1 2 3 4	When can we stop wearing masks? Agent-based modeling to identify when vaccine coverage makes nonpharmaceutical interventions for reducing SARS-CoV-2 infections redundant in indoor gatherings
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11	
12	Abstract: As vaccination efforts to combat the COVID-19 pandemic are ramping up worldwide,
13	there are rising concerns that individuals will begin to eschew nonpharmaceutical interventions
14	for preventing SARS-CoV-2 transmission and attempt to return to pre-pandemic normalcy
15	before vaccine coverage levels effectively mitigate transmission risk. In the U.S.A., some
16	governing bodies have already weakened or repealed guidelines for nonpharmaceutical
17	intervention use, despite a recent spike in national COVID-19 cases and majority population of
18	unvaccinated individuals. Recent modeling suggests that repealing nonpharmaceutical
19	intervention guidelines too early into vaccine rollouts will lead to localized increases in COVID-
20	19 cases, but the magnitude of nonpharmaceutical intervention effects on individual-level SARS-
21	CoV-2 infection risk in fully- and partially-vaccinated populations is unclear. We use a
22	previously-published agent-based model to simulate SARS-CoV-2 transmission in indoor
23	gatherings of varying durations, population densities, and vaccination coverage levels. By

simulating nonpharmaceutical interventions in some gatherings but not others, we were able to

quantify the difference in SARS-CoV-2 infection risk when nonpharmaceutical interventions

24

26	were used, relative to scenarios with no nonpharmaceutical interventions. We found that
27	nonpharmaceutical interventions will often reduce secondary attack rates, especially during brief
28	interactions, and therefore there is no definitive vaccination coverage level that makes
29	nonpharmaceutical interventions completely redundant. However, the reduction effect on
30	absolute SARS-CoV-2 infection risk conferred by nonpharmaceutical interventions is likely
31	proportional to COVID-19 prevalence. Therefore, if COVID-19 prevalence decreases in the
32	future, nonpharmaceutical interventions will likely still confer protective effects but potential
33	benefits may be small enough to remain within "effectively negligible" risk thresholds.
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- 35 Keywords: agent-based model, COVID-19, indoor transmission, nonpharmaceutical
- 36 interventions, SARS-CoV-2, vaccine

37 Introduction

38 Global vaccine rollout to combat the Coronavirus Disease 2019 (COVID-19) pandemic is 39 well underway, with at least different seven vaccines approved for distribution by different 40 countries (WHO 2021). In the U.S.A., where three vaccines have been approved for distribution 41 (CDC 2021a), 24.8% of the population has been fully vaccinated against COVID-19 as of April 42 17th 2021 (CDC 2021b). Despite ongoing vaccine rollouts, as of April 17th 2021, there is an 43 indication that COVID-19 cases are surging in some U.S. states (NY Times 2021). In spite of 44 rising case numbers, several U.S. states have recently rescinded, or allowed to expire, policies 45 mandating use of nonpharmaceutical intervention in public spaces, with seemingly no intention 46 of reinstating them in the near future (State of Iowa 2021; State of Mississippi 2021; State of 47 Texas 2021). Population-level epidemiological models of vaccine rollout effects on COVID-19 48 transmission suggest that discontinuing nonpharmaceutical intervention use early into the 49 vaccination effort leads to a subsequent surge in COVID-19 cases and related hospitalizations 50 and deaths (Gozzi et al. 2021; Moore et al. 2021). 51 The magnitude of nonpharmaceutical intervention effects on individual-level SARS-52 CoV-2 infection risk in fully- and partially-vaccinated populations is unclear. This information is 53 crucial for identifying vaccination levels at which it would be appropriate to scale-back 54 guidelines for nonpharmaceutical interventions, as it would allow governing bodies to base 55 policies on concrete risk estimates. The United States Centers for Disease Control and 56 Prevention (CDC) has updated guidelines on safe gathering protocols, recommending that groups 57 of fully-vaccinated people can now safely interact amongst themselves, or with small groups of 58 unvaccinated people at low risk for developing severe COVID-19, without utilizing any 59 nonpharmaceutical Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2)

