

Individually Optimal Choices can be Collectively Disastrous in COVID-19 Disease Control

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1 **Individually optimal choices can be collectively disastrous in COVID-19**
2 **disease control**

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19

20 **Abstract:**

21 **Background:** The word ‘pandemic’ conjures dystopian images of bodies stacked in the streets
22 and societies on the brink of collapse. Despite this frightening picture, denialism and
23 noncompliance with public health measures are common in the historical record, for example
24 during the 1918 Influenza pandemic or the 2015 Ebola epidemic. The unique characteristics of
25 SARS-CoV-2—its high basic reproduction number (R_0), time-limited natural immunity and
26 considerable potential for asymptomatic spread—exacerbate the public health repercussions of
27 noncompliance with interventions (such as vaccines and masks) to limit disease transmission.
28 Our work explores the rationality and impact of noncompliance with COVID-19 disease control
29 measures.

30 **Methods:** In this work, we used game theory to explore when noncompliance confers a
31 perceived benefit to individuals. We then used epidemiological modeling to predict the impact
32 of noncompliance on control of COVID-19, demonstrating that the presence of a noncompliant
33 subpopulation prevents suppression of disease spread.

34 **Results:** Our modeling demonstrating that noncompliance is a Nash equilibrium under a broad
35 set of conditions, and that the existence of a noncompliant population can result in extensive
36 endemic disease in the long-term after a return to pre-pandemic social and economic activity.
37 Endemic disease poses a threat for both compliant and noncompliant individuals; all
38 community members are protected if complete suppression is achieved, which is only possible
39 with a high degree of compliance. For interventions that are highly effective at preventing
40 disease spread, however, the consequences of noncompliance are borne disproportionately by
41 noncompliant individuals.

42 **Conclusions:** In sum, our work demonstrates the limits of free-market approaches to
43 compliance with disease control measures during a pandemic. The act of noncompliance with
44 disease intervention measures creates a negative externality, rendering COVID-19 disease
45 control ineffective in the short term and making complete suppression impossible in the long
46 term. Our work underscores the importance of developing effective strategies for prophylaxis
47 through public health measures aimed at complete suppression and the need to focus on
48 compliance at a population level.

49

50 **Keywords**

51 COVID-19, game theory, noncompliance, infectious disease control, epidemiological modeling

52

53 **Background**

54 As we enter the unfamiliar territory of the worst global pandemic in a century, the worldwide
55 emergence of noncompliance with public health measures aimed at limiting the spread of
56 COVID-19 is not as surprising as it may seem at first blush [1, 2]. During the 1918 Influenza
57 pandemic, for example, resistance to public health measures aimed at reducing the spread of
58 disease manifested at the individual level, leading to violence [3] and stiff punishments for
59 “mask slackers” [4, 5]. Anti-mask protesters led large demonstrations [6], and city councils
60 questioned the value of mask ordinances [7, 8] with emotionally charged language: “under no
61 circumstances will I be muzzled like a hydrophobic dog” [9]. The phrasing may be dated, but the
62 sentiment echoes precisely across a century [10].

63

64 For COVID-19, a number of features of the disease facilitate noncompliance with disease
65 control measures such as masking and vaccination. Hospitalization and death happen away
66 from the public eye, and our changing understanding of the mechanism of transmission, the
67 risk of mortality and the long-term consequences of the disease have favored the spread of
68 misinformation. The spread of confusion and misinformation has been a common feature for
69 other novel pathogen-induced pandemics such as Ebola [1, 11, 12] and the 1918 Flu [13]. While
70 the existence of pandemic denialism was easy to anticipate [14], the unique characteristics of
71 SARS-CoV-2 amplify its effect. Studies suggest that asymptomatic or presymptomatic patients
72 account for up to 40% of SARS-CoV-2 transmission [15], severely limiting the utility of more
73 traditional and intuitive disease control measures such as symptomatic isolation [16]. The high
74 R_0 of SARS-CoV-2 (reported to be 5.7 in the early days of the pandemic in Wuhan [17]) creates
75 the potential for explosive growth in situations where the virus has not been completely
76 eradicated, as has been demonstrated by a massive second wave in many European countries
77 [18, 19]. Making matters worse, estimates for natural immunity as a consequence of SARS-CoV-
78 2 infection range from six to twenty-four months [20–22], creating the potential for multiple
79 waves of disease in the short term.

80

81 Thus, the unique characteristics of SARS-CoV-2 raise the possibility that noncompliance with
82 public health measures may create conditions that make disease control in the short term
83 impossible or prevent any return to pre-pandemic lifestyles in the long run. With this in mind,
84 we asked three questions: First, in the specific case of SARS-CoV-2, are there circumstances that
85 lead to a perceived benefit to noncompliance with public health measures for a substantial

86 portion of the population? Second, what is the impact of noncompliance on the attainability of
87 complete disease suppression for SARS-CoV-2? Third, what is the magnitude of the negative
88 externality (a cost incurred by them that is not of their choosing) created for the compliant
89 population as a result of noncompliance of others?

90

91 We approached the first question from the perspective of game theory, which has previously
92 been applied to decision-making around vaccine uptake [23]. A number of studies have
93 examined noncompliance with measures to control COVID-19 through a social-sciences lens,
94 exploring social and psychological risk factors associated with this behavior. These studies, from
95 a range of different countries, have linked noncompliance to Dark Triad traits (i.e.,
96 Machiavellianism, Psychopathy Factor 1, and narcissistic rivalry [24]), antisocial behaviors [25],
97 higher levels of impulsivity [26] and a prior record of delinquent behaviors [27]. A positive,
98 rather than normative, framing of the question involves exploring the set of conditions for
99 which the perceived benefit of noncompliance to the individual is simply greater than the
100 perceived benefit of compliance. This allows us to examine the problem of compliance from the
101 limited perspective of individuals optimizing for their own benefit without accounting for the
102 common good, particularly relevant in the context of arguments based on personal liberty
103 being used as a justification for noncompliance [28].

104

105 For the next two questions, we used a Susceptible-Exposed-Infected-Recovered-Susceptible
106 (SEIRS) epidemiological modeling framework with a duration of immunity ranging from six to
107 twenty-four months to explore the range of levels of compliance and intervention efficacy

108 required for disease suppression. Our intent in this study was to establish a link between the
109 free optimization of individuals' outcomes as a result of noncompliance, the externalities
110 generated by those choices, and the implications for epidemic control in the short and long
111 term.

