Update on COVID-19 Epidemiology and Management Strategies



Harvard T.H. Chan School of Public Health, United States

The pandemic is global



The pandemic is hyper-local

nd deaths counts. Risk Levels ases for actual number of cont earn more	are calculated base	d on daily cases per	es. Hover over a county 100,000 population (7 d	for detailed information on cases ay rolling average). See Daily New	State/County	Rank	Daily new cases per 100k people (7d moving avg.)	Daily new cases (7d moving avg.)
isk Levels by County					🕂 South Dakota	1	131.2	1,160.
d→ /){h⊔ ở ∖					🕂 North Dakota	2	116.2	885
					+ Wisconsin	3	79.8	4,643
				🕂 Montana	4	75.2	803	
					+ Wyoming	5	68.3	395
					+ Iowa	6	65.6	2,069
					+ Nebraska	7	57.5	1,112
		┝╌╞╍╌╌┼┶┫╌╸ ╕╴┓╸╴╴╴╴╴╴ ╞╧╌╕╹┝╌╌┝╼╴╤┽╴╞╧╤┶┵┿┷┷┷			🕂 Utah	8	53.6	1,719
					+ Idaho	9	51.1	913
					🕂 Alaska	10	51.1	373
					+ Illinois	11	48.5	6,149
					+ Minnesota	12	46.7	2,633
					+ Kansas	13	42.2	1,229
			推击注意		+ Indiana	14	41.5	2,792
	N				+ Missouri	15	40.0	2,452
S. F.	1				+ Kentucky	16	37.4	1,671
Tayl and					+ Colorado	17	36.8	2,118
					+ New Mexico	18	36.6	767
					+ Arkansas	19	35.7	1,076
_					+ Rhode Island	20	33.8	357
Risk Levels:	Green	Yellow	Orange	Red	+ Michigan	21	31.2	3,113
Microsoft Al for Health				about Sources k Here			ai4hc19@m	Contact u

https://globalepidemics.org/key-metrics-for-covid-suppression/

Horrific racial/ethnic disparities

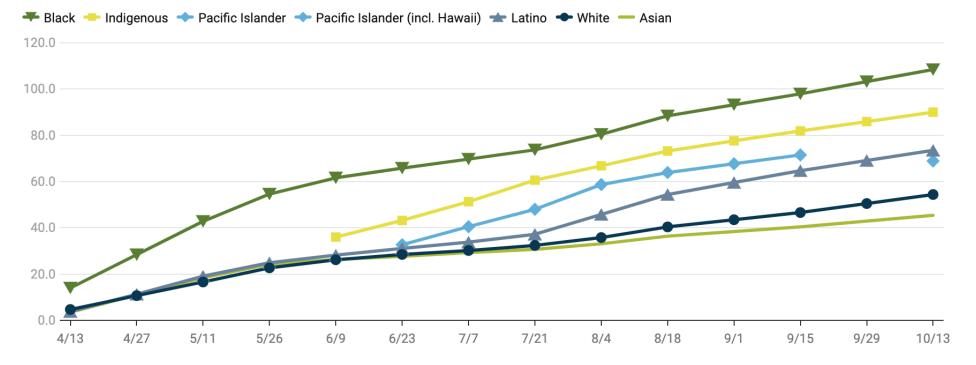
A P M RESEARCH LAB

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Black & Indigenous Americans experience highest death tolls from COVID-19

Cumulative actual COVID-19 mortality rates per 100,000, by race and ethnicity, April 13-Oct. 13, 2020



Note: All intervals are 14 days apart, except for 5/11-5/26, which is a 15-day period. 9/1 and 9/29 data has been interpolated. Pacific Islander data prior to 10/13 did not include Hawaii, as it was not releasing data; its inclusion resulted in an overall drop in the Pacific Islander rate, which begins a new series at 10/13. Source: APM Research Lab • Get the data • Created with Datawrapper

