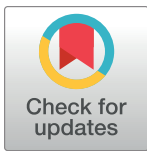


RESEARCH ARTICLE

Social distancing to slow the US COVID-19 epidemic: Longitudinal pretest–posttest comparison group study

Mark J. Siedner^{1,2,3}*, Guy Harling^{3,4,5,6,7}, Zahra Reynolds¹, Rebecca F. Gilbert¹, Sebastien Haneuse⁸, Atheendar S. Venkataramani^{9,10}, Alexander C. Tsai^{1,2,7}

1 Massachusetts General Hospital, Boston, Massachusetts, United States of America, **2** Harvard Medical School, Boston, Massachusetts, United States of America, **3** Africa Health Research Institute, KwaZulu-Natal, South Africa, **4** University College London, London, United Kingdom, **5** MRC/Wits Agincourt Unit, Rural Public Health and Health Transitions Research Unit, University of the Witwatersrand, Johannesburg, South Africa, **6** Department of Epidemiology, Harvard T.H. Chan School of Public Health, Boston, Massachusetts, United States of America, **7** Harvard Center for Population and Development Studies, Cambridge, Massachusetts, United States of America, **8** Department of Biostatistics, Harvard T.H. Chan School of Public Health, Boston, Massachusetts, United States of America, **9** Department of Medical Ethics and Health Policy, Perelman School of Medicine, University of Pennsylvania, Philadelphia, Pennsylvania, United States of America, **10** Leonard Davis Institute of Health Economics, University of Pennsylvania, Philadelphia, Pennsylvania, United States of America



OPEN ACCESS

Citation: Siedner MJ, Harling G, Reynolds Z, Gilbert RF, Haneuse S, Venkataramani AS, et al. (2020) Social distancing to slow the US COVID-19 epidemic: Longitudinal pretest–posttest comparison group study. *PLoS Med* 17(8): e1003244. <https://doi.org/10.1371/journal.pmed.1003244>

Academic Editor: Donald A. Redelmeier, University of Toronto, CANADA

Received: April 27, 2020

Accepted: July 2, 2020

Published: August 11, 2020

Copyright: © 2020 Siedner et al. This is an open access article distributed under the terms of the [Creative Commons Attribution License](https://creativecommons.org/licenses/by/4.0/), which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

Data Availability Statement: The data on social distancing policies are contained within the Supplementary Appendix (Table A in [S1 Text](#)). The data on COVID-19 cases and deaths are publicly available from <https://github.com/nytimes/covid-19-data>.

Funding: This study was funded in part by a Wellcome Trust Sir Henry Dale Fellowship to GH (<https://wellcome.ac.uk/grant-funding/schemes/sir-henry-dale-fellowships>) and a grant from the

* These authors contributed equally to this work.

* msiedner@mgh.harvard.edu

Abstract

Background

Social distancing measures to address the US coronavirus disease 2019 (COVID-19) epidemic may have notable health and social impacts.

Methods and findings

We conducted a longitudinal pretest–posttest comparison group study to estimate the change in COVID-19 case growth before versus after implementation of statewide social distancing measures in the US. The primary exposure was time before (14 days prior to, and through 3 days after) versus after (beginning 4 days after, to up to 21 days after) implementation of the first statewide social distancing measures. Statewide restrictions on internal movement were examined as a secondary exposure. The primary outcome was the COVID-19 case growth rate. The secondary outcome was the COVID-19-attributed mortality growth rate. All states initiated social distancing measures between March 10 and March 25, 2020. The mean daily COVID-19 case growth rate decreased beginning 4 days after implementation of the first statewide social distancing measures, by 0.9% per day (95% CI –1.4% to –0.4%; $P < 0.001$). We did not observe a statistically significant difference in the mean daily case growth rate before versus after implementation of statewide restrictions on internal movement (0.1% per day; 95% CI –0.04% to 0.3%; $P = 0.14$), but there is substantial difficulty in disentangling the unique associations with statewide restrictions on internal movement from the unique associations with the first social distancing measures. Beginning

Sullivan Family Foundation to ACT. The funders had no role in study design, data collection and analysis, decision to publish, or preparation of the manuscript.

Competing interests: I have read the journal's policy and the authors of this manuscript have the following competing interests: GH reports having received funding within the past five years from the U.S. National Institutes of Health, the Wellcome Trust, the Royal Society, and the International Initiative for Impact Evaluation (3ie). ACT receives a stipend as a Specialty Consulting Editor for PLOS Medicine and serves on the journal's editorial board. All other authors have declared that no competing interests exist.

Abbreviations: CI, confidence interval; COVID-19, coronavirus disease 2019; IQR, interquartile range.

7 days after social distancing, the COVID-19-attributed mortality growth rate decreased by 2.0% per day (95% CI -3.0% to -0.9% ; $P < 0.001$). Our analysis is susceptible to potential bias resulting from the aggregate nature of the ecological data, potential confounding by contemporaneous changes (e.g., increases in testing), and potential underestimation of social distancing due to spillover effects from neighboring states.

Conclusions

Statewide social distancing measures were associated with a decrease in the COVID-19 case growth rate that was statistically significant. Statewide social distancing measures were also associated with a decrease in the COVID-19-attributed mortality growth rate beginning 7 days after implementation, although this decrease was no longer statistically significant by 10 days.

Author summary

Why was this study done?

- There are few empirical data about the population health benefits of imposing statewide social distancing measures to reduce transmission of severe acute respiratory syndrome coronavirus 2, which causes coronavirus disease 2019 (COVID-19).

What did the researchers do and find?