60 transmission interventions (e.g., face coverings, 2-m social distancing, etc.) (CDC 2021c). 61 However, the guidelines also recommend to continue avoiding medium to large gatherings, and 62 the use of nonpharmaceutical interventions in public and when gathering with unvaccinated 63 individuals. This caution stems from the incomplete knowledge of vaccine effectiveness across 64 different populations, their effects on transmission, and the potential change on vaccine 65 effectiveness caused by the emergence of new SARS-Cov-2 variants. The problem with citing vaccination efforts as a justification for discontinuing 66 nonpharmaceutical interventions is twofold. First and foremost, the majority of the U.S. 67 68 population is not yet fully vaccinated (CDC 2021b), and therefore presumably has little-to-no 69 immunity from SARS-CoV-2 infections. Secondly, while there is growing evidence that these 70 vaccines reduce SARS-CoV-2 infection risk in addition to COVID-19 incidence, vaccines may 71 not confer complete immunity or block transmission (Hall et al. 2021; Lipsitch & Kahn 2021; 72 Yellen et al. 2021). Data suggest that the BNT162b2 mRNA vaccine (i.e., the vaccine developed 73 by Pfizer-BioNtech) may be \approx 72% effective at preventing laboratory-confirmed SARS-CoV-2 74 infections after a single dose, and $\approx 86-92\%$ two weeks following the second dose (Hall et al. 75 2021; Yellen et al. 2021). Furthermore, this vaccine may reduce viral loads, a potential proxy for 76 infectiousness, in infected individuals by 3-4 times (Levine-Tiefenbrun et al. 2021). Less 77 information is available on the ability of the other two vaccines approved for U.S. distribution to 78 reduce SARS-CoV-2 infections, but Lipsitch & Kahn (2021) do estimate that mRNA-1273 (i.e., 79 the vaccine developed by Moderna and NIAID) can reduce individual-level infection risk by at 80 least 61% following the first dose. Despite potentially-high infection-reduction efficacies, 81 without vaccines that confer complete immunity from infection or prevent transmission from 82 infectious individuals, it will be difficult to halt SARS-CoV-2 circulation in the population

83	through vaccination efforts alone (Gozzi et al. 2021; Moore et al. 2021). Considering that most
84	people also have yet to be fully vaccinated, guidelines that advocate phasing out
85	nonpharmaceutical interventions during interpersonal interactions may be premature at this time.
86	In Farthing & Lanzas (2021), we described an agent-based model (ABM) for simulating
87	indoor respiratory pathogen transmission. We previously used this model to quantify effects of
88	nonpharmaceutical interventions on reducing SARS-CoV-2 transmission risk during an indoor
89	superspreading event (Farthing & Lanzas 2021). Here, we use it to simulate SARS-CoV-2
90	transmission in indoor gatherings of varying durations, population densities, and proportional
91	vaccination coverage. By simulating nonpharmaceutical interventions in some gatherings but not
92	others, we were able to quantify the difference in SARS-CoV-2 infection risk when
93	nonpharmaceutical interventions were used in conjunction with vaccination efforts, relative to
94	scenarios with no nonpharmaceutical interventions. Using these data, we demonstrate how
95	interested parties can easily estimate the potential reduction in SARS-CoV-2 infection risk
96	attributable to nonpharmaceutical interventions, and try to answer the question: "at what point
97	during vaccine rollout are gatherings without non-pharmaceutical measures safe?"
98	
99	Methods
100	We used the ABM we first described in Farthing & Lanzas (2021) to simulate the effect
101	of increasing vaccination coverage and nonpharmaceutical interventions on SARS-CoV-2
102	transmission risk during indoor gatherings. The simulation input levels and parameter values we
103	used are given in Table 1. We made the assumptions that any infectious individuals at gatherings
104	would be asymptomatic because symptomatic people would consciously decide to stay away,

105 and that no one with partial immunity exists within the group of attendees. Vaccinated people

had a fixed probability of becoming completely immune to SARS-CoV-2 infection (Table 1),
and those that did not become immune remained susceptible to infection (i.e, 'all-or-nothing'
vaccine). Finally, we only simulated use of cloth face coverings, rather than notably moreeffective masks like N95s, because we make the assumption that the majority of Americans have
ready access to, and are more-likely to use cloth masks.
All simulations were carried out within the open-source modeling software, NetLogo

112 (Ver. 6. 1. 1 – Wilensky 1999). We executed a factorial simulation run in the NetLogo 113 BehaviorSpace using our specified input levels, and ran 200 simulations replicates of each 114 parameter set combination when the nonpharmaceutical interventions were included and when 115 they were not. We ran these factorial combination sets separately in order to save computation 116 time as there were two inputs (i.e., mask efficacy, attempted social distance) that only changed 117 when nonpharmaceutical interventions were simulated. We ultimately produced 1,612,800 118 simulations without nonpharmaceutical interventions, and 9,676,800 including them (i.e., 119 11,289,600 total simulations). We recorded the number of susceptible individuals infected in 120 each simulation, and aggregated this information into a single data set prior to analysis.

121 We reported the mean probability of observing ≥ 1 successful infection event(s) and 122 mean secondary attack rates in indoor gatherings when an asymptomatic person was also in 123 attendance across factorial combinations of "between-group comparison" variables (Table 1). 124 Secondary attack rates here were calculated by dividing the number of people that were infected 125 at the gathering by the number of "healthy" people at the start of the gathering, and can also be 126 considered to be the individual-level probability of a previously healthy attendee being infected 127 at the gathering. To assess the difference between protection conferred by the simultaneous 128 deployment of pharmaceutical and nonpharmaceutical interventions, versus use of only

129 nonpharmaceutical interventions, we first smoothed the observed mean secondary attack rates 130 (μ) by fitting them to a beta regression model with a fixed unknown precision parameter, ϕ using 131 a logit link function to map (0,1) values (Ferrari & Cribari-Neto 2004). The specific model is 132 given by:

133
$$ln\left(\frac{\mu}{1-\mu}\right)(\phi) = (\phi)\,\beta_0 + \beta_1(Gathering\ duration) + \beta_2(Intervention\ level) +$$

134
$$\beta_3(Vaccine\ coverage) + \beta_4(Vaccine\ efficacy) + \beta_5(Vaccine\ coverage\ *\ Vaccine\ efficacy),$$

136 where "Intervention level" is a categorical variable containing the following mutually-exclusive 137 levels: "cloth face masks & vaccination," "cloth face masks & 2-m social distancing & vaccination," and "vaccination only." Additionally, "Vaccine efficacy" here refers to the ability 138 139 of vaccines to induce complete immunity to infection. "Vaccine coverage" and "Vaccine 140 efficacy" are given in terms of decimal percent, not percentage points (e.g., 0.1, not 10%). 141 Because beta regression models assume all dependent variable values fall between 0 and 1, we 142 used the data transformation procedure described by (Cribari-Neto & Zeiles, 2010) to reconstruct 143 our proportion data without these extremities prior to model fitting. We used the pseudo- R^2 144 calculation procedure given by Ferrari & Cribari-Neto (2004) to assess the goodness of fit for 145 our regression model.

After fitting our data, we used the regression model to predict the mean secondary attack rates during a 60-minute gathering with a single asymptomatic person in attendance across the complete factorial combination of covariate inputs described in Table 2. We report the difference between predicted values when all interventions (i.e., cloth face masks & 2-m social distancing & vaccination) are utilized, and predicted values assuming vaccinations are the only interventions. All analyses and plotting were carried out using functions from the "betareg"

152 (Ferrari & Cribari-Neto 2004) and "ggplot2" (v. 3.3.2, Wickham 2016) R packages, respectively,

153 in RStudio (v. 1.1.463, RStudio Team, Boston, MA) (RStudio Team 2018) running R (v. 3.6.2, R

- 154 Foundation for Statistical Computing, Vienna, Austria) (R Core Team 2020).
- 155

156 **Results & Discussion**

157 We found that the probability of ≥ 1 successful transmission event generally increased 158 with population density (Fig. 1). This is unsurprising, as SARS-CoV-2 transmission in this ABM 159 is highly sensitive to within-room population density (Farthing & Lanzas 2021). We observed 160 that at low population densities and/or short-duration gatherings, the use of nonpharmaceutical 161 interventions can significantly reduce the probability of successful transmission. Furthermore, it 162 is clear that at low population densities, 2-m social distancing confers additional protective 163 effects when used in conjunction with cloth face coverings, even during relatively-long duration 164 gatherings. This is consistent with what we observed when we used the same ABM to directly compare the effectiveness of varied nonpharmaceutical interventions to prevent SARS-CoV-2 165 166 transmission during a superspreading event (Farthing & Lanzas 2021). We found that cloth face 167 masks alone conferred few protective effects in long-duration gatherings.

The probability of transmission events occurring was unlikely to reach $\approx 0\%$ outside of scenarios with low population density and multiple nonpharmaceutical interventions, or $\geq 95\%$ vaccine coverage and vaccines that were 100% effective at preventing infections. Given that 1) current estimates place SARS-CoV-2 vaccine efficacies against infection between 60-90% (Hall et al. 2021; Lipsitch & Kahn 2021; Yellen et al. 2021), 2) historical precedence suggesting adult populations will fall well short of these high vaccination levels (Applewhite et al. 2020; CDC 2020), and 3) the difficulty government institutions have had enforcing nonpharmaceutical

intervention policies (Jacobs & Ohinmaa 2020; Pedersen & Favero 2020), it is unlikely that these
scenarios will be representative of average real-world gatherings. Moreover, in 60-min gathering
scenarios, the probability of ≥ 1 successful transmission event occurring is relatively high even
when gathering attendees utilize nonpharmaceutical interventions and most are vaccinated.
The probability that ≥1 SARS-CoV-2-positive individual is in attendance at a gathering

180 can be calculated as

181

$$1 - (1 - p)^n$$
, (2)

where p is the local COVID-19 prevalence, and n is the number of people at the gathering
(Chande et al. 2020). The prevalence of infectious cases (p) can be highly uncertain because of
the variable testing effort across time and space, but it can be estimated by assuming that any
SARS-CoV-2-positive individuals are infectious at time of testing and will remain infectious for
a given period of time. Additionally, ascertainment bias can be factored in. The probability that a
given individual will be infected at a gathering is then

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$$(1 - (1 - p)^n)q_i,$$
 (3)

where q_i is the probability that individual *i* will be infected given exposure to an asymptomatic 189 190 individual at the gathering. Effectively, what we report in Fig. 2 are estimates of q_i under 191 different circumstances. Our findings suggest that cloth-based mask use, with or without 2-m 192 social distancing, often does not confer significant protective effects during long-duration 193 gatherings (Fig. 2), we have also shown that implementing these nonpharmaceutical 194 interventions can reduce overall transmission probability (Fig. 1) and secondary attack rates (Fig. 195 2, Table 3) during brief interactions or gatherings with relatively-few people (e.g., fewer than 10 196 people, the limit for indoor and/or outdoor social gatherings enforced by some U.S. states 197 (MultiState 2021)). This effectively means that strict guidelines for continued nonpharmaceutical

198 intervention use will likely help to mitigate SARS-CoV-2 spread, and therefore COVID-19 199 incidence, for as long as these policies are in effect.