Spread has depended on individual & local responses and risk factors

New York City virus PCR in delivering mothers Maternal virus prevalence, Mar-Apr

Maternal virus prevalence

Reduction in movement

Correlation



Kissler et al. Nature Communications 2020

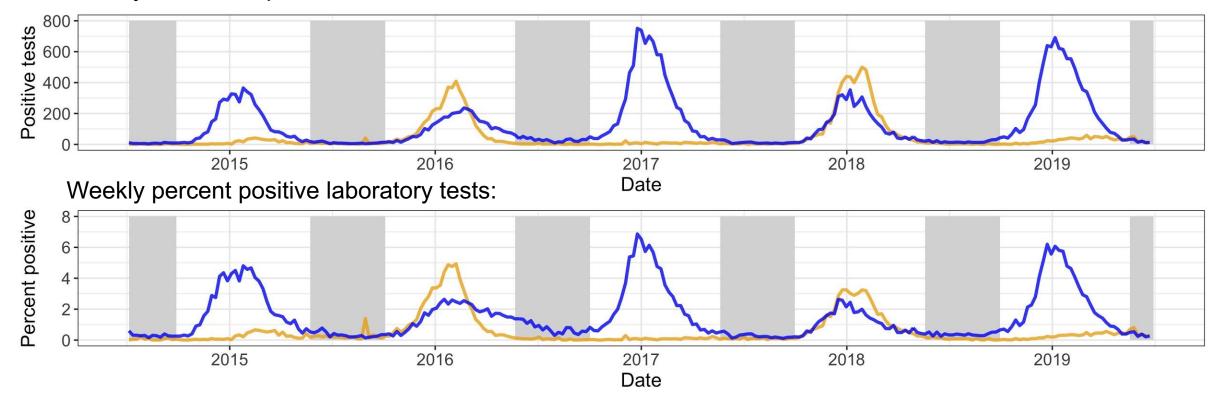
Where we left off...

Spring, 2020:

- Some places "crushed the curve" using a combination of lockdown, masking, and extensive testing, tracing, and isolation/quarantine
- Other places used lockdowns to "flatten the curve"
- Questions on what will happen in the summer and fall

Is there seasonality? What about for the other human coronaviruses?

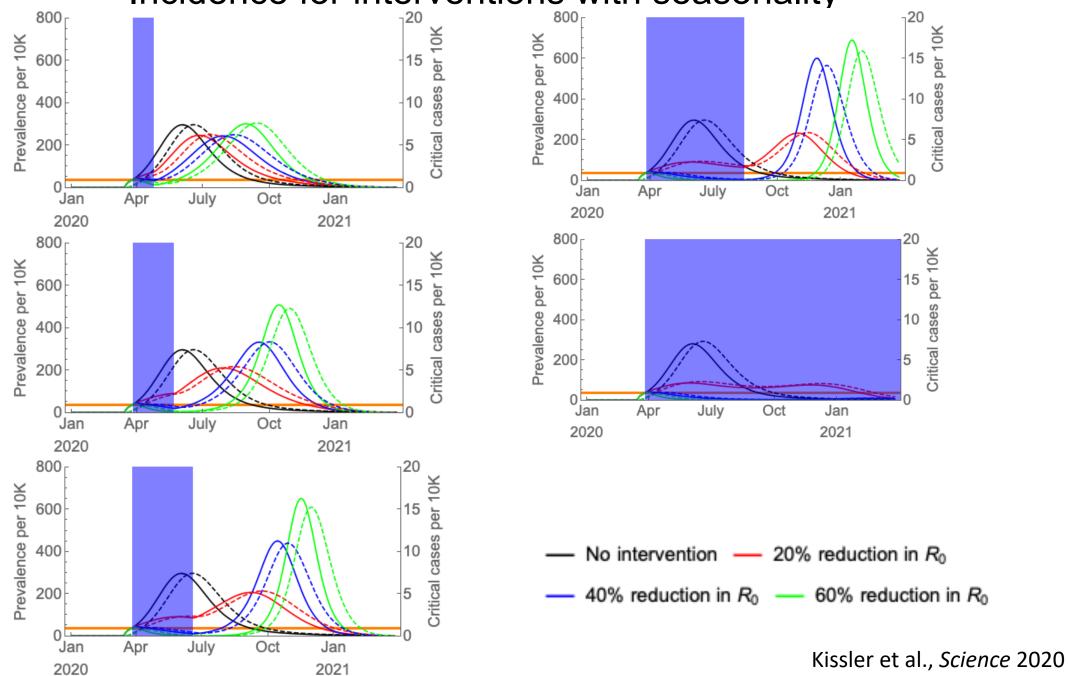
Weekly number of positive tests from NREVSS:



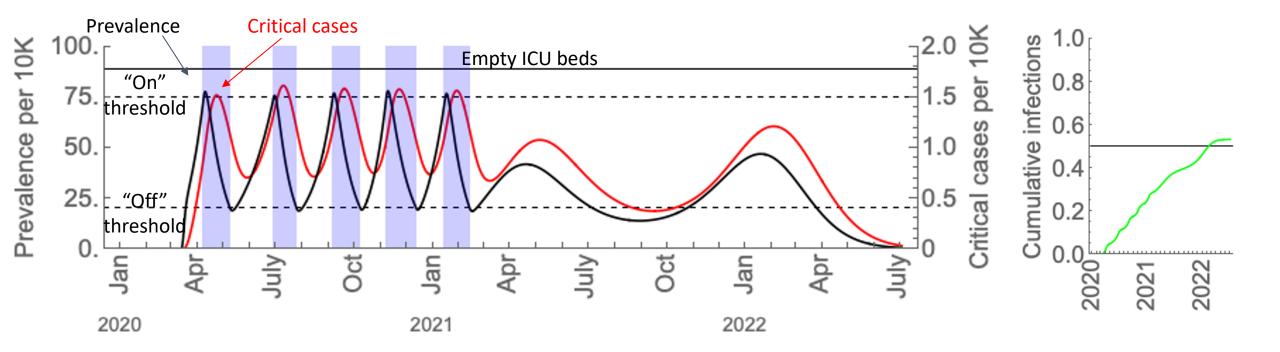
strain — CoVHKU1 — CoVOC43

Kissler et al., Science 2020

Incidence for interventions with seasonality



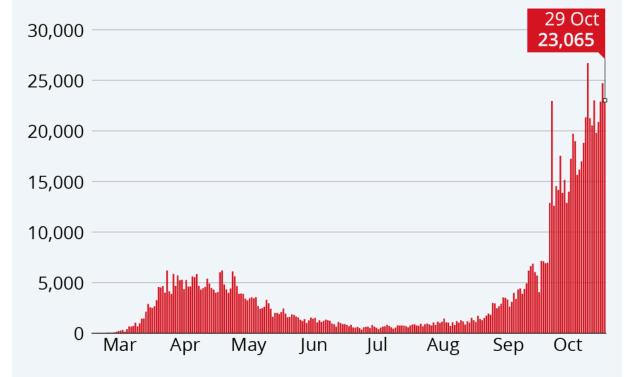
Cycles of lockdowns...



Example from the UK

Daily UK Covid-19 cases

Daily reported lab-confirmed Covid-19 cases in the United Kingdom^{*}



* Figures represent the date on which cases were recorded, not the date of the test. Source: data.gov.uk

Example from Israel

Daily new confirmed COVID-19 cases

The number of confirmed cases is lower than the number of actual cases; the main reason for that is limited testing.







"Reduce. Relax. Repeat"



Replying to @kakape

Europe meanwhile is using lockdowns to avoid health care system from collapsing. The strategy if there is one seems to be: Reduce numbers to manageable levels, then relax measures. I call it the 3Re ,strategy': Reduce. Relax. Repeat.