- We compared data from each state before versus after implementation of statewide social distancing measures to estimate changes in the mean daily COVID-19 case growth rate.
- The case growth rate declined by approximately 1% per day beginning 4 days (approximately 1 incubation period) after statewide social distancing measures were implemented.
- Our model implies that social distancing reduced the total number of COVID-19 cases by approximately 1,600 reported cases at 7 days after implementation, by approximately 55,000 reported cases at 14 days after implementation, and by approximately 600,000 reported cases at 21 days after implementation.

What do these findings mean?

- Statewide social distancing measures were associated with a reduction in the growth rate of COVID-19 cases in the US.
- Our analysis is susceptible to potential bias resulting from the aggregate nature of the data, potential confounding by other changes that occurred during the study period (e.g., increases in testing), and potential underestimation of social distancing due to spillover effects from neighboring states.

Introduction

Modeling estimates suggested that up to 80% of Americans would be infected with severe acute respiratory syndrome coronavirus 2 (SARS CoV-2) if no preventive interventions were implemented [1]. In response, US state governments implemented social distancing measures in an attempt to limit its transmission and reduce morbidity and mortality from coronavirus disease 2019 (COVID-19). Such measures have been implemented during prior pandemics, with moderate success [2–5], and are predicted to prevent a rapid, overwhelming epidemic in modeling studies [6]. Data on social distancing and its associations with the course of the COVID-19 pandemic are now beginning to emerge, although no studies to our knowledge have examined changes in COVID-19-attributed mortality as an outcome. Recent studies have shown slowed COVID-19 epidemic growth coinciding with spontaneous reductions in internal movement [7,8], as well as after implementation of social distancing measures [9–11]. Because of the economic and social costs associated with social distancing measures [12,13], there is immense value in using different modalities to quantify the extent to which they have benefits for epidemic control. Herein, we contribute to this body of evidence, using a longitudinal pretest–posttest comparison group study design to compare the daily growth rate of COVID-19 cases, and the daily growth rate of COVID-19-attributed deaths, before versus after implementation of social distancing measures in the US. Our primary aim was to empirically estimate the public health impact of government-mandated non-pharmacological interventions in the period after their initial implementation and prior to their recent staged relaxation.

Methods

Data collection

We searched government websites and third-party sources to identify all statewide social distancing measures implemented between January 21 and May 1, 2020 (see [S1 Text](#)). We applied a broad definition of social distancing measures, using a previously published typology [14]: closures of schools, closures of workplaces, cancellations of public events, restrictions on internal movement, and closures of state borders. Restrictions on internal movement, i.e., shelter-in-place orders (often referred to colloquially as “lockdowns”), are generally the most restrictive of these in terms of their impacts on daily movement. To obtain daily state-specific reported COVID-19 cases and deaths, we used the *New York Times* COVID-19 database (<https://github.com/nytimes/covid-19-data>; last accessed May 26, 2020). Reporting of cases and deaths in the *New York Times* database varies by state, but typically includes both laboratory-confirmed and suspected cases, as recommended by the Council of State and Territorial Epidemiologists [15].

Our primary outcome was the rate of change in daily COVID-19 cases in each state, calculated as the natural log of cases on each date minus the natural log of cases on the prior date. Analysis was restricted to days on which a state had at least 30 cumulative cases reported, to minimize any effects of volatile rate changes early in the epidemic. The primary exposure of interest was time, measured as a continuous variable and divided into 2 periods: pre-implementation (14 days prior to, and through 3 days after, implementation of the first statewide social distancing measure) versus post-implementation (4 days after, to up to 21 days after implementation). We selected 4 days after implementation as the transition point based on previously published estimates of the lower end of the 95% confidence interval (CI) of the COVID-19 incubation period [16–18], which is when cases at a population level should be expected to decline in the setting of a structural intervention. For COVID-19, the latent period (between infection and the onset of infectiousness) is shorter than the incubation period

(between infection and the onset of symptoms), but during the time period covered by this study—and as of the time of this writing—active asymptomatic screening programs have not been implemented in the US, so case counts would not have been expected to change until individuals became symptomatic and presented for testing. We limited our analysis to 21 days after implementation to prevent any potential diluting effects resulting from relaxation of some social distancing measures, which in some states began less than 4 weeks after implementation (e.g., Alabama, Alaska, Mississippi, and South Carolina).

The secondary outcome was the rate of change in daily COVID-19-attributed deaths in each state, calculated as the natural log of deaths on each date minus the natural log of deaths on the prior date. Analysis was restricted to days on which a state had at least 30 cumulative deaths reported. In the analysis of COVID-19-attributed deaths, there was less certainty about the hypothesized lag between implementation of social distancing and observed changes in the outcome. The median time from symptom onset to death varies widely in the literature. The COVID-19 Surveillance Group [19] has estimated a median of 8 days from symptom onset to death in Italy, while estimates from China are approximately twice that amount [20–22]. Incorporating an incubation period of 3–5 days, we hypothesized that a beneficial impact of social distancing measures on COVID-19-attributed mortality, if any, would be observed no sooner than 7 days and no later than 14 days after implementation. Therefore, we took a more exploratory approach to modeling the time from enactment of statewide social distancing, selecting time periods for the spline terms (described below) every 3 days during days 4–17 after implementation (i.e., 4, 7, 10, 14, and 17 days), and we allowed for observation in the post-intervention period up through 30 days.

As a secondary exposure, we examined statewide restrictions on internal movement. For states that did not implement statewide restrictions on internal movement during the study period, we set day 0 as 14 days after the last state implemented a restriction on internal movement (April 21, 2020) and attributed the prior 14 days to the pre-implementation period.

