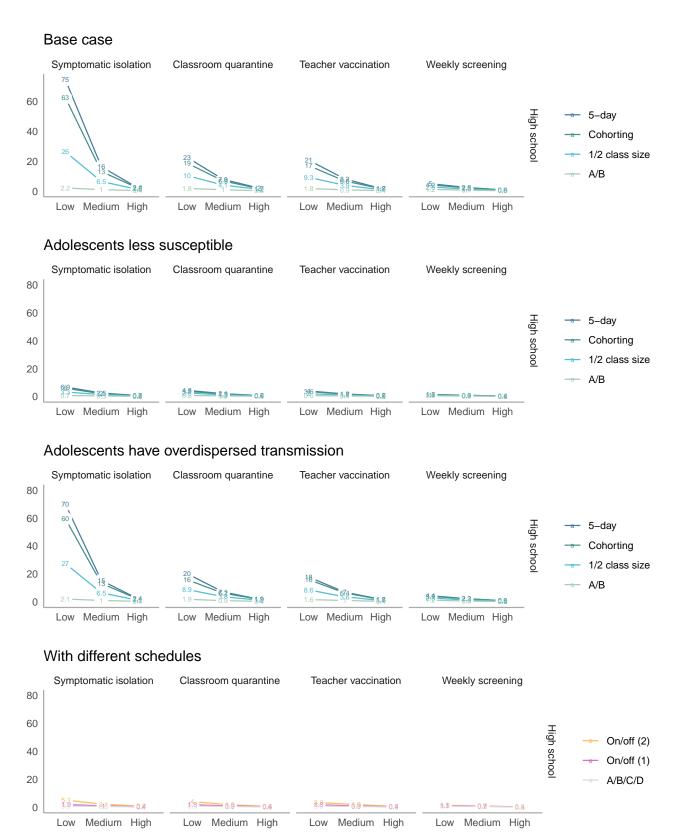


Figure S5: Sensitivity analyses (high schools) – average number of total secondary transmissions over 30 days (outside of the index case's household) following a single introduction into a school community. The columns vary the level of prevention measure uptake, with low uptake assuming minimal interventions and high uptake assuming intensive interventions. Line colors correspond to scheduling strategies.

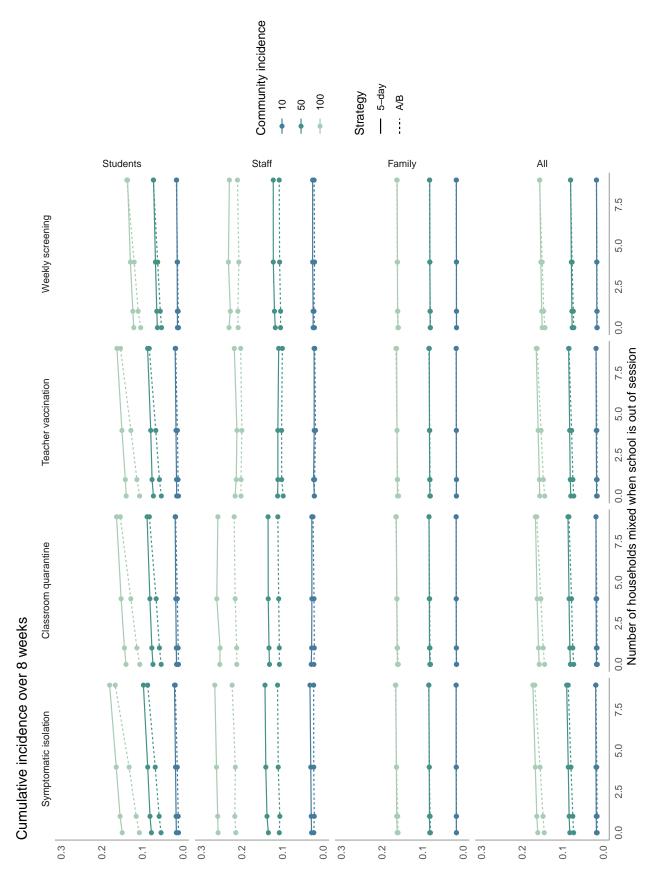


Figure S6: Cumulative incidence over 8 weeks in elementary schools across different levels of out-of-school mixing. The x-axis shows the average daily community incidence per 100,000 population. The y-axis shows cumulative incidence over 8 weeks. Columns denote different isolation, quarantine, vaccination, and detection strategies, while rows show different population subgroups.