As vaccine coverage increases, the question now becomes "how much elevated risk is 200 acceptable in the absence of nonpharmaceutical interventions?" If we let q'_i denote the 201 202 probability that individual *i* will be infected given exposure to an asymptomatic individual at a gathering where no nonpharmaceutical interventions were in place, and q_i^* denote the probability 203 204 that individual *i* will be infected given exposure to an asymptomatic individual at a gathering 205 where some level of nonpharmaceutical interventions were in place, then the relative effect of 206 nonpharmaceutical interventions on reducing infection risk is equal to 207

7
$$\frac{q_i}{q'_i} * 100\%.$$
 (4)

208 By quantifying covariate effects in our beta-regression model, we provide interested parties with a formula that can be used to quickly determine generalized q'_i or q^*_i values, without 209 210 the need for running a large number of simulations. Due to the logit link function we used, the 211 mean secondary attack rates in our ABM simulations (μ) can be predicted using the equation 212

 $e^{\beta_0+\beta_1}(Gathering duration)+\beta_2(Intervention level)+\beta_3(Vaccine coverage)+\beta_4(Vaccine efficacy)+\beta_5(Vaccine coverage*Vaccine efficacy)$ 213 $\mu = \frac{\sigma}{1 + \rho \beta_0 + \beta_1 (\text{Gathering duration}) + \beta_2 (\text{Intervention level}) + \beta_3 (\text{Vaccine coverage}) + \beta_4 (\text{Vaccine efficacy}) + \beta_5 (\text{Vaccine coverage} + \text{Vaccine efficacy}) + \beta_5 (\text{Vaccine coverage} + \beta_4 (\text{Vaccine efficacy}) + \beta_5 (\text{Vaccine eff$ 214 (5)

(Ferrari & Cribari-Neto 2004). Our regression model had a pseudo-R² of 0.37. Given the number 215 216 of stochastic processes in our ABM and the variability purposely introduced into simulations 217 (Table 1), we believe the explanatory power of the model is acceptable for our purposes here. 218 Assuming mean population-level vaccine efficacies of 60% and 80%, which we believe are 219 conservative estimates for U.S.-approved vaccine efficacies, our regression model consistently

220 predicts that secondary attack rates decrease by 55-58% when attendees utilize cloth masks and 221 2-m social distancing, regardless of gathering duration (Fig. 3). However, it is important to 222 reiterate that here we estimate the probability or infection given contact with an infectious 223 individual at a gathering (q_i) and comment on the relative risk difference attributable to 224 intervention use. This should not be confused with the absolute risk of becoming infected at a 225 gathering (see Equation 3). We demonstrate the difference in Figure 4, which is a simplistic 226 example intended to show that even at relatively high COVID-19 prevalence levels, 20 people 227 gathering indoors for 60 minutes have a substantially-lower individual-level risk of SARS-CoV-228 2 infections than is suggested by q_i alone. Though predicting intervention effects on community-229 level COVID-19 prevalence and infection-related events (e.g., symptom-onset, mortality, or hospitalization) is outside the scope of our model, our simulations do suggest that secondary 230 231 attack rates are negatively correlated with vaccine coverage. Given that we expect local COVID-232 19 prevalence to eventually follow similar trends (Gozzi et al. 2021), the relative impact of 233 nonpharmaceutical interventions on infection risk reduction will likely decrease over time as 234 vaccine rollouts continue.

235 In addition to being unable to comment on community-level infection metrics, there are a 236 few other limitations associated with our results that we must acknowledge. Aside from the 237 ABM design limitations outlined in Farthing et al. (2021), we make a number of assumptions in 238 our simulations. Most of these assumptions are directly tied to our parameter space detailed in 239 Table 1, and include such things as: in simulated gatherings only one asymptomatic individual 240 was in attendance, no individuals wear masks with exposure-reduction efficacies > 50% and 241 therefore we are not simulating the use of N95 or similar masks, and there is no simulated 242 forced-air ventilation or infectious individuals that produce superspreader-level of contaminated