Statistical analysis

We fitted mixed effects linear regression models, specifying the log difference in daily cases as the outcome of interest and including a random effect for state, to allow for within-state correlation of cases over time. Explanatory variables included time in days, implementation period, and a time-by-implementation-period product term. This analysis was not conducted as part of a preplanned/registered study protocol. However, we followed a clear analysis plan (see [S1 Text](#)), with minor changes made in response to a rapidly changing epidemic and in response to editorial and reviewer feedback. The initial analysis, described in the manuscript we deposited with the medRxiv preprint server on April 8, 2020 [23], included data on statewide social distancing measures implemented between January 21 and March 30, 2020, and COVID-19 cases through March 31. Prior to submission of the current manuscript, we updated the dataset and extended the study period to include social distancing measures implemented up to April 8 and COVID-19 cases and deaths up to April 8. In response to editorial and reviewer feedback, we further updated the dataset and extended the study period to include social distancing measures implemented up to May 1 and COVID-19 cases and deaths up to May 26, expanded the analysis to include growth in COVID-19-attributed deaths as a secondary outcome, and added an event study specification and sensitivity analyses investigating a range of plausible incubation periods (see below).

To assess the robustness of our findings, we conducted several sensitivity analyses. First, to adjust for potential confounding by population density, we adjusted our estimates by state-level population density [24,25]. Second, to account for weekly periodicity that could also

coincide with implementation of social distancing measures, we also adjusted for day of the week [26,27]. Third, to assess the extent to which early versus late implementation of social distancing measures modified the effects of these actions [28], we stratified our estimates by the size of the epidemic in the state at the time of implementation. Fourth, although implementation of social distancing measures is most likely to show population-level effectiveness in reducing coronavirus transmission beginning 3 days later, at the lower bound of its estimated incubation period [16], we refitted the regression models specifying a range of different incubation periods.

Finally, to assess the extent to which our estimates may have been driven by model specification, we replaced the longitudinal pretest–posttest comparison group study design with a multivariable regression model in which the primary explanatory variables of interest were a series of binary indicators denoting each day before versus after implementation of the first statewide social distancing measures (often described in the econometrics literature as an event study specification [29–31]). This approach compares daily case growth before versus after implementation of the first statewide social distancing measures in states that implemented such measures versus daily case growth in states that did not implement such measures. Our regression model included state fixed effects, to adjust for potential confounding from time-invariant state-level factors or baseline differences in population socioeconomic or health characteristics, and linear and quadratic terms for time (days) to adjust for nationwide secular trends in the outcomes. We computed 95% CIs adjusted for clustering within states, the geographical level at which exposure occurred [32].

Ethics statement

This ecological analysis was based on publicly available data and was exempt from ethical review.

Results

A complete list of dates of statewide social distancing measures, by type of measure and state, is contained in Table A in [S1 Text](#). During March 10–25, all 50 states and the District of Columbia implemented at least 1 statewide social distancing measure (Fig A in [S1 Text](#)). The most widely enacted measures on the first date of implementation were cancellations of public events (34/51 [67%]) and closures of schools (26/51 [51%]). The first social distancing measures were implemented when the median statewide epidemic size was 35 cases (interquartile range [IQR] 17–72).

[Fig 1A](#) shows the mean daily COVID-19 case growth rate mapped against the date of the first statewide social distancing measures. At the date of implementation of the first social distancing measure, states had a mean daily case growth rate of 30.8% (95% CI 29.1–32.6; [Table 1](#)), corresponding to a doubling of total cases every 3.3 days. From 14 days prior to, and through 3 days after, implementation of the first social distancing measure, the mean daily case growth rate did not change (–0.2% per day; 95% CI –0.6% to 0.3%; $P = 0.51$). Beginning 4 days after implementation of the first statewide social distancing measure, the mean daily case growth rate decreased by an additional 0.9% per day (95% CI –0.4% to –1.4%; $P < 0.001$). This estimate corresponds to a mean daily case growth rate that had declined to 26.5% (doubling of total cases every 3.8 days) by day 7 after enactment of the first statewide social distancing measures, to 19.6% (doubling time of 5.1 days) by day 14, and to 12.7% (doubling time of 7.9 days) by day 21.

As of May 1, nearly all (45 [90%]) states had implemented statewide restrictions on internal movement. These restrictions on internal movement were implemented a median of 11 days