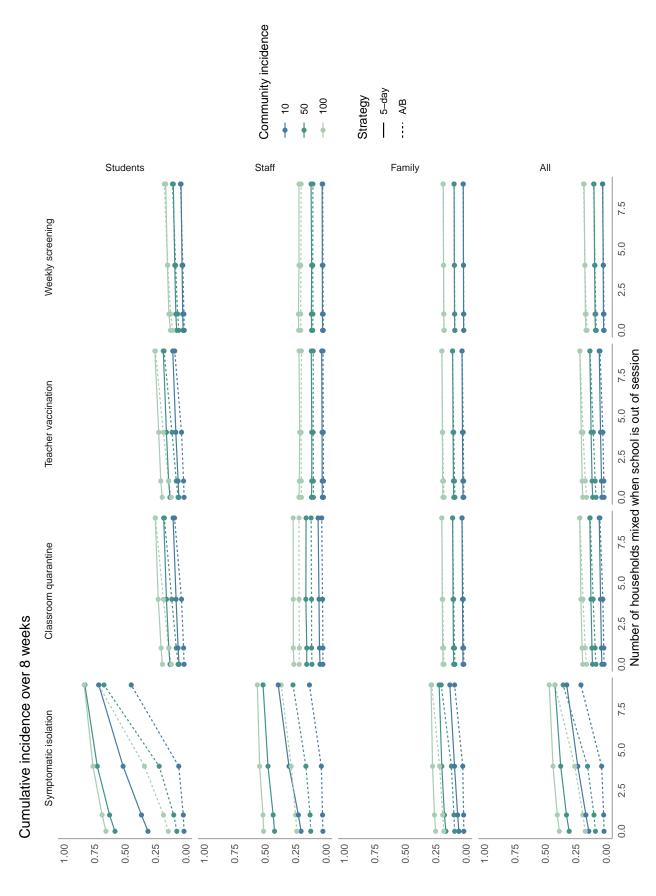


Figure S7: Cumulative incidence over 8 weeks in high schools across different levels of out-of-school mixing. The x-axis shows the average daily community incidence per 100,000 population. The y-axis shows cumulative incidence over 8 weeks. Columns denote different isolation, quarantine, vaccination, and detection strategies, while rows show different population subgroups.

MODEL

We created an agent-based stochastic SEIR model of COVID-19 transmission, as depicted in Figure S8, including susceptible (S), exposed (E), infectious (with clinical symptoms (I_S), subclinical symptoms (I_S), or asymptomatic disease (I_A), and recovered individuals (R). Only individuals in the susceptible compartment could contract a new infection, and only those in an infectious compartment could transmit disease. Individuals with clinical symptoms (I_S) were assumed to self-isolate after the appearance of symptoms, an average of 2 days after the onset of infectiousness.

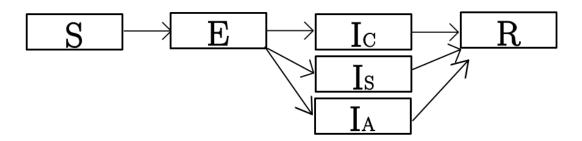


Figure S8: Model compartments.