243	aerosols (e.g., 970 quanta (Miller et al. 2020)). Additionally, we do not simulate activity-specific
244	behaviors and individuals in our simulations were unmoving. Finally, we based the
245	infectiousness of asymptomatic individuals on the estimate given by Buonanno et al. (2020) (i.e.,
246	142 quanta/hr), and to relate this estimate to ABM parameters we used the linear model
247	described in Farthing et al. (2021). However, this parameterization procedure may have over-
248	inflated virion transmissibility in certain scenarios because quanta-estimates are room-size
249	specific, and the Farthing et al. (2021) linear model was based on simulations of gatherings
250	within a relatively large room. In short, our results must be viewed through the lens of simulated
251	world parameters and behaviors, and likely will not wholly reflect all variability that may exist in
252	real-world transmission events. This is very common for ABM-based studies however, and we
253	feel that our model is sufficiently accurate to highlight general trends in indoor SARS-CoV-2
254	transmission and infection risk.
255	
256	Conclusions
257	We found that nonpharmaceutical interventions will often reduce secondary attack rates,
258	especially during brief interactions, and therefore there is no definitive vaccination coverage
259	level that makes nonpharmaceutical interventions completely redundant. However, the beneficial
260	effect on absolute SARS-CoV-2 infection risk reduction conferred by nonpharmaceutical
261	interventions used during indoor gatherings is likely proportional to COVID-19 prevalence.
262	Therefore, if U.S. COVID-19 prevalence decreases in the future, nonpharmaceutical
263	interventions will likely still confer protective effects, but any potential benefits may be small

enough to remain within "effectively negligible" risk thresholds.

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268	
269	Author Contributions
270	Trevor Farthing led the model creation, data analysis, and manuscript writing, but both
271	authors conceived the ideas presented herein, contributed to model development and writing
272	efforts, and gave final approval for publication. Cristina Lanzas secured the funding.
273	
274	Data availability
275	We first made our ABM publicly available for download in Farthing et al. (2021). The
276	current iteration can be downloaded from the Lanzas lab's github repository at
277	https://github.com/lanzaslab/droplet-ABM.
278	
279	References
280	1. Adams WC. Measurement of breathing rate and volume in routinely performed daily
281	activities. 1993. Final Report, Contract No. A033-205. California Air Resources Board,
282	Sacramento, CA, USA.
283	https://ww2.arb.ca.gov/sites/default/files/classic//research/apr/past/a033-205.pdf.
284	2. Applewhite A, Stancampiano FF, Harris DM, Manaois A, Dimuna J, Glenn J, Heckman MG
285	Brushaber DE, Sher T, & Valery JR. (2020). A retrospective analysis of gender-based
286	difference in adherence to influenza vaccination during the 2018-2019 season. J Prim Care
287	& Commun Heal 11:1-6. https://doi.org/10.1177%2F2150132720958532.

288 3. Buonanno G, Stabile L, & Morawska L. (2020). Estimation of airborne v	viral	emission:
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289 quanta emission rate of SARS-CoV-2 for infection risk assessment. *Environ Internat*

290 141:105794. https://doi.org/10.1016/j.envint.2020.105794.

- 4. Castillo JE, & Weibel JA. (2018). A point sink superposition method for predicting droplet
- interaction effects during vapor-diffusion-driven dropwise condensation in humid air. Int J
- 293 *Heat Mass Trans* 118:708-719. https://doi.org/10.1016/j.ijheatmasstransfer.2017.11.045.
- 5. Chande A, Lee S, Harris M, Nguyen Q, Beckett SJ, Hilley T, Andris C, & Weitz JS. (2020).
- 295 Real-time, interactive website for US-county-level COVID-19 event risk assessment. Nat
- 296 Hum Behav. 4:1313-1319. https://doi.org/10.1038/s41562-020-01000-9.
- 297 6. Cribari-Neto F, Zeileis A. (2010). Beta regression in R. J Stat Soft. 34(2):1-24.
- 298 https://doi.org/10.18637/jss.v034.i02.
- 299 7. Das SK, Alam J, Plumari S, Greco V. (2020) Transmission of airborne virus through sneezed
 300 and coughed droplets. Phys Fluids. 32:097102. https://doi.org/10.1063/5.0022859.
- 301 8. van Doremalen N, Bushmaker T, Morris DH, Holbrook MG, Gamble A, Williamson BN,
- 302 Tamin A, Harcourt JL, Thornburg NJ, Gerber SI, Lloyd-Smith JP, de Wit E, & Munster VJ.
- 303 (2020). Aerosol and surface stability of SARS-CoV-2 as compared with SARS-CoV-1. N
- 304 *Engl J Med* 2020(382):1564-1567. https://doi.org/10.1056/NEJMc2004973.
- 305 9. Farthing TS, & Lanzas C. (2021). Assessing the efficacy of interventions to control indoor
- 306 SARS-Cov-2 transmission: an agent-based modeling approach. Preprint available at:
- 307 https://doi.org/10.1101/2021.01.21.21250240.
- 308 10. Ferrari SLP, Cribari-Neto F. (2004). Beta regression for modelling rates and proportions. J
- 309 Appl Stat. 31(7): 799-815. https://doi.org/10.1080/0266476042000214501.