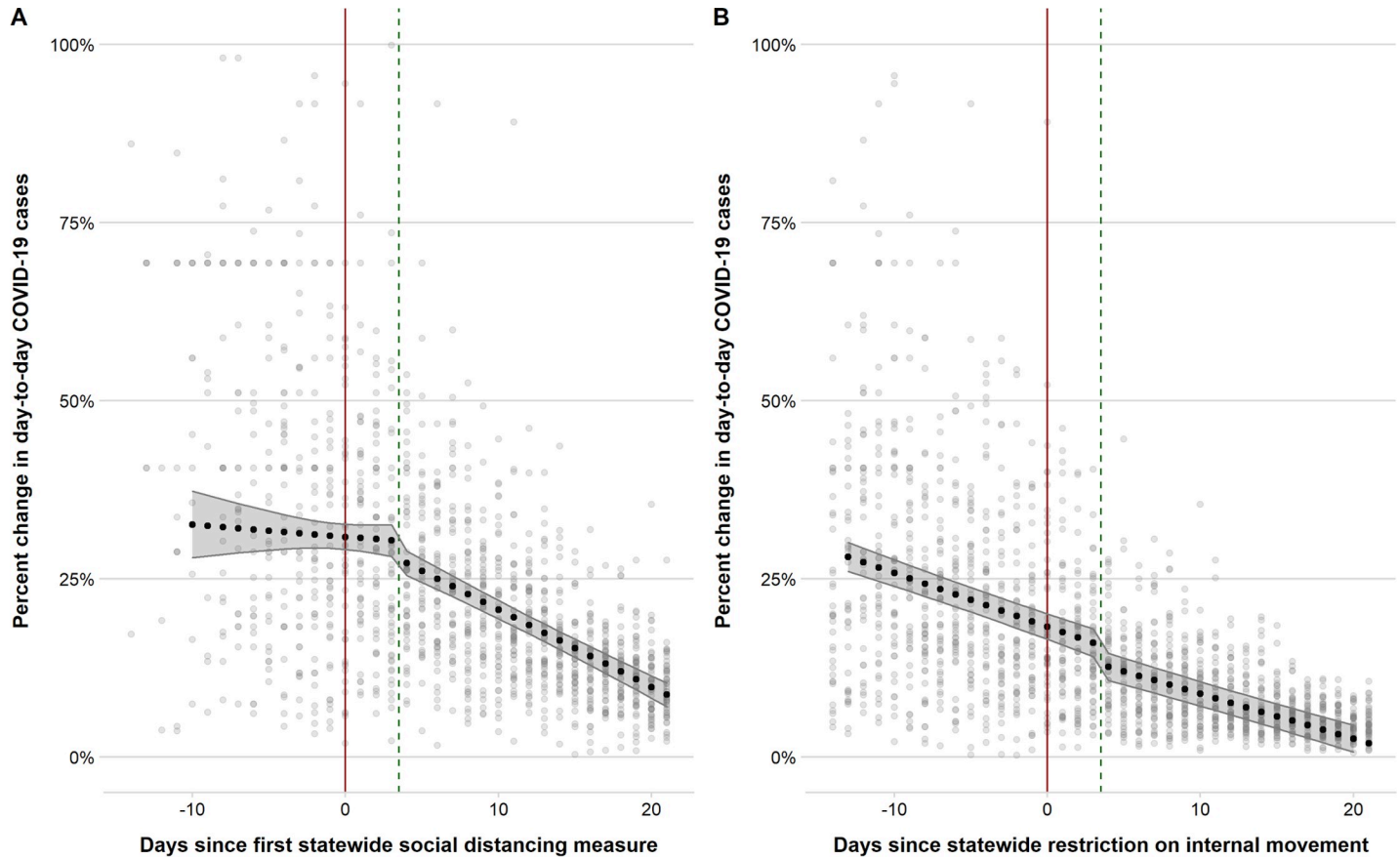


Fig 1. Scatter plots and predictive margins with 95% confidence interval derived from regression models of the daily COVID-19 growth rate before versus after implementation of the first statewide social distancing measures and statewide restrictions on internal movement. (A) First statewide social distancing measure. (B) Statewide restrictions on internal movement. The red line indicates the date of implementation in each state. The green dashed line is 4 days after implementation of the social distancing measure. COVID-19, coronavirus disease 2019.

<https://doi.org/10.1371/journal.pmed.1003244.g001>

(IQR 8–15) after the first statewide social distancing measure was implemented in the respective states, when the median statewide epidemic size was 937 cases (IQR 225–1,414). The mean daily case growth rate was already declining, at a mean rate of -0.8% per day, during the 14 days prior to implementation of statewide restrictions on internal movement (95% CI -0.9% to -0.7% ; $P < 0.001$) (Table 1; Fig 1B). There was a drop detected 3 days after statewide restrictions on internal movement were implemented (-3.1% ; 95% CI -4.7% to -1.5% ; $P <$

Table 1. Linear regression models for the growth in mean daily COVID-19 cases before versus after implementation of the first statewide social distancing measure and statewide restrictions on internal movement.

Variable	First statewide social distancing measure			Statewide restrictions on internal movement		
	<i>b</i>	95% CI	<i>P</i> value	<i>b</i>	95% CI	<i>P</i> value
Constant	0.309	0.291, 0.326	<0.001	0.183	0.164, 0.201	<0.001
Time (days relative to intervention)	-0.002	-0.006, 0.003	0.44	-0.008	-0.009, -0.006	<0.001
Post-intervention period	0.006	-0.017, 0.029	0.59	-0.031	-0.047, -0.015	<0.001
Time × post-intervention period	-0.009	-0.014, -0.004	<0.001	0.001	-0.0004, 0.003	0.14

The post-intervention period begins 4 days after social distancing measures were implemented, to account for the disease incubation period. *b*, estimated regression coefficient. CI, confidence interval; COVID-19, coronavirus disease 2019.

<https://doi.org/10.1371/journal.pmed.1003244.t001>

Table 2. Linear regression models for growth in mean daily COVID-19-attributed deaths before versus after implementation of the first statewide social distancing measure and statewide restrictions on internal movement, assuming a range of expected days between symptom onset and death.

Time after implementation at which post-intervention period begins	First statewide social distancing measure			Statewide restriction on internal movement		
	<i>b</i>	95% CI	<i>P</i> value	<i>b</i>	95% CI	<i>P</i> value
4 days	-0.018	-0.033, -0.002	0.02	-0.005	-0.009, -0.002	0.005
7 days	-0.020	-0.030, -0.009	<0.001	-0.002	-0.006, 0.002	0.29
10 days	-0.010	-0.021, 0.015	0.09	-0.000	-0.002, 0.002	0.98
14 days	0.001	-0.004, 0.007	0.78	0.002	0.000, 0.004	0.02
17 days	0.001	-0.003, 0.006	0.53	0.003	0.001, 0.006	0.001

Each row corresponds to a separate regression model fitted to the data that includes 3 variables: time in days, implementation period, and a time-by-implementation-period product term. The start of the post-intervention period specified in the regression models varies from 4 to 17 days after social distancing measures were implemented, to account for the disease incubation period as well as uncertainty in the expected time from symptom onset to death.

b, estimated regression coefficient; CI, confidence interval; COVID-19, coronavirus disease 2019.