112

113 **Methods**

114 *Game Theory Modeling of Compliance with COVID-19 Interventions*

115 For the purposes of this work, we defined an "intervention" as being a public health measure
116 that reduces the transmission of COVID-19. This may be a nonpharmaceutical intervention,
117 such as masks, or a biomedical intervention, such as a vaccine. Compliance with an intervention
118 is defined as a binary choice. An individual can choose whether or not to comply with an
119 intervention based on the perceived costs and benefits of the intervention. We modeled this
120 choice using a game theoretic framework, which compares the perceived cost of compliance
121 (reduction of quality of life resulting from the intervention) in relation to perceived cost of
122 infection (risk-weighted morbidity/mortality burden) to the individual. Individuals derive a
123 benefit or cost (i.e., a payoff) from interactions with other individuals in the population, who
124 can also either be compliers or noncompliers.

125

126 We sought to determine the conditions under which noncompliance is the Nash equilibrium, or
127 optimal behavior strategy for individuals seeking to maximize their own payoff. In a Nash
128 equilibrium, the expected payoff to noncompliers is higher than the payoff to compliers when
129 interacting with any other individual in the population [29].

130

131 For this two-strategy “game”, the payoffs to compliers and noncompliers are given in Table 1,
132 where q is the cost of the intervention, α_i is the fraction of infected individuals of type i , and m_i
133 is the perceived cost of infection for type i individuals, where i can either be u (noncompliers) or
134 v (compliers). The cost m_i is the perceived risk of a negative health outcome given exposure to
135 an infected individual. Other parameter definitions are given in Table 2. As in the SEIRS model,
136 the efficacy of the intervention in protecting the individual from getting infected (b) is equal to
137 the efficacy in preventing transmission (c) (i.e. $b = c$).

138

139 **Table 1:** Payoff matrix for compliers/noncompliers.

	Noncompliant interaction partner	Compliant interaction partner
Noncomplier payoff	$-\alpha_u m_u$	$-\alpha_v m_v c$
Complier payoff	$-q - \alpha_u m_u b$	$-q - \alpha_v m_v b c$

140 α_i : fraction of infected individuals of type i , m_i : perceived cost of infection for type i individuals,
141 m_i : the perceived risk of a negative health outcome given exposure to an infected individual,
142 where i can either be u (noncompliers) or v (compliers). All other parameter definitions are
143 given in Table 2.

144

145 Noncompliance is a Nash equilibrium if and only if both of the following conditions are met:

146
$$-\alpha_u m_u > -q - \alpha_v m_v b$$

147
$$-\alpha_u m_u c > -q - \alpha_v m_v b.$$

148 Or, equivalently

149

$$\alpha_u < \frac{d}{(m_u - m_v b)}$$

150

$$c\alpha_v < \frac{d}{(m_u - m_v b)}.$$

151

152 Since noncompliers are much more likely to be infected than compliers, $\alpha_u > c\alpha_v$. Therefore,
153 meeting the first condition alone (noncompliers receive a greater payoff than compliers when
154 interacting with other noncompliers) is sufficient for noncompliance to be a Nash equilibrium.

155

156 *SEIRS Model*

157 To support predictions of short- and long-term outcomes for the COVID-19 pandemic, we built
158 an SEIRS ordinary differential equations (ODE) model to account for disease spread, waning
159 immunity in the recovered population, and the acceptance of a vaccine or non-pharmaceutical
160 intervention (NPI) in a fraction of the population. The model consists of two parallel sets of SEIR
161 compartments representing the vaccinated or NPI-compliant (“compliant”) and unvaccinated or
162 NPI-noncompliant (“noncompliant”) populations. The compliant population has a reduced risk
163 of infection which is conferred by the vaccine or NPI (“protective efficacy”). The compliant
164 population may also have a reduced risk of transmission to others upon infection resulting from
165 physiological or behavioral changes (“transmission reduction.”) All compartments were
166 assumed to be well-mixed, meaning that compliant and noncompliant individuals are in contact
167 within and between groups. Vaccination or NPI compliance-based reductions in susceptibility,
168 transmissibility, or contact rate were assumed to be time-invariant, reflecting the most

169 optimistic case for disease control. Similarly, individuals do not move between the compliant
 170 and noncompliant compartments. Model equations are summarized below:

171
$$\frac{dS_v}{dt} = -\beta b S_v (c I_v + I_u) + \delta R_v + f\mu - \lambda S_v$$

172
$$\frac{dE_v}{dt} = -\alpha E_v + \beta b S_v (c I_v + I_u) - \lambda E_v$$

173
$$\frac{dI_v}{dt} = -\gamma I_v + \alpha E_v - \lambda I_v$$

174
$$\frac{dR_v}{dt} = \gamma I_v (1 - \sigma) - \delta R_v - \lambda R_v$$

175
$$\frac{dS_u}{dt} = -\beta S_u (c I_v + I_u) + \delta R_u + (1 - f)\mu - \lambda S_u$$

176
$$\frac{dE_u}{dt} = -\alpha E_u + \beta S_u (c I_v + I_u) - \lambda E_u$$

177
$$\frac{dI_u}{dt} = -\gamma I_u + \alpha E_u - \lambda I_u$$

178
$$\frac{dR_u}{dt} = \gamma I_u (1 - \sigma) - \delta R_u - \lambda R_u$$

179 Where S represents the susceptible population, E the exposed population, I the infectious
 180 population, and R the recovered population. Subscript v represents the vaccinated or compliant
 181 sub-population, while subscript u represents the unvaccinated or noncompliant sub-population.

182 Model parameters are summarized in Table 2.

183

184 **Table 2:** Model parameters for SEIRS model.

Parameter	Symbol	Value	Source
Latency period	$1/\alpha$	3 days	[30]

Reproductive number	R_0	5.7 individuals	[17]
Infectious period	$1/\gamma$	10 days	[31]
Natural immunity duration	$1/\delta$	18 months	[32]
Infection fatality rate	σ	0.68%	[33]
Population birth rate	μ	1% annually	[34]
Population death rate	λ	0.9% annually	[35]
Fraction compliant	f	Variable	
Protective efficacy	1-b	Variable	
Transmission reduction	1-c	Variable	

185 All parameters defining the ODE-based SEIRS model. In this analysis, the fraction compliant,
 186 protective efficacy against infection, and reduction in transmission are treated as independent
 187 variables.

188

189 According to the CDC, R_0 for SARS-CoV-2 under pre-pandemic social and economic conditions is
 190 estimated to be approximately 5.7 [17]. For the purpose of this study, an R_0 of 5.7 is used to
 191 represent epidemiological conditions under a theoretical full return to pre-pandemic activity.

192 The contact rate β is derived from the relationship between R_0 and the infectious period:

$$193 \quad \beta = \gamma R_0$$

194 In this “normal” scenario, disease reduction interventions reduce the compliant population’s
 195 infection rate by the factor b, which represents the intervention’s protective efficacy, and the
 196 compliant population’s transmission rate by the factor c, representing the intervention’s
 197 reduction in transmissibility. For simplicity, the reduction of transmission was assumed to be

198 equivalent to the protective efficacy (reduction of susceptibility) of each intervention. This is an
199 optimistic assumption; in some cases, an intervention may provide little or no reduction in
200 transmission in compliant infected individuals.