- 310 11. Fryar CD, Kruszon-Moran D, Gu Q, & Ogden CL. (2018). Mean body weight, height, waist
- 311 circumference, and body mass index among adults: United States, 1999–2000 through 2015–
- 312 2016. National Health Statistics Reports, No. 122. United States National Center for Health
- 313 Statistics, Hyattsville, MD, USA. https://stacks.cdc.gov/view/cdc/61430.
- 314 12. Gozzi N, Bajardi P, & Perra N. (2021). The importance of non-pharmaceutical interventions
- 315 during the COVID-19 vaccine rollout. Preprint available at:
- 316 ttps://doi.org/10.1101/2021.01.09.21249480.
- 317 13. Hall VJ, Foulkes S, Saei A, Andrews N, Oguti B, Charlett A, Wellington E, Stowe J, Gillson
- 318 N, Atti A, Islam J, Karagiannis I, Munro K, Khawam J, The Siren Study Group, Chand MA,
- Brown C, Ramsey ME, Bernal JL, & Hopkins S. (2021). Effectiveness of BNT162b2 mRNA
- 320 vaccine against infection and COVID-19 vaccine coverage in healthcare workers in England,
- 321 multicentre prospective cohort study (the SIREN Study). Preprint available at:
- 322 https://dx.doi.org/10.2139/ssrn.3790399.
- 323 14. Jacobs P, & Ohinmaa AP. (2020). The enforcement of statewide mask wearing mandates to
- 324 prevent COVID-19 in the US: an overview. *F1000Res* 9:1100.
- 325 https://dx.doi.org/10.12688%2Ff1000research.25907.1.
- 326 15. Kwon S-B, Park J, Jang J, Cho Y, Park D-S, Kim C, Bae G-N, & Jang A. (2012). Study on
- 327 the initial velocity distribution of exhaled air from coughing and speaking. *Chemosphere*
- 328 87(11):1260-1264. https://dx.doi.org/10.1016/j.chemosphere.2012.01.032.
- 329 16. Levine-Tiefenbrun M, Yelin I, Katz R, Herzel E, Golan Z, Scheiber L, Wolf T, Nadler V,
- 330 Ben-Tov A, Kuint J, Gazit S, Patalon T, Chodick G, & Kishony R. (2021). Decreased SARS-
- 331 CoV-2 viral load following vaccination. Preprint available at:
- 332 https://doi.org/10.1101/2021.02.06.21251283.

- 333 17. Lipsitch M, & Kahn R. (2021). Interpreting vaccine efficacy trial results for infection and
- transmission. Preprint available at: https://doi.org/10.1101/2021.02.25.21252415.
- 18. Miller SL, Nazaroff WW, Jimenez JL, Boerstra A, Buonanno G, Dancer SJ, Kurnitski J,
- 336 Marr LC, Morawska L, & Noakes C. (2020). Transmission of SARS-CoV-2 by inhalation of
- respiratory aerosol in the Skagit Valley Chorale superspreading event. *Indoor Air* 00:1-10.
- 338 https://doi.org/1010.1111/ina.12751.
- 339 19. Moore S, Hill EM, Tildesley MJ, Dyson L, & Keeling M. (2021). Vaccination and non-
- 340 pharmaceutical interventions: when can the UK relax about COVID-19. Preprint available at:
- 341 https://doi.org/10.1101/2020.12.27.20248896.
- 342 20. MultiState. (2021). COVID-19 State and Local Policy Dashboard.
- 343 https://www.multistate.us/research/covid/public. [cited 2021 Apr 14].
- 344 21. O'kelly E, Pirog S, Ward J, Clarkson PJ. (2020). Ability of fabric face mask materials to
- filter ultrafine particles at coughing velocity. *BMJ Open* 10(9):e039424.
- 346 http://dx.doi.org/10.1136/bmjopen-2020-039424.
- 347 22. Pedersen MJ, & Favero N. (2020). Social distancing during the COVID-19 pandemic: who
- 348 are the present and future noncompliers. *Public Admin Rev* 80(5):805-814.
- 349 https://doi.org/10.1111/puar.13240.
- 350 23. R Core Team. (2020). R: A language and environment for statistical computing. R
- Foundation for Statistical Computing, Vienna, Austria. https://www.R_project.org. [cited
 2021 Mar 26].
- 353 24. RStudio Team. (2018). RStudio: integrated development Environment for r. RStudio Team,
- Boston, Massachusetts, USA. http://www.rstudio.com. [cited 2021 Mar 2].