<https://doi.org/10.1371/journal.pmed.1003244.t002>

0.001), but no statistically significant difference in the rate of change before versus after implementation (0.1% per day; 95% CI -0.04% to 0.3%; $P = 0.14$). As discussed in more detail below, there is substantial difficulty in disentangling the unique associations with statewide restrictions on internal movement from the unique associations with the first social distancing measures.

In the analysis of the secondary outcome, change in daily COVID-19-attributed deaths, given the uncertainty in the hypothesized lag between implementation of social distancing and observed changes (if any) in daily COVID-19-attributed deaths, we explored a range of lag times. As shown in Table 2, by 7 days after implementation of the first statewide social distancing measure, the mean daily growth rate in COVID-19-attributed deaths decreased by 2.0% per day (95% CI -3.0% to -0.9%; $P < 0.001$). By 14 days, the estimated association was no longer statistically significant (-1.0% per day; 95% CI -0.2% to 0.1%; $P = 0.09$). No additional statistically significant benefit was found after 7 days after implementation of statewide restrictions on internal movement.

Sensitivity analyses suggested our estimates were not sensitive to inclusion of additional covariates, did not differ by the size of the epidemic at implementation, and were consistent with the known incubation period (Tables B, C, and D in S1 Text). In the event study specification, mean daily case growth was negative by day 4, and the estimates were statistically significant by day 8, consistent with the primary analysis (Fig B in S1 Text). The event study analysis for change in daily COVID-19-attributed deaths also produced estimates qualitatively similar to the primary analysis, although with slightly larger CIs given the smaller number of events (Fig C in S1 Text).

Discussion

In this longitudinal pretest–posttest comparison group study, we found that implementation of social distancing measures was associated with a reduction in the mean daily growth rate of COVID-19 cases and in the mean daily growth rate of COVID-19-attributed deaths. Our estimates imply a more than doubling in the doubling time (from 3.8 days to 8.0 days) by 3 weeks following the implementation of social distancing measures. Assuming a cumulative epidemic size of 4,125 reported cases (equivalent to the cumulative number of cases in the US at the time of implementation in each state), the reduction in growth rate we estimated corresponds to a difference between 26,281 reported cases with no social distancing versus 24,625 reported cases with social distancing, at 7 days after implementation; a difference between 158,518

reported cases with no social distancing versus 102,223 reported cases with social distancing, at 14 days after implementation; and a difference between 904,773 reported cases with no social distancing versus 283,161 reported cases with social distancing, at 21 days after implementation. Stated differently, our model implies that social distancing reduced the total number of reported COVID-19 cases by approximately 1,600 cases at 7 days after implementation, by approximately 56,000 reported cases at 14 days after implementation, and by approximately 621,000 reported cases at 21 days after implementation. It can be inferred that earlier implementation of social distancing measures would likely have reduced morbidity and mortality even further.

These results are consistent with both the theoretical effect of social distancing on epidemic spread [6] and the historical benefit observed with the implementation of such interventions during prior epidemics of communicable diseases [28]. They also are largely in keeping with recent data on the impacts of social distancing measures in the US on both mobility [7,8] and case growth rates [9–11], with generally similar effect sizes. Our study extends this literature by further examining COVID-19-attributed mortality as an outcome. The association between social distancing and case growth rate was most apparent at the lower bound of the incubation period that has been estimated based on publicly available data, with some evidence that the change in growth rate may have started even earlier. We suspect that this observation may have resulted from self-imposed social distancing, which reportedly occurred prior to government-issued mandates [33].

Our findings should be interpreted with the following limitations in mind. Our estimates would be biased toward the null if (1) state and local governments intensified social distancing measures in response to a worsening epidemic, (2) there were substantial violations of the stable unit treatment value assumption (e.g., workplace closures of large employers that had spill-over effects across state lines), or (3) surveillance and testing intensified during the study period (thereby resulting in increased case reporting). Moreover, statewide restrictions on internal movement were often implemented after other social distancing measures had already been applied, further biasing our estimate toward the null. Estimates of cases and deaths in our model include both those that are laboratory-confirmed and those that are suspected by health departments, but both are likely to be underestimates due to limitations in testing, the presence of asymptomatic cases, and the occurrence of COVID-19-related deaths that are not attributed to COVID-19 [34,35]. Nonetheless, our analyses focus on day-to-day changes in the growth rate of cases and deaths, so underreporting would only bias our results if reported versus true outcomes systematically differed prior to versus after the implementation of social distancing measures. In contrast, our projected estimates of cases prevented are likely to be highly conservative, because they are modeled based on reported cases. While some studies have suggested that cases have been underreported in the US by 1–2 orders of magnitude [36], deaths appear to be undercounted by approximately 25% [34]. Finally, our estimate of the association between social distancing and change in mean daily case growth rate (and the corresponding number of cases averted) cannot be extrapolated linearly beyond the 21-day post-implementation analysis window. At the time of this analysis, states had begun to relax social distancing measures, which necessarily prevents drawing conclusions about the long-term associational effects of these interventions in isolation.

Our study does not ascertain which of the statewide social distancing measures were most effective in reducing mean daily COVID-19 case growth. A variety of social distancing measures were used, often simultaneously, making it difficult to disentangle their independent associations. It is also possible that some state residents changed their behaviors in response to local (e.g., county-level) social distancing measures enacted prior to the statewide measures, or that some state residents changed their behaviors independently of social distancing measures enacted at any level of jurisdiction. The latter phenomenon has been observed empirically in

other settings [37,38]. Such behavior change would be consistent with our sensitivity analysis suggesting a signal that extends back to a 0-day incubation period.