201 The model's initial conditions are set to approximate current United States disease prevalence
202 and seroprevalence (as of September 2020):

203 $I(t = 0) = 0.2\%$

204 $R(t = 0) = 8\% [36]$

205 Our model lacks a seasonal component for SARS-CoV-2 transmission, as such associations have
206 been conjectured [37] but not proven, and it also assumes a 18-month duration of natural
207 immunity, as an optimistic estimate based on the duration of antibody responses currently
208 reported [20–22]. The disease-preventing interventions and return to normalcy (which would
209 correspond to a return to the pre-pandemic R_0 of 5.7) are assumed to occur at the beginning of
210 the simulation interval.

211

212 ***Compliance Sweeps***

213 To gauge the impact of NPI or vaccine compliance on population outcomes, we varied the
214 compliant fraction under a series of simulated vaccine or NPI deployment schemes with varying
215 degrees of protective efficacy. The model allows tracking of outcomes for the population as a
216 whole and for the compliant and noncompliant sub-populations.

217

218 **Results**

219 *Structural incentives for noncompliance with interventions aimed at controlling COVID-19*

220 In Figure 1, we modeled the decision to comply with public health measures in terms of its
221 perceived short-term impact to individuals. In game theory, a Nash equilibrium is a strategy
222 which has a higher payoff for the individual than all other possible strategies (“no regrets”)[29].
223 Individuals using a strategy that is a Nash equilibrium are unable to improve their outcome by
224 switching strategies. Strikingly, for a large region of parameter space in this model,
225 noncompliance is a Nash equilibrium. Even so, one can make the case that, using realistic
226 estimates for risk of infection and risk of adverse outcomes given infection, compliance would
227 still be a rational choice for the vast majority of the population. For example, for an
228 intervention that is 50% effective at reducing the risk of infection, when 2% of individuals are
229 infected, compliance is a Nash equilibrium at a 1% relative cost (ratio of the loss of quality of
230 life associated with the intervention over the cost of infection in terms of risk of mortality,
231 morbidity, and disability). While the decision to comply is determined by the perceived cost of
232 infection and the perceived cost of intervention, the actual costs may be very different. The
233 cost of wearing a mask, for example, is likely to be much less than the risk-weighted cost of
234 death or disability due to COVID-19 (see Tables S1, S2 for a more detailed analysis).

235

236 *Failure to suppress COVID-19 results in waves of disease*

237 As shown in Figure 2, insufficient reduction in COVID-19 transmission allows the disease to
238 persist upon a rapid return to pre-pandemic activity and spread in multiple waves over time.
239 The model does not account for changes in behavior or environmental factors over time, so
240 these oscillations in transmission are caused by a predator-prey dynamic within the SEIRS
241 system rather than triggered by external factors. This dynamic is driven by the time-variant

242 availability of susceptible hosts as immunity wanes in former COVID-19 infected individuals.
243 Panels 2A-C represent a high compliance (95%) scenario with a 50% effective intervention. The
244 efficacy of an intervention describes the fraction of possible transmission events it prevents. In
245 this case, the oscillations and variability in risk for the compliant population are relatively small
246 because the intervention serves to dampen the oscillations in transmission rate. However, in
247 panels 2D-F, representing a low compliance (50%) scenario with a 50% effective intervention,
248 the oscillatory pattern is much more pronounced and risk to the compliant population is
249 variable over time (Fig. 2F). Additionally, the cumulative risk to the compliant population
250 relative to the noncompliant population is higher when more of the population is noncompliant
251 (Fig. 2C, F).

252

253 *Near-term suppression of COVID-19 requires a high degree of compliance*
254 In the short term, to suppress COVID-19 while returning to pre-pandemic social and economic
255 activity, an intervention with a high degree of efficacy and compliance is required (Fig. 3).
256 Although effective suppression can be achieved with an intervention with as low as 65%
257 efficacy, at least 80% compliance is required for even the most effective interventions. The
258 predicted number of cases in the next year span three orders of magnitude, from less than one
259 million cases to hundreds of millions of cases, depending on the effectiveness and the degree of
260 compliance with transmission reduction interventions.

261

262 *If COVID-19 becomes endemic, steady-state yearly spread depends on population compliance*

263 If immunity to SARS-CoV-2 by natural infection is not life-long, as suggested by many studies
264 [20–22], and if effective interventions are not undertaken at a large scale, the virus will become
265 endemic. As shown in Figure 4, this means that in the long term, SARS-CoV-2 will reach a
266 steady-state prevalence in the population. For a 50% effective intervention, the disease will
267 become endemic even if the entire population complies with the intervention. As expected, the
268 benefit of compliance for an individual is smaller for a 50% effective intervention (Fig. 4A)
269 relative to a 95% effective intervention (Fig. 4B). The full compliance scenario for the 95%
270 effective intervention is an example of disease suppression.

271

272 *Failure to suppress COVID-19 in the long-term results in persistent high disease burden*
273 If complete suppression of disease is not achieved, a high annual disease burden persists
274 indefinitely in most scenarios (Fig. 5). The marginal cost in terms of yearly cases for failures to
275 suppress disease is highest for near-success cases and is steeply dependent on the degree of
276 compliance (Fig. S1, see Supplementary Figures). This suggests that the best strategic objective
277 for a stable return to pre-pandemic activity is complete suppression of SARS-CoV-2 spread.

278

279 *Complete suppression requires high compliance and at least 60% efficacy*
280 In Figure 5, the steady-state yearly caseload of SARS-CoV-2 is plotted against the fraction of the
281 population complying with a variety of theoretical interventions. For interventions with greater
282 than 60% efficacy, complete suppression can be achieved if compliance is above a certain high
283 threshold, depending on the intervention's efficacy. For example, with an intervention with
284 70% efficacy, complete suppression can be achieved with at least 92% compliance. Increasingly

285 effective interventions reduce the compliance threshold for complete suppression and reduce
286 the yearly caseload in endemic scenarios. However, the impact of progressive improvements in
287 efficacy shrinks as 100% efficacy is approached. Even for a 99% effective vaccine, greater than
288 80% compliance is required to achieve complete suppression of SARS-CoV-2. This suggests that
289 a high degree of intervention efficacy cannot compensate for the epidemiological impact of a
290 large noncompliant population.

291

292 Additionally, we note that the duration of immunity does not impact these compliance
293 thresholds for achieving complete suppression (Fig. S3, see Supplementary Figures). However,
294 the duration of immunity does impact the expected yearly disease burden. To further
295 demonstrate this point, Figures 2, 3, 4, 5, and S2 were reimplemented in the additional
296 materials assuming a shorter (6-month) or longer (36-month) duration of immunity (Figs. S5-
297 S14).