At each daily time-step, we modeled dyadic interactions between individuals according to household, classroom, school, and childcare relationships, drawing parameter values from the distributions specified in Table 1 in the main text. A SARS-CoV-2-infected individual i transmitted to susceptible individual j at time-step t with Bernoulli probability equal to:

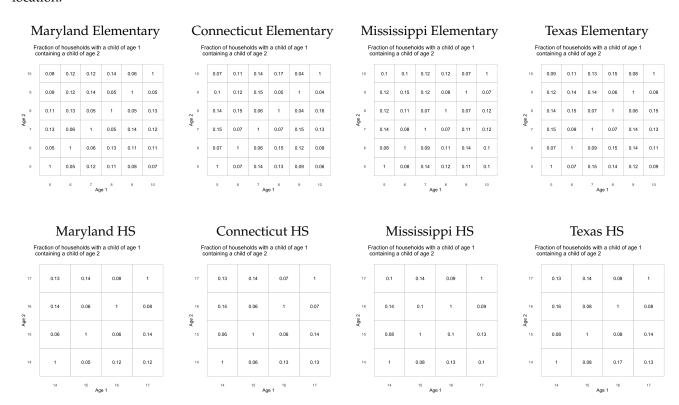
$$p_{ijt} = c_{ijkt}q_k s_j r_i a_i d_i$$

where c_{ijk} was an indicator variable equal to 1 if individuals i and j had contact type k at time t, q_k was the probability of transmission given one day of contact type k, r_i was the relative infectiousness of individual i (compared to full adult infectiousness), s_j was the relative susceptibility of individual j (compared to full adult infectiousness), and a_i indicated whether individual i had asymptomatic disease and d_i was a dispersion factor representing individual-level heterogeneity in transmissibility. Considerations for these parameters are discussed further in the text.

HOUSEHOLD STRUCTURE

We use a Framework for Reconstructing Epidemiological Dynamics (FRED) to generate household structures (58). For elementary schools, we sampled from households containing at least one child aged 5 to 10 to identify siblings attending the same school. For high schools, we sampled from households with students

aged 14 to 17. For each student, we included two adults in the household, based on the average number of household members over 25. For each staff member, we also included a household adult contact, representing a partner or roommate with whom they had close contact. For computational simplicity, we used Maryland as a representative state, as sibling structure (the main parameter of interest) did not appear sensitive to location.



COMPARISON TO OBSERVED OUTBREAKS

A number of factors make formal calibration challenging for this paper. First, most data collection has been ad hoc, with some sources biased toward reporting large outbreaks and others toward high-mitigation schools who voluntarily collect and report data. Without comprehensive data on school mitigation efforts, interpretation can be challenging, as our results emphasize that a large range of outcomes is possible due to both variation in parameters and stochastic uncertainty. Testing practices vary across schools, affecting reporting procedures, and the definition of "contact" also varies, with some sources including even brief contacts while others require more sustained interaction (e.g. >15 minutes without a mask). Overall, this section describes available data sources in comparison to our results. We emphasize that substantial uncertainty persists, and that weekly screening or other surveillance is one of the best tools available for understanding a specific context as well as early detection of outbreaks.

DIFFERENCES IN INFECTIOUSNESS AND SUSCEPTIBILITY BY AGE

In our model, we assumed that young children (10 and under) were both less susceptible and less infectious than adults. To inform these assumptions, we used a meta-analysis on child susceptibility on those under 18-20 (62), which was consistent with best-fit model estimates (76) and another study on child infectiousness (77). We also used a number of contact tracing studies suggesting not just a difference between children and adults but also an age gradient in susceptibility and infectiousness. These included a study from B'nei Brak, Israel on household infections (Figure 4) (63) and two studies from France contrasting minimal elementary school outbreaks (despite introductions) with a larger high school cluster in an area with early COVID-19 exposure (8,64). Limited data from Iceland, with comprehensive contact tracing and sequencing, suggested a similar difference between young children and adolescents in infectiousness (78). Contact tracing data from South Korea on infectiousness, while more difficult to interpret due to concurrent exposures among household members and high PPE usage of guardians of infected children (79,80), was not inconsistent with this finding (33).