Finally, our analysis cannot answer questions about the appropriate time for rescinding social distancing measures. While our findings demonstrate that early mitigation efforts have yielded a substantial population health benefit, such benefits should be weighed against their costs. The costs of social distancing are likely to exacerbate the confluence of longstanding economic, social, and health decline that is already occurring in the US [39–41], brought into even sharper relief given emerging data about racial, ethnic, and socioeconomic disparities in the incidence of COVID-19-related burden [42,43]. Moreover, there is currently robust debate about the extent to which legal authority to initiate or rescind social distancing measures resides with the federal or local government [44]. Additional study may be possible in the coming weeks to months if some statewide social distancing measures are relaxed while others are retained, or if measures are relaxed in some local jurisdictions while being retained in others. The use of cross-country data may also be helpful in this regard, albeit limited by any substantial within-country heterogeneity (such as has been observed in the diversity of uncoordinated US federal and state responses to the epidemic).

Our finding that implementation of statewide social distancing was associated with a reduction in the mean daily growth rate of COVID-19-attributed deaths should be interpreted with more caution, given the uncertainty in published estimates of the median time from symptom onset to death [19–21]. The strongest association we estimated, in terms of both statistical significance and magnitude, occurred at 7 days after implementation. While this association could be reflective of what is known about time to death for the median hospitalized patient, other plausible lag times were not associated with a statistically significant reduction in the mean daily growth rate of COVID-19-attributed deaths.

The finding that implementation of statewide restrictions on internal movement was associated neither with a statistically significant reduction in the mean daily growth rate of COVID-19 cases nor with a statistically significant reduction in the mean daily growth rate of COVID-19-attributed deaths warrants additional scrutiny. Identification of the unique effect of statewide restrictions on internal movement, separate from that of the initial social distancing measures, is a difficult undertaking given that restrictions on internal movement were implemented a median of 10 days later. There were no states in which statewide restrictions on internal movement were implemented without the prior implementation of other social distancing measures (e.g., cancellations of public events and closures of schools). As a result, we could not test hypotheses about the independent causal effects of statewide restrictions on internal movement, but rather could only ascertain the extent to which these restrictions were associated with additional reductions in the COVID-19 case growth rate, i.e., beyond those generated by the initial statewide social distancing measures. Moreover, the null finding should be interpreted in light of the other likely biases toward the null previously discussed.

In summary we demonstrate that the US COVID-19 epidemic growth rate began to decline within approximately 1 incubation period following the initiation of statewide social distancing measures. Exploratory findings suggest that social distancing may also have had lagged mortality benefits. Future work should consider the potential public health benefits of relaxing these measures and, specifically, to what extent and how differential relaxation of certain measures promotes persistent epidemic control.

Supporting information

S1 STROBE Checklist.

(DOC)

S1 Text.
(DOCX)

Author Contributions

Conceptualization: Mark J. Siedner, Guy Harling, Sebastien Haneuse, Atheendar S. Venkataramani, Alexander C. Tsai.

Data curation: Mark J. Siedner, Guy Harling, Zahra Reynolds, Rebecca F. Gilbert, Atheendar S. Venkataramani, Alexander C. Tsai.

Formal analysis: Mark J. Siedner, Guy Harling, Rebecca F. Gilbert, Sebastien Haneuse, Atheendar S. Venkataramani, Alexander C. Tsai.

Funding acquisition: Guy Harling, Alexander C. Tsai.

Investigation: Mark J. Siedner, Guy Harling, Zahra Reynolds, Rebecca F. Gilbert, Sebastien Haneuse, Atheendar S. Venkataramani, Alexander C. Tsai.

Methodology: Mark J. Siedner, Guy Harling, Sebastien Haneuse, Atheendar S. Venkataramani, Alexander C. Tsai.

Supervision: Mark J. Siedner, Guy Harling, Alexander C. Tsai.

Visualization: Zahra Reynolds, Rebecca F. Gilbert.

Writing – original draft: Mark J. Siedner, Guy Harling.

Writing – review & editing: Mark J. Siedner, Guy Harling, Zahra Reynolds, Rebecca F. Gilbert, Sebastien Haneuse, Atheendar S. Venkataramani, Alexander C. Tsai.

References

1. Kissler SM, Tedijanto C, Goldstein E, Grad YH, Lipsitch M. Projecting the transmission dynamics of SARS-CoV-2 through the postpandemic period. *Science*. 2020; 368(6493):860–8. <https://doi.org/10.1126/science.abb5793> PMID: 32291278
2. Markel H, Lipman HB, Navarro JA, Sloan A, Michalsen JR, Stern AM, et al. Nonpharmaceutical interventions implemented by US cities during the 1918–1919 influenza pandemic. *JAMA*. 2007; 298(6):644–54. <https://doi.org/10.1001/jama.298.6.644> PMID: 17684187
3. Bootsma MC, Ferguson NM. The effect of public health measures on the 1918 influenza pandemic in U.S. cities. *Proc Natl Acad Sci U S A*. 2007; 104(18):7588–93. <https://doi.org/10.1073/pnas.0611071104> PMID: 17416677
4. Ferguson NM, Cummings DA, Cauchemez S, Fraser C, Riley S, Meeyai A, et al. Strategies for containing an emerging influenza pandemic in Southeast Asia. *Nature*. 2005; 437(7056):209–14. <https://doi.org/10.1038/nature04017> PMID: 16079797
5. Taubenberger JK, Morens DM. 1918 influenza: the mother of all pandemics. *Emerg Infect Dis*. 2006; 12(1):15–22. <https://doi.org/10.3201/eid1201.050979> PMID: 16494711
6. Teslya A, Pham TM, Godijk NG, Kretzschmar ME, Bootsma MCJ, Rozhnova G. Impact of self-imposed prevention measures and short-term government-imposed social distancing on mitigating and delaying a COVID-19 epidemic: a modelling study. *PLoS Med* 17(7): e1003166. <https://doi.org/10.1371/journal.pmed.1003166>
7. Andersen M. Early evidence on social distancing in response to COVID-19 in the United States. SSRN Working Paper No. 3569368 [preprint]. Amsterdam: Elsevier; 2020 [cited 2020 Jun 2]. Available from: <https://ssrn.com/abstract=3569368>.
8. Abouk R, Heydari B. The immediate effect of COVID-19 policies on social distancing behavior in the United States. medRxiv [preprint]. 2020 Apr 28. <https://doi.org/10.1101/2020.04.07.20057356>
9. Courtemanche C, Garuccio J, Le A, Pinkston J, Yelowitz A. Strong social distancing measures in the United States reduced the COVID-19 growth rate. *Health Aff (Millwood)*. 2020 May 14. <https://doi.org/10.1377/hlthaff.2020.00608> PMID: 32407171