298

299 *If complete suppression is not achieved, compliant populations remain at risk without a highly
300 effective vaccine*

301 Although improvements in vaccine or intervention efficacy face diminishing returns on the
302 population level and cannot fully compensate for poor compliance, compliant individuals stand
303 to gain from even small improvements in efficacy (Fig. 6). This means that although the
304 compliance threshold required for complete suppression may not shift substantially as efficacy
305 improves, the incentive for individuals to comply or seek vaccination on a voluntary basis will
306 increase as the efficacy increases.

307

308 **Discussion**

309 Our work makes the case that noncompliance is embedded deeply in human nature, as
310 individuals optimizing their own self-interest can justify their actions in terms of their own
311 perceived cost-benefit.

312

313 Individuals may perceive noncompliance as favorable for a number of reasons [38, 39]. They
314 may view their own risk of being infected as lower than average (the optimism bias [40], which
315 has been documented as a risk factor in predicting noncompliance for COVID-19 mitigation
316 measures [41]), or they may view their own risk of adverse outcomes as a result of infection as
317 being lower than average [27]. Globally, the public health messaging around noncompliance
318 has focused on the low risk of death for younger individuals [42–44] and has invoked the
319 imagery of “shielding” highly vulnerable populations from the disease [45] as an altruistic
320 motive [46]. To the extent that many Western countries at present are facing uncontrolled
321 disease spread, it is likely that invoking altruism may not be the most effective means of disease
322 control. Underestimating the risk of infection may also lead to individuals believing that
323 noncompliance is the better choice.

324

325 The interplay between risk perception and compliance is complex, and fear may also play a
326 paradoxical role in noncompliance. A number of studies have demonstrated a link between
327 emotions and cognitive assessment of risk. In particular, high levels of fear coupled with a low
328 sense of efficacy may create a defensive response in individuals who then proceed to dismiss

329 the risk (“we’re all going to die anyway”)[47]. Studies have also shown that psychological affect
330 plays a role in risk perception in individuals who are less comfortable and/or experienced
331 interpreting probability [48, 49].

332

333 Regardless of the underlying causes, a Nash equilibrium of noncompliance creates a Tragedy of
334 the Commons situation, where individuals acting according to their own self-interest create
335 outcomes that are suboptimal for the common good, by spoiling the shared resource through
336 their collective actions. The term Tragedy of the Commons dates back to an influential article
337 written over fifty years ago [50], which in turn was inspired by a nineteenth-century essay
338 describing grazing practices of farmers. Tragedy of the Commons situations are indeed common
339 in the fields of economics, politics, environmental policy and sociology. What makes Tragedy of
340 the Commons situations particularly intractable is that it usually only takes a small proportion
341 of individuals optimizing for their own self-interest to create devastating externalities for the
342 rest of the population. This behavior underscores the limitations of the laissez-faire,
343 individualistic, approach to disease control during a pandemic. While the underpinnings of such
344 a laissez-faire approach are often said to lie in the economic theory of utilitarianism [51], put
345 forward by John Stuart Mill, such an approach actually violates the standard originally laid out
346 by Mill by which a person’s liberty may be restricted: “The only purpose for which power can
347 rightfully be exercised over any member of a civilized community, against his will, is to prevent
348 harm to others” [52].

349

350 Given the ubiquity of the problem, some public policy solutions can be found that have close
351 analogies to successful interventions in other spheres of human activity. First, public health
352 messaging that seeks to alter the Nash equilibrium at an individual level are worth exploring. In
353 individualistic societies, this may be accomplished by de-emphasizing altruism and focusing on
354 the individual cost-benefit. One way this may be achieved is by emphasizing the long-term
355 morbidity costs (such as cryptic heart, lung, brain and kidney damage) as have been
356 documented to occur in even asymptomatic COVID-19 patients in an age-independent manner
357 [53–55]. An additional approach is to provide an accurate and current picture of the risk of
358 contracting the disease. Second, public health policy that creates costs for noncompliers may
359 serve to shift some of the externalities back on to the originator (as was the case with mask
360 ordinances during the 1918 Flu [4] and fines imposed for noncompliance with COVID-19
361 prevention measures in some countries [56, 57]). Third, public health interventions should
362 engage at the level of the community. Public health and communications experts could test a
363 number of different messages that underscore the downside of negative externality-creating
364 behavior at a societal level. Some of these approaches have been used previously in the context
365 of vaccine acceptance [58]. It is worth noting that our analysis points out a potential
366 mechanism for the high levels of compliance observed in countries such as Korea [23], with
367 strong societal norms and a positive view of COVID-19 restrictions such as mask-wearing [59,
368 60]. In these cultures, the prevailing cultural beliefs may serve to lower the cost of the
369 intervention. In this context, we note that there is a modest association (Fig. S4, see
370 Supplementary Figures) between societies with strong societal norms (“tight cultures” [61]) and
371 the total case count per million at this point in the pandemic ($p=0.04$).

372

373 From the perspective of biomedical interventions, our work points out that interventions with a
374 high degree of protective efficacy are required for complete suppression of SARS-CoV-2, making
375 this disease particularly challenging to control. Highly effective interventions have the dual
376 effect of making the creation of negative externalities less beneficial for the noncompliant
377 population (Fig. 4), and also increasing the benefit to the compliant population (Fig. 6). Notably,
378 highly effective interventions also provide more wiggle room for public policy, as the threshold
379 level of compliance required for the complete suppression of COVID-19 drops from
380 approximately 95% for a minimally effective intervention to approximately 80% for highly
381 effective interventions. Another path to disease suppression lies in implementing passive
382 interventions that reduce the R_0 , such as improving ventilation. Such passive interventions,
383 being not subject to the problem of individual noncompliance, can serve to lower the bar for
384 compliance for any given intervention to achieve complete suppression (Fig. S2, see
385 Supplementary Figures).

386

387 Notably, our work optimistically assumes that the impact of biomedical and nonpharmaceutical
388 interventions is not variable over time. In some settings, “pandemic fatigue” may drive
389 increased noncompliance with nonpharmaceutical interventions over time [62], and relaxations
390 of individual caution and public health guidelines may follow improvements in regional
391 transmission, creating reactive variability in intervention effectiveness. Although biomedical
392 interventions such as vaccines are less susceptible to variability in day-to-day decision-making,
393 immunity to SARS-CoV-2 is expected to wane over a period of months [20–22], which can be

394 expected to impact the duration of vaccine protection. Challenges in vaccine distribution and
395 compliance may compound this waning immunity, reducing the apparent effectiveness of
396 vaccines at the individual and population scale. Additionally, we assumed that compliant
397 individuals have a reduced risk of transmission upon infection, equal to their reduction in risk of
398 infection. This indirect benefit is challenging to measure in clinical trials, and preclinical studies
399 show that some vaccine candidates are capable of reducing nasal viral load (and by implication,
400 risk of transmission) in vaccinated animals [63, 64], while others are not [65, 66].