We focused on data from contact tracing studies that used comprehensive testing of contacts because these were less likely to be biased. In particular, we did not want to interpret evidence that children were rarely identified as index cases (77) or had lower seroprevalence in some contexts as evidence that they were less susceptible or infectious (81). These differences could have been driven by the fact that children are less likely to have symptoms and be tested and/or that their contacts were markedly reduced by school closures early in the pandemic. While household contact tracing studies with comprehensive testing avoided this issue, they could have had other biases. For example, children were unlikely to be caretakers of sick individuals

compared to adults in the household, and may have been shielded in houses with known cases, particularly in the midst of an unprecedented pandemic. Nevertheless, in general, higher household attack rates in children have been observed for seasonal influenza and H1N1, making lower estimates for COVID-19 particularly notable (82–87).

These findings nevertheless cannot differentiate between biological explanations for lower susceptibility in younger children (e.g., lower density of ACE2 receptor) and behavioral ones (e.g., easier to restrict socialization). In addition, some studies suggest higher susceptibility and/or infectiousness of young children than we include in our base case (88–92). While we model these possibilities in sensitivity analyses, the bulk of evidence on well-studied school outbreaks has pointed to important distinctions between elementary and high school-aged children. For example, when Israel experienced significant outbreaks upon return to school in the early summer, there was a significant outbreak of 178 cases in a middle/high school (concentrated in grades 7-9), but elementary school outbreaks were generally reported as smaller (e.g. 33 cases) (10,18). This difference was also apparent in informal databases, with high schools largely responsible for outbreaks of more than 50 people (e.g. in New Zealand and the US prior to social distancing and Australia, where a school outbreak was reported to be driven by high schoolers) (93–95). In the Netherlands, a health official was quoted saying that significant outbreaks occurred mainly in high schools and universities prior to an elementary school outbreak with B.1.1.7 (75). Exceptions often included significant outbreaks among teachers (e.g., in Chile (12) and Singapore (96)).

SECONDARY CASES

Our results are broadly consistent a few key features of observed data. First, well-studied cases have led to no or minimal outbreaks in a number of settings. In passive surveillance from the United Kindom during the summer, in-school transmission was identified from 39% of index cases in secondary schools and 26% of cases in primary schools "in the context of small class or bubble sizes, half empty schools, and extensive hygiene measures." This is similar to what we predicted with an A/B model in secondary schools with medium mitigation (36%) and a low mitigation scenario in elementary schools (23%). No onward transmission was found in Singapore or Ireland, each with 3 seed cases (9,97). In Rhode Island childcare settings, which had small class sizes, onward transmission was documented in 4/29 index cases (14%), consistent with 1/2 class size scenario and high mitigation (12%) (68). In North Carolina, minimal transmission was documented with under a hybrid model with strong mitigation measures in place (32 secondary transmissions identified through 773 contact traced cases for 0.04 average secondary transmissions per index case) (98). Similarly, with masking and cohorting, rural Wisconsin schools reported few cases linked to in-school transmission (99).

In this context, COVID-19 incidence was lower than the local community rates, although this comparison was not age-adjusted for the fact that children generally have lower reported COVID-19 case rates than the general population.

However limited testing in some of these studies may have missed subclinical cases. A recent paper estimated the number of direct secondary infections to students and staff from each introduced case in the context of weekly or biweekly testing in high-mitigation K-12 independent schools to be about 0.2 to 0.5, which is within the range of our estimates of average direct secondary transmission of with weekly testing either and high mitigation (0.09 to 0.45) or medium mitigation (0.16 to 0.85) (100) (Figure S9). Similarly a study from Norway with rigorous quarantine and comprehensive testing estimated a 0.9% child-to-child school attack rate and 1.7% child-to-adult attack rate in primary schools. This is most congruent with our model of high mitigation under a 5-day schedule, under which we would expect approximately a 0.9% child-child attack rate and 1.8% child-adult attack rate for full-day contact over the course of infection (101).