- 355 25. State of Iowa. (2021). Public health proclamation -2021.02.05.
- 356 https://governor.iowa.gov/sites/default/files/documents/Public%20Health%20Proclamation%
- 357 20-%202021.02.05.pdf. [cited 2021 Mar 26].
- 358 26. State of Mississippi. (2021). Executive order No. 1549.
- 359 https://www.sos.ms.gov/content/executiveorders/ExecutiveOrders/1549.pdf [cited 2021 Mar
- 360 26].
- 361 27. State of Texas. (2021). Executive order GA 34.
- 362 https://open.texas.gov/uploads/files/organization/opentexas/EO-GA-34-opening-Texas-
- 363 response-to-COVID-disaster-IMAGE-03-02-2021.pdf [cited 2021 Mar 26].
- 364 28. The New York Times [NY Times]. (2021). Coronavirus in the U.S.: latest map and case
- 365 count. https://www.nytimes.com/interactive/2020/us/coronavirus-us-cases.html#states. [cited
 366 2021 Apr 17].
- 367 29. United States Centers for Disease Control and Prevention [CDC]. (2020). Flu vaccination

368 coverage, United States, 2019-20 influenza season.

- 369 https://www.cdc.gov/flu/fluvaxview/coverage-1920estimates.htm. [cited 2021 Mar 26].
- 370 30. United States Centers for Disease Control and Prevention [CDC]. (2021a). Different
- 371 vaccines. https://www.cdc.gov/coronavirus/2019-ncov/vaccines/different-vaccines.html.
- 372 [cited 2021 Mar 26].
- 373 31. United States Centers for Disease Control and Prevention [CDC]. (2021b). COVID data
- 374 tracker. https://covid.cdc.gov/covid-data-tracker/#vaccinations. [cited 2021 Apr 17].
- 375 32. United States Centers for Disease Control and Prevention [CDC]. (2021c). CDC issues first
- 376 set of guidelines on how fully vaccinated people can visit safely with others.

- 377 https://www.cdc.gov/media/releases/2021/p0308-vaccinated-guidelines.html. [cited 2021
- 378 Mar 26].
- 379 33. Wickham, H. (2016). ggplot2: elegant graphics for data analysis. Springer-Verlag, New
- 380 York, USA. https://ggplot2-book.org/. [cited 2021 Mar 26].
- 381 34. Wilensky U. NetLogo. (1999). Center for Connected Learning and Computer-Based
- 382 Modeling, Northwestern University, Evanston, IL. http://ccl.northwestern.edu/netlogo/.
- 383 [cited 2021 Mar 26].
- 384 35. Wölfel R, Corman VM, Guggemos W, Sailmaier M, Zange S, Müller MA, Niemeyer D,
- Jones TC, Vollmar P, Rothe C, Hoelscher M, Bleicker T, Brünink S, Schneider J, Ehmann R,
- 386 Zwirglmaier K, Drosten C, & Wendtner C. (2020). Virological assessment hospitalized
- 387 patients with COVID-2019. *Nature* 581:465-469. https://doi.org/10.1038/s41586-020-2196-
- 388 x.
- 389 36. World Health Organization [WHO]. (2021). COVID-19 vaccines.
- 390 https://www.who.int/emergencies/diseases/novel-coronavirus-2019/covid-19-vaccines. [cited

391 2021 Mar 26].

- 392 37. Yelin I, Katz R, Herzel E, Berman-Zilberstein T, Ben-Tov A, Kuint J, Gazit S, Patalon T,
- 393 Chodick G, & Kishony R. (2021). Associations of the BNT162b2 COVID-19 vaccine
- 394 effectiveness with patient age and comorbidities. Preprint available at:
- 395 https://doi.org/10.1101/2021.03.16.21253686.

397 Tables

Parameter/Model Input	Purpose¶	Value(s)	Reference(s)			
Infectiousness parameters						
Droplet count (droplets/expectoration) [†]	Fixed value	1.42e ⁵	Buonanno et al. 2020, Farthing & Lanzas 2021			
Droplet spread angle – not	Fixed value	63.5	Kwon et al. 2012			
Droplet travel distance – not	Fixed value	0.55 (0.068) ^{‡§}	Das et al. 2020			
Vaccine-induced infectiousness reduction (%)	Within-group variation	0, 25, 50, 75	Vaccination may reduce infectiousness of asymptomatic individuals by as much as 75%, but effects are unclear (Levine-Tiefenbrun et al. 2021)			
Scenario environment and individ	2021).					
Area (m ²)*	Within-group variation	36, 81, 225	_			
Expectoration height (m)	Fixed value	1.7	Fryar et al. 2018			
Inhalation rate (m ³ air/min)	Fixed value	0.023	Adams 1993			
Maximum people in a single 1- m ² patch (people)	Fixed value	2	_			
Number of asymptomatic infectious individuals (people)	Fixed value	1	-			
Scenario virion behavior inputs	Fixed value					
Virion count (virions/mL fluid)	Fixed value	2.35e ⁹	Wölfel et al. 2020			
Virion decay rate (%/min)	Fixed value	1.05	van Doremalen et al. 2020			