401

402 Taken together, our work demonstrates that noncompliance with measures to control COVID-
403 19 is at once easy to justify on an individual level and leads to devastating public health
404 consequences. Even under optimistic assumptions about the transmission benefit and
405 durability of preventive interventions, noncompliance presents a significant obstacle to COVID-
406 19 suppression. Three key messages are worth keeping in mind. First, the importance of
407 focusing on complete suppression as a desirable end goal for SARS-CoV-2 (Fig. 5) and as a
408 prerequisite for a return to a pre-pandemic lifestyle. SARS-CoV-2 is highly transmissible and can
409 be expected to circulate at high rates if it becomes endemic. Second, the need for public policy
410 to focus on compliance as a key prerequisite for both short-term suppression (Fig. 3) and long-
411 term complete suppression (Fig. 5) of COVID-19, and to seek ways to alter the space where
412 compliance is the Nash equilibrium by increasing the cost of noncompliance. Finally, the need
413 to focus on highly effective interventions from a biomedical perspective and to view partially
414 effective prophylactics as contributors to the solution rather than the solution in its entirety.

415

416 It is our hope that this work draws the attention of the biomedical community to how high the
417 bar is actually set for us to return to normalcy, and to public policymakers to highlight the need
418 for concerted action that is focused on complete disease suppression.

419

420 **Conclusions**

421 In this study, we analyzed the impact of noncompliance with COVID-19 disease control
422 measures (such as masks and vaccines) on public health. Using a game-theoretic framework,
423 we demonstrated that noncompliance can be rationalized in terms of benefit to the individual.
424 We further demonstrated, using SEIRS model-based analyses that this noncompliance is
425 a significant impediment to COVID-19 disease control. The compliance threshold for achieving
426 complete suppression of SARS-CoV-2 disease spread is a function of the intervention's efficacy.
427 A minimum of 82% compliance is required to achieve complete suppression of SARS-CoV-2
428 spread under an R_0 of 5.7 for an intervention that is 100% effective. These findings demonstrate
429 that a successful COVID-19 suppression strategy must involve highly effective interventions for
430 which broad compliance can be achieved. Interventions that provide benefit to individuals in
431 compliance may inspire greater uptake by shifting the externality onto noncompliant
432 individuals.

433

434 **Declarations**

435 *Ethics approval and consent to participate*

436 Not applicable. No human subjects, human tissues, or human data were involved in the
437 execution of this study.

438 *Consent for publication*

439 Not applicable.

440 *Availability of data and materials*

441 Data sharing is not applicable to this article as no datasets were generated or analysed

442 during the current study. Please contact AC for simulation code.

443 *Competing interests*

444 AC, MS are employees of and shareholders in Fractal Therapeutics. DvE, DW and RN

445 are advisors to and shareholders of Fractal Therapeutics. KJ, SR, JF, and NH have no

446 conflicts of interest to report.

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449 *Authors' contributions*

450 MS provided epidemiological modeling and related figures. DVE provided game

451 theoretic modeling and related analysis. MS, DVE, and AC contributed substantially to

452 manuscript writing. KJ, SR, JF, DW, RN, and NH read, revised, and approved the final

453 manuscript.

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455 Not applicable.

456 **List of Abbreviations**

457 R₀: Basic reproduction number

458 SEIRS: Susceptible-Exposed-Infected-Recovered-Susceptible

459 ODE: Ordinary differential equations

460 NPI: Nonpharmaceutical intervention

461

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642

643

644 **Figure 1:** Noncompliance is a Nash equilibrium when infection rates are low or prevention is
645 costly or ineffective. Intervention efficacy and intervention cost conditions for which
646 noncompliance is a Nash equilibrium (red) or not a Nash equilibrium (blue) if the disease is
647 present in 2% of individuals in the population. Intervention cost relative to infection cost is
648 defined as the ratio of intervention cost to risk-weighted infection cost.

649

650 **Figure 2:** Failure to eradicate COVID-19 results in waves of disease upon rapid return to pre-
651 pandemic activity. Panels A and D represent the fraction of the population, including both
652 compliant and noncompliant individuals, that is susceptible, exposed, infectious, and recovered
653 populations over time after a return to pre-pandemic conditions under (A-C) 95% compliance or
654 (D-F) 50% compliance with a 50% effective intervention. Panels B and E demonstrate the
655 fraction of compliant and noncompliant individuals who are infected over time. Panels C and F
656 demonstrate the cumulative hazard ratio for infection in noncompliant versus compliant
657 individuals.

658

659 **Figure 3:** Short-term suppression of COVID-19 requires a high degree of compliance with a
660 highly effective measure. Total US COVID-19 infections in the next year under interventions
661 with varying efficacy and compliance. Black box on panel A shows region expanded in panel B.
662 Panel B is displayed on a log scale.

663

664 **Figure 4:** Steady-state individual risk is impacted by individual and population compliance.
665 Cumulative infections per individual under a 50% (A) or 90% (B) effective intervention. Three
666 scenarios are simulated: full noncompliance, full compliance, and 70% compliance (with
667 outcomes for compliant and noncompliant individuals shown).

668

669 **Figure 5:** Population-level impact of interventions is highly dependent on compliance. Yearly US
670 cases at steady-state under interventions with varying degrees of efficacy and compliance.

671

672 **Figure 6:** More effective interventions provide greater benefit to compliant individuals if
673 disease spread persists. Reduction in yearly likelihood of infection for compliant individuals as a
674 function of the overall fraction of the population in compliance and the efficacy of the
675 intervention.