Even accounting for stochastic variability, it is nevertheless possible that our results overestimate or underestimate transmission in environments with extremely well-controlled or poorly controlled transmission. For example, Hong Kong reported negligible transmission even in a high schools with very strong mitigation (102). While difficult to interpret due to differences in "close contact" defintions across studies, some data may suggest higher attack rates than were modeled, such as 27% of contacts testing positive in data from Florida (103). (Even accounting for that fact that only 43% of contacts were tested, the implied lower bound of a 12% attack rate exceeds our modeled attack rates, particularly when accounting for reduced transmission from children and asymptomatic individuals and from non-full-day contacts.) Nevertheless with a large range of modeled attack rates and sensitivity analyses, we aim to capture much of the distribution of school-based attack rates.

Second, several data sources showed signs of overdispersion with the possibility of large outbreaks alongside cases without apparent transmission. In the Rhode Island example, one outbreak involved 10 cases among contacts (10 children, four staff members, and one parent); in another study from Australia, 9 cases in early childhood education centers led to no onward transmission while one led to 13 infections (7). If these were indeed caused by a single index case, the level of overdispersion in our model may not capture such extreme outcomes.

Last, teachers were often overrepresented in outbreaks even in well-studied outbreaks, with 16% of staff and 10% of students having antibodies in a Chilean outbreak across multiple school levels (12). In the Australian study, adults comprised 8/18 of secondary cases identified (7).

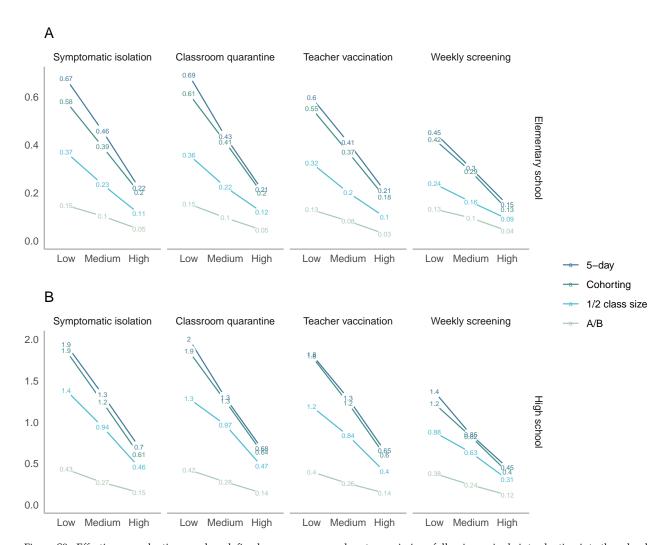


Figure S9: Effective reproduction number, defined as average secondary transmissions following a single introduction into the school (students and staff). The top panel shows elementary schools, where children are assumed to be less susceptible and less infectious, while the bottom panel shows high schools. Note that axes differ across rows. The x-axes vary the level of prevention measure uptake, with low uptake assuming minimal interventions and high uptake assuming intensive interventions. Line colors correspond to scheduling strategies.

FREQUENCY OF INFECTIONS AND SUBCLINICAL INFECTIONS IN CHILDREN COMPARED TO ADULTS

Our model predicted both lower incidence of infections in children and a higher rate of underdiagnosis. With these combined, we would expect to see fewer cases in children, but a smaller relative difference when comprehensive surveillance and/or random testing is conducted, which is consistent with observed data. For example, in passive surveillance from the United Kingdom, staff had more than 4 times the COVID incidence of students per 100,000 across all age groups; however in random testing, observed prevalence was roughly equal among students and staff (104,105). In elementary schools in New York in fall 2020, schools reported substantially fewer cases in elementary school students than in staff per population, but in random surveillance testing from the same period in New York City (manually extracted and analyzed by others), prevalence was roughly equal in students and staff (106,107). While these are difficult to compare directly as people who self-isolate for symptoms are not present for random testing, the contrast remains striking. Less systematically, a major outbreak in an Israeli high school was detected from wide-scale testing after observing 2 unlinked cases (10), and the first Ontario school to participate in voluntary mass asymptomatic screening closed after uncovering a substantial number of previously undetected cases (108). In an analysis of two schools with biweekly or weekly testing, 3% of cases in elementary school-aged children, 25% in middle school-aged children, and 9% in high school-aged children were symptomatic at the time of testing, compared to 48% in adults (100).