10. Dave DM, Friedson AI, Matsuzawa K, Sabia JJ. When do shelter-in-place orders fight COVID-19 best? Policy heterogeneity across states and adoption time. NBER Working Paper No. 27091 [preprint]. Cambridge: National Bureau of Economic Research; 2020 [cited 2020 Jun 2]. Available from: <https://www.nber.org/papers/w27091>.
11. Hsiang S, Allen D, Annan-Phan S, Bell K, Bolliger I, Chong T, et al. The effect of large-scale anti-contagion policies on the COVID-19 pandemic. *Nature*. 2020 Jun 8. <https://doi.org/10.1038/s41586-020-2404-8> PMID: 32512578
12. Bayham J, Fenichel EP. Impact of school closures for COVID-19 on the US health-care workforce and net mortality: a modelling study. *Lancet Public Health*. 2020; 5(5):e271–78. [https://doi.org/10.1016/S2468-2667\(20\)30082-7](https://doi.org/10.1016/S2468-2667(20)30082-7) PMID: 32251626
13. Eichenbaum MS, Rebelo S, Trabandt M. The macroeconomics of epidemics. NBER Working Paper No. 26882 [preprint]. Cambridge: National Bureau of Economic Research; 2020 [cited 2020 Jun 2]. Available from: <https://www.nber.org/papers/w26882>.
14. Petherick A, Hale T, Phillips T, Webster S. Variation in government responses to COVID-19. Blavatnik School Working Paper [preprint]. Oxford: University of Oxford Blavatnik School of Government; 2020 [cited 2020 Jun 2]. Available from: <https://www.bsg.ox.ac.uk/research/publications/variation-government-responses-covid-19>.
15. Council of State and Territorial Epidemiologists. Standardized surveillance case definition and national notification for 2019 novel coronavirus disease (COVID-19). Washington (DC): Office of the Assistant Secretary for Preparedness and Response; 2020 [cited 2020 Jun 2]. Available from: <https://asprtracie.hhs.gov/technical-resources/resource/8322/standardized-surveillance-case-definition-and-national-notification-for-2019-novel-coronavirus-disease-covid-19>.
16. 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. *Ann Intern Med*. 2020; 172(9):577–82. <https://doi.org/10.7326/M20-0504> PMID: 32150748
17. Backer JA, Klinkenberg D, Wallinga J. Incubation period of 2019 novel coronavirus (2019-nCoV) infections among travellers from Wuhan, China, 20–28 January 2020. *Euro Surveill*. 2020; 25(5):2000062.
18. Guan WJ, Ni ZY, Hu Y, Liang WH, Ou CQ, He JX, et al. Clinical characteristics of coronavirus disease 2019 in China. *N Engl J Med*. 2020; 382(18):1708–20. <https://doi.org/10.1056/NEJMoa2002032> PMID: 32109013
19. COVID-19 Surveillance Group. Characteristics of COVID-19 patients dying in Italy. Report based on available data on March 20th, 2020 [preprint]. Rome: Istituto Superiore di Sanità; 2020 [cited 2020 Apr 12]. Available from: https://www.epicentro.iss.it/coronavirus/bollettino/Report-COVID-2019_20_marzo_eng.pdf.
20. Zhou F, Yu T, Du R, Fan G, Liu Y, Liu Z, et al. Clinical course and risk factors for mortality of adult inpatients with COVID-19 in Wuhan, China: a retrospective cohort study. *Lancet*. 2020; 395(10229):1054–62. [https://doi.org/10.1016/S0140-6736\(20\)30566-3](https://doi.org/10.1016/S0140-6736(20)30566-3) PMID: 32171076
21. Verity R, Okell LC, Dorigatti I, Winskill P, Whittaker C, Imai N, et al. Estimates of the severity of coronavirus disease 2019: a model-based analysis. *Lancet Infect Dis*. 2020; 20(6):669–77. [https://doi.org/10.1016/S1473-3099\(20\)30243-7](https://doi.org/10.1016/S1473-3099(20)30243-7) PMID: 32240634
22. Wu JT, Leung K, Bushman M, Kishore N, Niehus R, de Salazar PM, et al. Estimating clinical severity of COVID-19 from the transmission dynamics in Wuhan, China. *Nat Med*. 2020; 26(4):506–10. <https://doi.org/10.1038/s41591-020-0822-7> PMID: 32284616
23. Siedner MJ, Harling G, Reynolds Z, Gilbert RF, Venkataramani A, Tsai AC. Social distancing to slow the U.S. COVID-19 epidemic: an interrupted time-series analysis. medRxiv [preprint]. 2020 Apr 8. <https://doi.org/10.1101/2020.04.03.20052373>
24. Lee VJ, Aguilera X, Heymann D, Wilder-Smith A, Lancet Infectious Diseases Commission. Preparedness for emerging epidemic threats: a Lancet Infectious Diseases Commission. *Lancet Infect Dis*. 2020; 20(1):17–9. [https://doi.org/10.1016/S1473-3099\(19\)30674-7](https://doi.org/10.1016/S1473-3099(19)30674-7) PMID: 31876487
25. Grassly NC, Fraser C. Mathematical models of infectious disease transmission. *Nature Rev Microbiol*. 2008; 6(6):477–87.
26. Lewis MD, Pavlin JA, Mansfield JL, O'Brien S, Boomsma LG, Elbert Y, et al. Disease outbreak detection system using syndromic data in the greater Washington DC area. *Am J Prev Med*. 2002; 23(3):180–6. [https://doi.org/10.1016/s0749-3797\(02\)00490-7](https://doi.org/10.1016/s0749-3797(02)00490-7) PMID: 12350450
27. Buckingham-Jeffery E, Morbey R, House T, Elliot AJ, Harcourt S, Smith GE. Correcting for day of the week and public holiday effects: improving a national daily syndromic surveillance service for detecting public health threats. *BMC Public Health*. 2017; 17(1):477. <https://doi.org/10.1186/s12889-017-4372-y> PMID: 28525991