676

677 **Additional data**

678 Supplementary Figures.docx

679 Additional figures and tables supporting the text and exploring modeling assumptions.

Figures

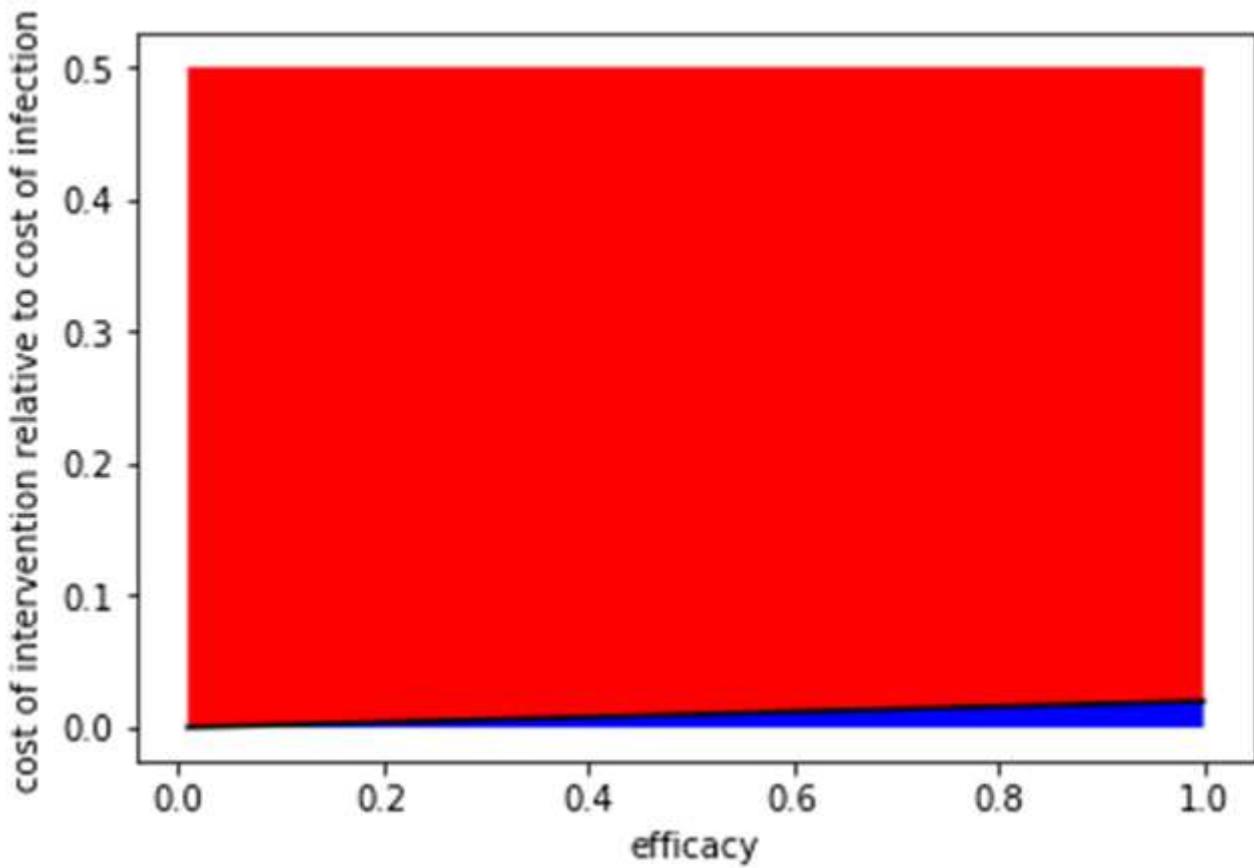


Figure 1

Noncompliance is a Nash equilibrium when infection rates are low or prevention is costly or ineffective. Intervention efficacy and intervention cost conditions for which noncompliance is a Nash equilibrium (red) or not a Nash equilibrium (blue) if the disease is present in 2% of individuals in the population. Intervention cost relative to infection cost is defined as the ratio of intervention cost to risk-weighted infection cost.

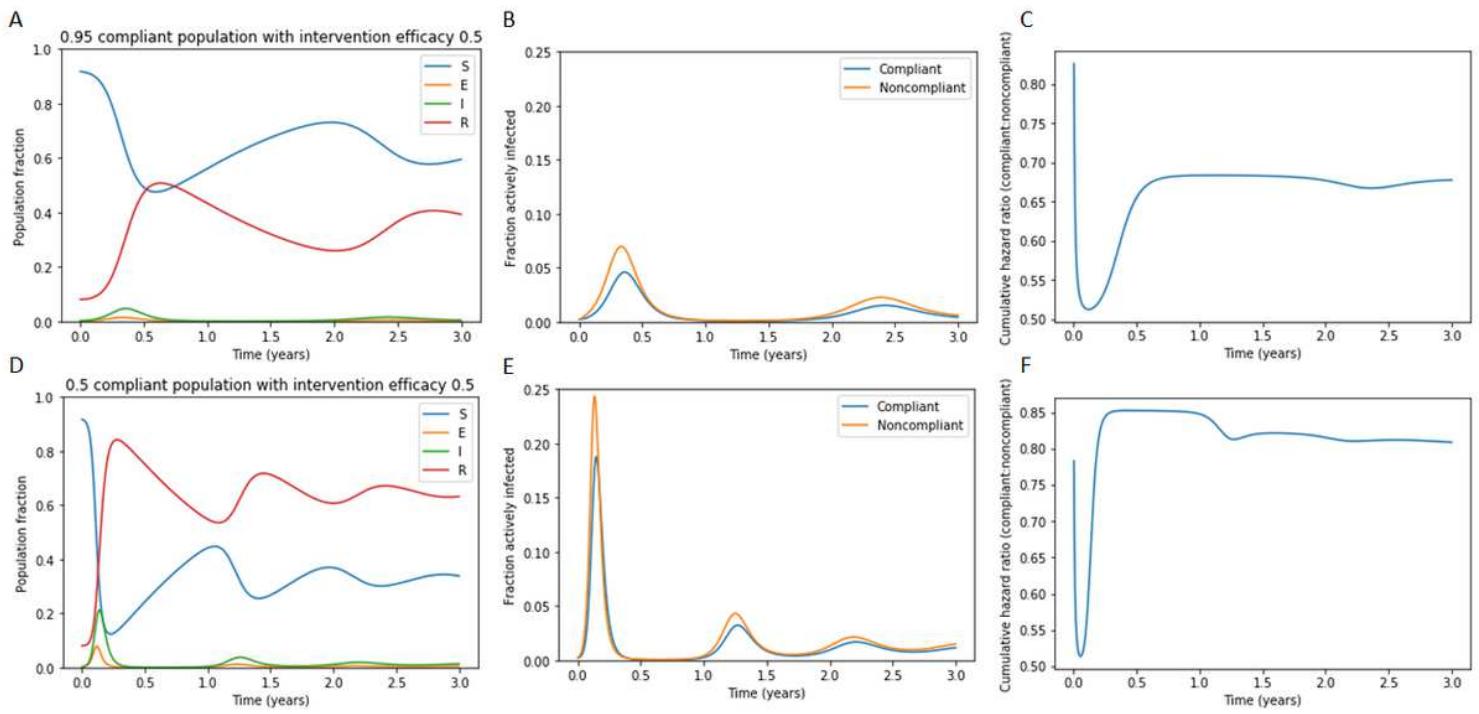


Figure 2

Failure to eradicate COVID-19 results in waves of disease upon rapid return to pre-pandemic activity. Panels A and D represent the fraction of the population, including both compliant and noncompliant individuals, that is susceptible, exposed, infectious, and recovered populations over time after a return to pre-pandemic conditions under (A-C) 95% compliance or (D-F) 50% compliance with a 50% effective intervention. Panels B and E demonstrate the fraction of compliant and noncompliant individuals who are infected over time. Panels C and F demonstrate the cumulative hazard ratio for infection in noncompliant versus compliant individuals.