EFFECTIVE REPRODUCTION NUMBER

In Figure S10, we display the effective reproduction number associated with different scenarios. One modeling study estimated that from August through October 2020, there was an average effective reproduction number of 0.54 [0.44-0.62] for children 0-9 and 0.75 [0.59-0.89] for children 10-19 (109). As noted in the introduction of this article, school openings varied considerably across the country, but both of these fall within our range of estimates. For elementary schools, it is consistent with full opening and high mitigation, hybrid opening or limited attendance and medium mitigation, or some combination of these with remote school. For high schools, it is most consistent with high mitigation limited attendance or a hybrid model or some combination of these with remote school.

POPULATION-LEVEL STUDIES

While we do not directly model full community incidence, two recent studies using quasi-experimental data to study the impact of school reopening on transmission were consistent with our observation that there is a higher risk of increased community transmission following school reopenings when initial transmission was

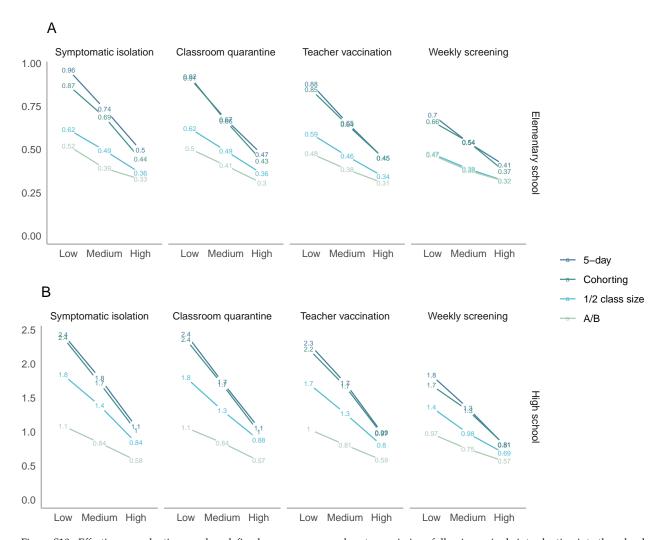


Figure S10: Effective reproduction number, defined as average secondary transmissions following a single introduction into the school community (students, staff, and families). The top panel shows elementary schools, where children are assumed to be less susceptible and less infectious, while the bottom panel shows high schools. Note that axes differ across rows. The x-axes vary the level of prevention measure uptake, with low uptake assuming minimal interventions and high uptake assuming intensive interventions. Line colors correspond to scheduling strategies.

high, contrasted to tight null effects when initial transmission was lower (110,111).

INTERVENTIONS

It is challenging to discern the importance of specific interventions from available data, and we modeled a package of interventions, intended to reflect a combination of masking, distancing, and ventilation. In practice, school districts must discern how to select interventions to reduce transmission while maximizing educational time. For example, space constraints may be a barrier to returning full-time in person if 6′ distancing is required. A recent paper examined the difference between 3′ and 6′ distancing in schools in Massachusetts, finding an IRR of 0.79 in students (95% CI: 0.53 to 1.18) and 0.92 in staff (95% CI: 0.67 to 1.25) after adjusting for demographics and community incidence (112). Applying a non-inferiority interpretation to these confidence intervals, they suggest that we can rule out with high confidence that 6′ distancing has an IRR of less than 0.53 in students and 0.67 in staff compared to 3′ distancing. Similarly, a systematic review found that mask-wearing by non-health care workers can reduce individual-level infection risk by 47%, although specific evaluation of the benefit in school settings is not available (113). Given the harms of lost educational time and the potential for other interventions like high adherence to masking, ventilation, and/or testing to be able to substitute for greater distancing, schools with high adherence to other prevention measures may feel comfortable moving from 6′ to 3′.