Virion infection risk (%/inhaled	Fixed value	6.24	Farthing & Lanzas 2021
virion)			
Scenario airflow inputs			
Diffusion rate (m ³ /min)	Fixed value	1.5e ⁻³	Castillo & Weibel 2018
Forced airflow	Fixed value	off	_
Scenario intervention inputs			
		• Mask use (10% exposure-	
		reduction efficacy), 2m	
		attempted social distancing	
		• Mask use (25% exposure-	
		reduction efficacy), 2m	
		attempted social distancing	
	D	• Mask use (50% exposure-	
	Between-group	reduction efficacy), 2m	
	comparison:	attempted social distancing	Mask use is intended to represent use of <i>cloth</i>
Nonpharmaceutical	intervention	• Mask use (10% exposure-	masks to prevent exposure to infectious media.
intervention scenarios	combinations	reduction efficacy), no attempted	Cloth mask efficacy is highly variable (O'kelly
	Within-group variation: mask efficacy	social distancing	et al. 2020).
		• Mask use (25% exposure-	
		reduction efficacy), no attempted	
		social distancing	
		• Mask use (50% exposure-	
		reduction efficacy), no attempted	
		social distancing	
		No nonpharmaceutical	
		interventions	

Gathering duration (min)	Between-group comparison	10, 60	-
Vaccine efficacy for preventing	Between-group comparison	50, 65, 80, 100	-
infection (%)	Between-group comparison	0:100 by 5	
Population density (people/m ²)**	Between-group comparison	0.17, 0.33, 0.67, 1	_

398
Table 1. Model parameter and scenario-specific input descriptions for transmission simulations.
 399 *All simulated worlds were square-shaped. The Purpose column describes why the parameter or 400 input was included as it relates to analyses. Specifically, "Fixed value" indicates that values are 401 unchanged across all simulations, and are thus irrelevant for analyses. "Between-group 402 comparison" indicates that levels were used in factorial combinations for data aggregation and 403 reporting. "Within-group variation" indicates that different levels were included to increase the 404 variation in simulation results, and by doing so increase model realism. [†]Based on linear 405 modeling described in Appendix S2 of Farthing & Lanzas (2021), this value equates to 142 406 quanta/hr, the average quanta emission rate for asymptomatic people calculated by Buonanno et 407 al. (2020). [‡]Standard deviation is given in parentheses. [§]Das et al. (2020) estimated the average 408 travel distance of a 100-micrometer droplet expelled from a height of 1.7 m at a velocity of 0.5 409 m/s to be 0.55 m. They also found that the majority of 100-µm droplets will fall 0.55-2.35 m 410 away from the expelling individual, depending on initial velocity, but droplets may settle up to 411 3.2 m away very rarely. A random draw of 10,000,000 samples from a log-normal distribution 412 parameterized using 1.7-m and 0.2095-m droplet spread distance mean and standard deviation 413 values, respectively, generated a distribution in line with this finding. The standard deviation we 414 use in simulations for non-coughing expectoration is proportionate to the one used in this random 415 draw. **Instead of specifying a fixed number of individuals in simulations, we scaled the 416 simulated population with world size.

Covariate	Value(s)
Gathering duration	60 min
Intervention level	 cloth face masks & 2-m social distancing & vaccination vaccination only
	• vaccination only
Vaccine coverage	0:1 by 0.1
Vaccine efficacy	0.6, 0.8

Table 2. Covariate values used for prediction in our example.

Coefficient	Estimate	р
Intercept	-3.786 (-3.857, -3.716)	-
ϕ	28.899 (28.336, 29.462)	_
Gathering duration (min)	0.012 (0.011, 0.012)	< 0.001
Intervention level		
Cloth face masks & 2-m social distancing & vaccination*	0 (0, 0)	_
Cloth face masks & vaccination	0.761 (0.737, 0.785)	< 0.001
Vaccination only	0.889 (0.866, 0.913)	< 0.001
Vaccine coverage	0.783 (0.660, 0.905)	< 0.001
Vaccine efficacy	0.385 (0.297, 0.472)	< 0.001
Vaccine coverage X Vaccine efficacy	-2.652 (-2.816, -2.487)	< 0.001

- **Table 3**. Logit scale estimates associated with 1-unit increases in covariate values given by our
- 421 beta-regression model. Wald 95% confidence intervals are given in parentheses. *This is the
- 422 reference level used to establish a baseline for binary dummy variables.

425 Figures



Additional interventions? 🖨 cloth face masks & 2-m social distancing 🖨 cloth face masks only 🖨 no nonpharmaceutical interventions

- 427 Figure 1. At low population densities and gathering duration limits, nonpharmaceutical
- 428 interventions to prevent infection and elevated vaccination rates consistently decrease the
- 429 probability of observing \geq 1 successful SARS-CoV-2 transmission events in simulations.













435 secondary attack rates by 55-58%. This effect was only modeled for vaccine efficacies of 60%436 and 80%.



439 Figure 4. Estimated absolute risk of being infected with SARS-CoV-2 during 60-minute

- 440 gatherings of varied sizes. Estimates were obtained by plugging Figure 3 predictions into
- 441 Equation 3 with fixed COVID-19 prevalence and n values. a) Absolute risk of SARS-CoV-2
- 442 transmission given that 10 people attend the gathering. b) Absolute risk of SARS-CoV-2
- 443 transmission given that 20 people attend the gathering.