28. Hatchett RJ, Mecher CE, Lipsitch M. Public health interventions and epidemic intensity during the 1918 influenza pandemic. *Proc Natl Acad Sci U S A*. 2007; 104(18):7582–7. <https://doi.org/10.1073/pnas.0610941104> PMID: 17416679
29. Sullivan D, von Wachter T. Job displacement and mortality: an analysis using administrative data. *Q J Econ*. 2009; 124(3):1265–306.
30. Jacobson L, LaLonde RJ, Sullivan DG. Earnings losses of displaced workers. *Am Econ Rev*. 1993; 83(4):685–709.
31. Goodman-Bacon A. Difference-in-differences with variation in treatment timing. NBER Working Paper No. 25018 [preprint]. Cambridge: National Bureau of Economic Research; 2018 [cited 2020 Jun 2]. Available from: <https://www.nber.org/papers/w25018>.
32. Bertrand M, Dufló E, Mullainathan S. How much should we trust differences-in-differences estimates? *Q J Econ*. 2004; 119(1):249–75.
33. Brzezinski A, Deiana G, Kecht V, Van Dijke D. The COVID-19 pandemic: government versus community action across the United States. *COVID Econ*. 2020; 7:115–156. Available from: <https://cepr.org/sites/default/files/news/CovidEconomics7.pdf>.
34. Weinberger DM, Chen J, Cohen T, Crawford FW, Mostashari F, Olson D, et al. Estimation of excess deaths associated with the COVID-19 pandemic in the United States, March to May 2020. *JAMA Intern Med*. 2020 Jul 1. <https://doi.org/10.1001/jamainternmed.2020.3391> PMID: 32609310
35. Rivera R, Rosenbaum J, Quispe W. Excess mortality in the United States during the peak of the COVID-19 pandemic. *medRxiv* [preprint]. 2020 Jun 27. <https://doi.org/10.1101/2020.05.04.20090324>
36. Lu FS, Nguyen AT, Link N, Santillana M. Estimating the prevalence of COVID-19 in the United States: three complementary approaches. *medRxiv* [preprint]. 2020 Jun 18. <https://doi.org/10.1101/2020.04.18.20070821> PMID: 32587997
37. Cowling BJ, Ali ST, Ng TWY, Tsang TK, Li JCM, Fong MW, et al. Impact assessment of non-pharmaceutical interventions against coronavirus disease 2019 and influenza in Hong Kong: an observational study. *Lancet Public Health*. 2020; 5(5):e279–88. [https://doi.org/10.1016/S2468-2667\(20\)30090-6](https://doi.org/10.1016/S2468-2667(20)30090-6) PMID: 32311320
38. Lewnard JA, Liu VX, Jackson ML, Schmidt MA, Jewell BL, Flores JP, et al. Incidence, clinical outcomes, and transmission dynamics of severe coronavirus disease 2019 in California and Washington: prospective cohort study. *BMJ*. 2020; 369:m1923. <https://doi.org/10.1136/bmj.m1923> PMID: 32444358
39. Case A, Deaton A. Deaths of despair and the future of capitalism. Princeton: Princeton University Press; 2020.
40. Venkataramani AS, Bair EF, O'Brien RL, Tsai AC. Association between automotive assembly plant closures and opioid overdose mortality in the United States: a difference-in-differences analysis. *JAMA Intern Med*. 2020; 180(2):254–62.
41. Muennig PA, Reynolds M, Fink DS, Zafari Z, Geronimus AT. America's declining well-being, health, and life expectancy: not just a white problem. *Am J Public Health*. 2018; 108(12):1626–31. <https://doi.org/10.2105/AJPH.2018.304585> PMID: 30252522
42. Chin T, Kahn R, Li R, Chen JT, Krieger N, Buckee CO, et al. U.S. county-level characteristics to inform equitable COVID-19 response. *medRxiv* [preprint]. 2020 Apr 11. <https://doi.org/10.1101/2020.04.08.20058248> PMID: 32511610
43. Unequal protection: American inequality meets covid-19. *The Economist*. 2020 Apr 18.
44. Fritze J. Authority to reopen U.S. for business in question. *USA Today*. 2020 Apr 14.