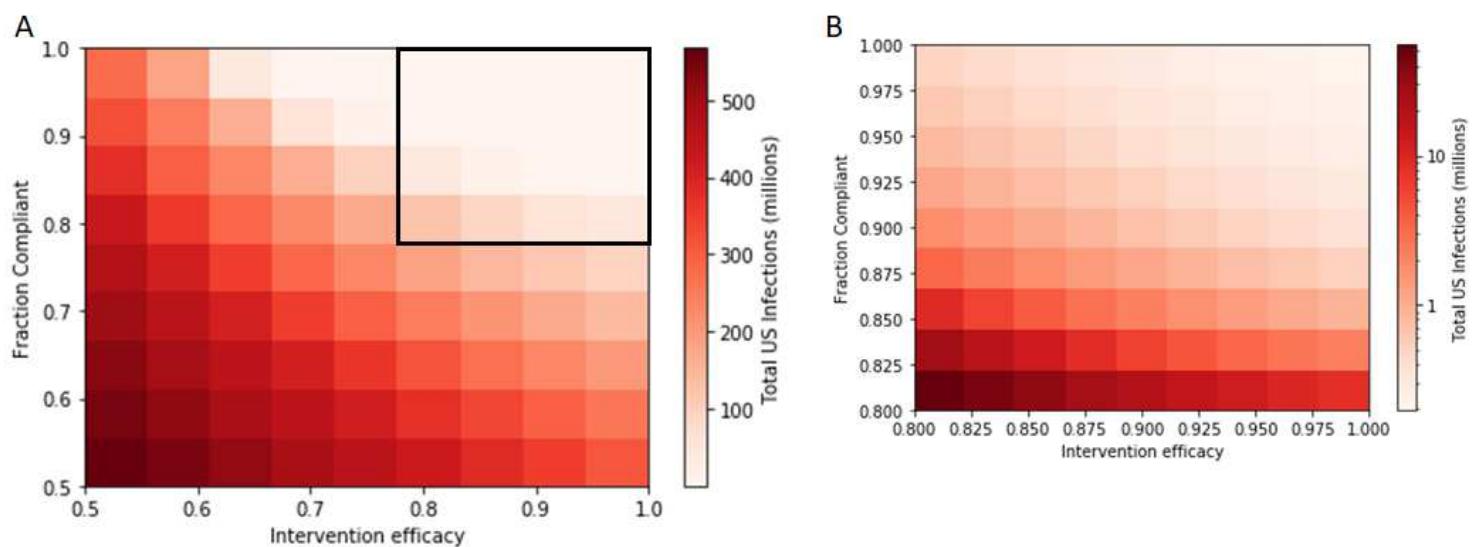


Figure 3

Short-term suppression of COVID-19 requires a high degree of compliance with a highly effective measure. Total US COVID-19 infections in the next year under interventions with varying efficacy and compliance. Black box on panel A shows region expanded in panel B. Panel B is displayed on a log scale.

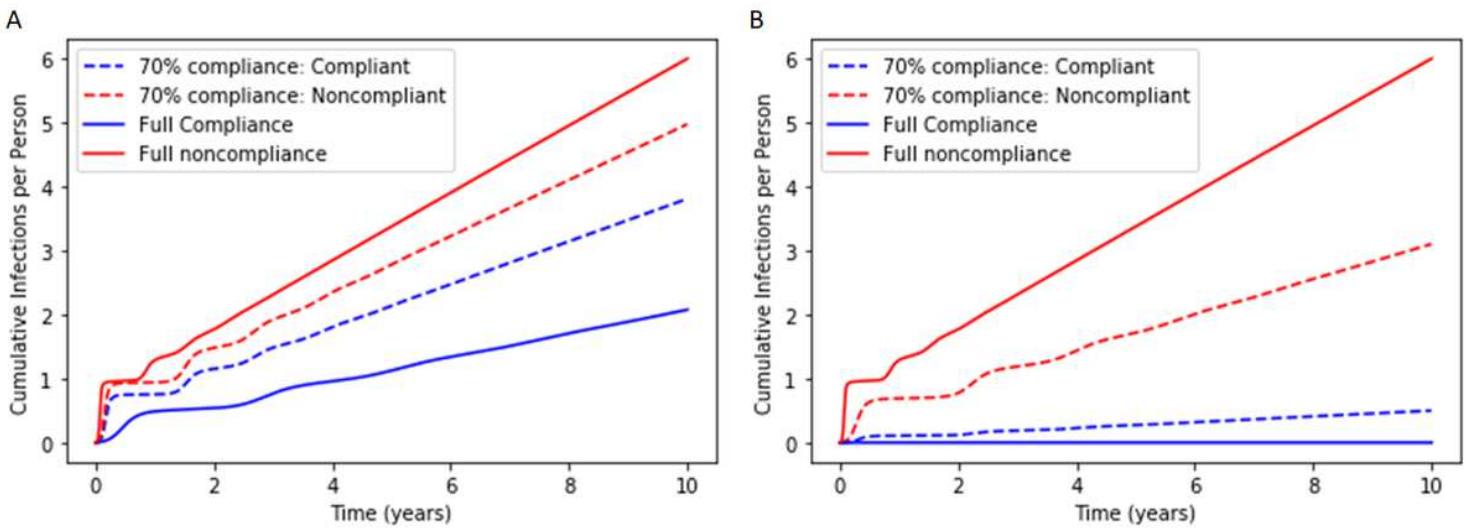


Figure 4

Steady-state individual risk is impacted by individual and population compliance. Cumulative infections per individual under a 50% (A) or 90% (B) effective intervention. Three scenarios are simulated: full noncompliance, full compliance, and 70% compliance (with outcomes for compliant and noncompliant individuals shown).

