

SUPPLEMENT: simulation model of causal pathways

In the main text results, we set the model tuning parameters a , b , and c to 1. Here we vary these parameters and examine model consistency with Zhou et al. (2020) data (Figure S1).

We simulate cohorts of 1000 hospitalized COVID-19 positive cases under a range of a, b, c scenarios, obtaining distributions for four possible outcomes: survivors with bacterial infection, survivors without bacterial infection, non-survivors with bacterial infection, and non-survivors without. We compare our modeled distributions to data from the Zhou et al study. Combinations of parameter values that were consistent with the data (within a 95% CI) are coded as green dots (Figure S1).

This model fitting shows that no parameter other than b can equal zero. This confirms our ruling out of the pipe structure (where $a = 0$) and allows us to rule out the fork structure (where $c = 0$) as very unlikely to be the true causal structure. Together, our models support both a collider or multi-path causal structure underpinning bacterial-associated mortality in COVID-19.

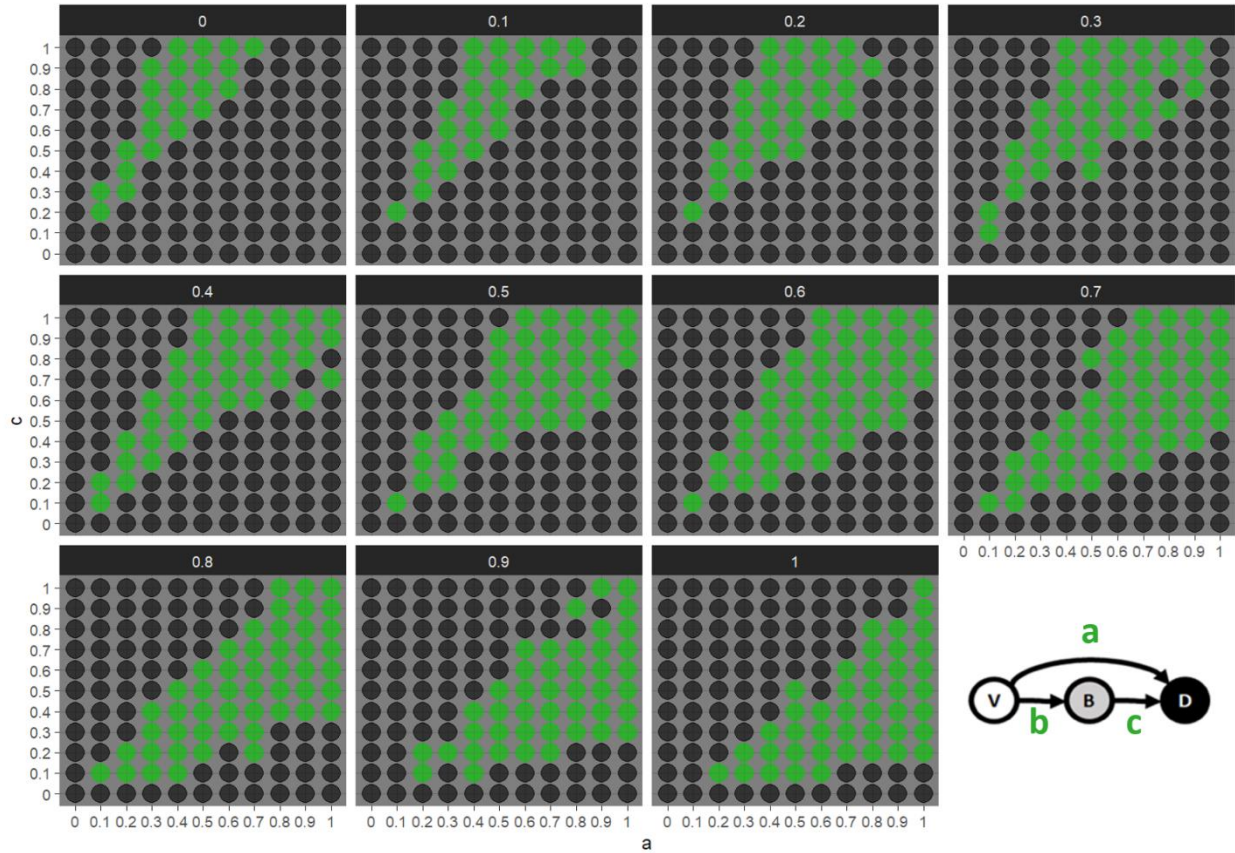


Figure S1. Fitting the general model. We fit a general model by introducing parameters a , b , and c to modulate the effect of $(V \rightarrow D)$, $(V \rightarrow B)$, and $(B \rightarrow D)$, respectively. For each combination of parameters a , b , and c , we run 100 simulations of cohorts of 1000 hospitalized patients each. For each cohort, obtaining distributions of the percentage of patients for each of four possible outcomes: survivors with bacterial infection, survivors without bacterial infection, non-survivors with bacterial infection, and non-survivors without. We then compare the observed percent distribution from the *Zhou et al* study to the simulated ones, and highlight where the observed percentage for each outcome falls within a 90% CI for the simulated distributions (green circles). Our model fitting shows that only b can equal zero and still be consistent with the observed infection and mortality data.