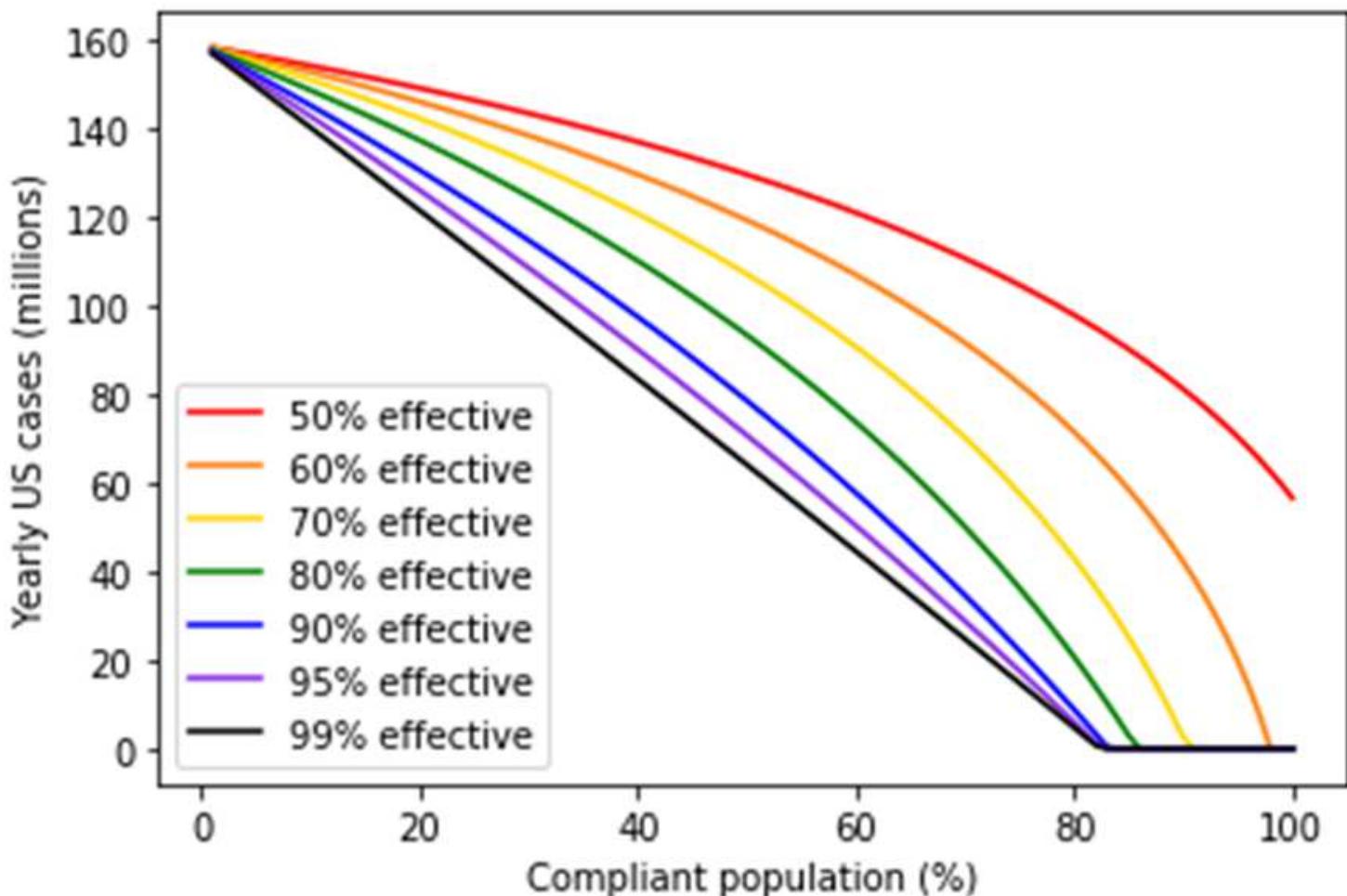


Figure 5

Population-level impact of interventions is highly dependent on compliance. Yearly US cases at steady-state under interventions with varying degrees of efficacy and compliance.

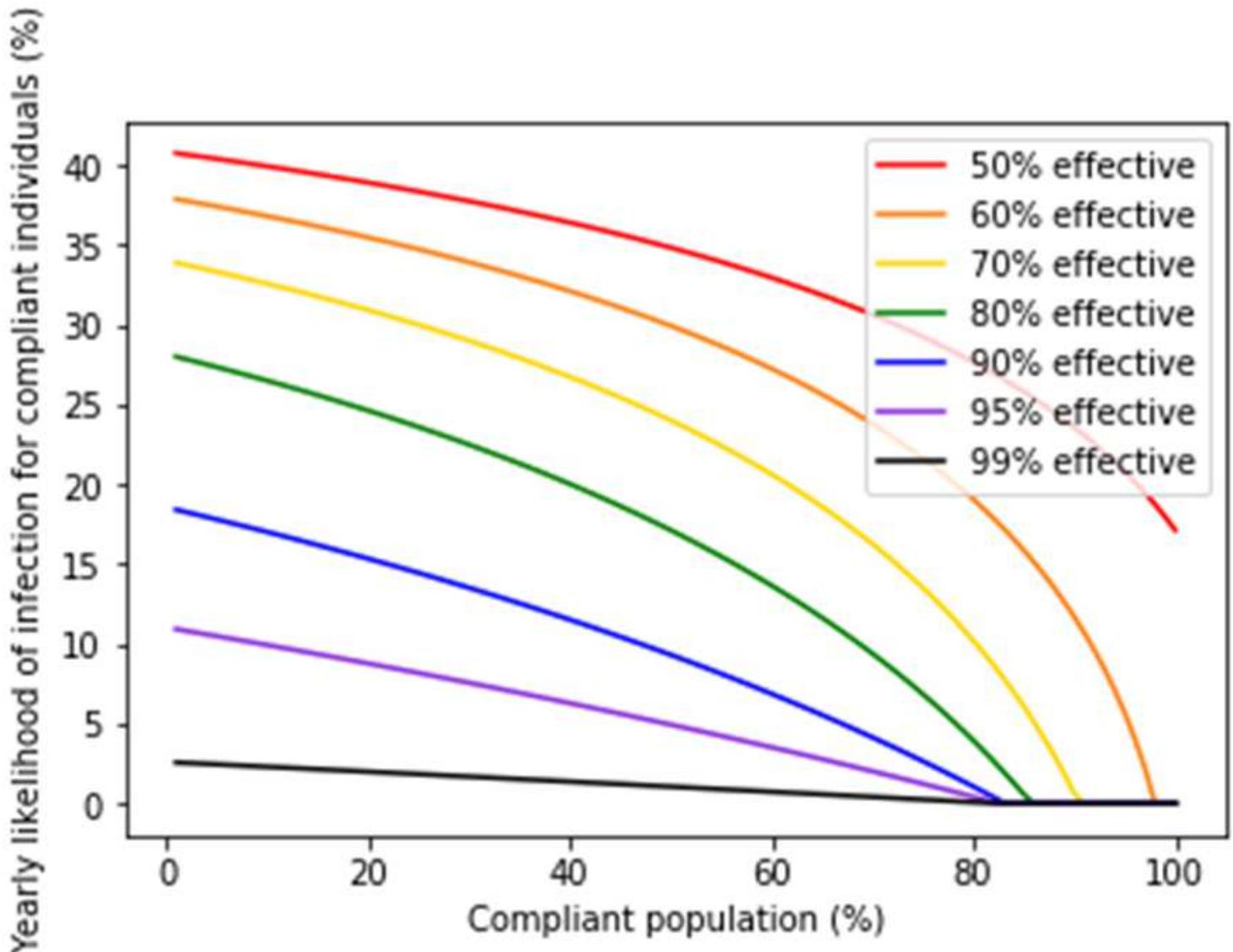


Figure 6

More effective interventions provide greater benefit to compliant individuals if disease spread persists. Reduction in yearly likelihood of infection for compliant individuals as a function of the overall fraction of the population in compliance and the efficacy of the intervention.

Supplementary Files

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- [SupplementaryFigures.docx](#)