6:26 AM · Nov 3, 2020 · Twitter for iPhone

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Other examples:

Mainland China Vietnam Taiwan South Korea Australia

So what do we do now?

- Repeated lockdowns, masking, distancing
- Testing, testing, testing... together with supported isolation and quarantine
- Await effective vaccines and therapeutics

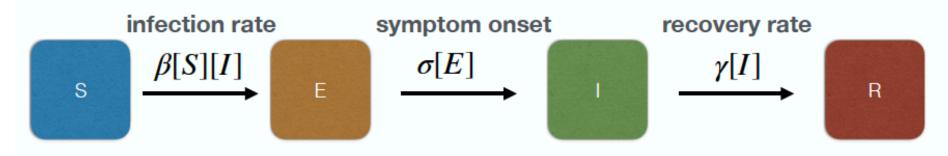
Model-informed COVID-19 vaccine prioritization strategies by age and serostatus

Kate M. Bubar,^{1,2}* Stephen M. Kissler,³ Marc Lipsitch^{3,4}, Sarah Cobey⁵, Yonatan H. Grad³, Daniel B. Larremore^{6,7}*

The model for SARS-CoV-2:

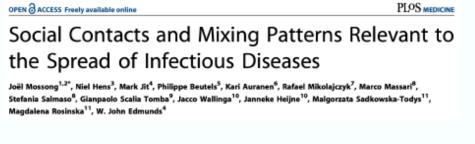
People move between the compartments of this "compartmental model":

Susceptible Exposed Infected Recovered

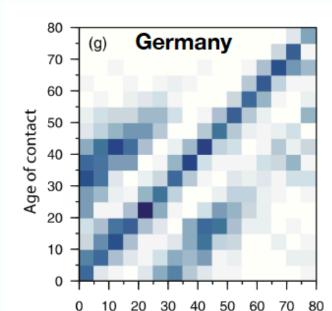


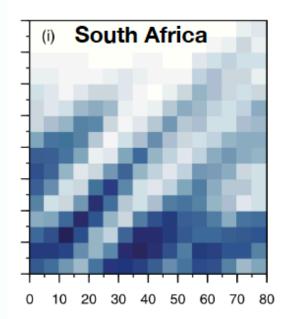
But in this kind of model, everyone is the same. We need more structure!

Stratified compartmental models e.g. POLYMOD-type age-structured SEIR models



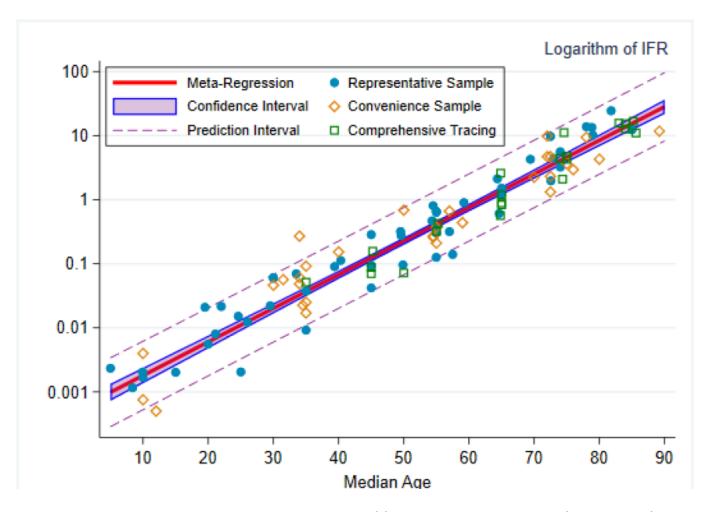
The POLYMOD study and others like it have mapped age-contact structure.





Age-stratified SEIR models allow us to ask more targeted questions!

Infection-Fatality Ratio (IFR) increases exponentially with age



Levin, Meyerowitz-Katz, et al., medRxiv https://www.medrxiv.org/content/10.1101/2020.07.23.20160895v6

knowns:

1. The vaccine will initially be scarce.

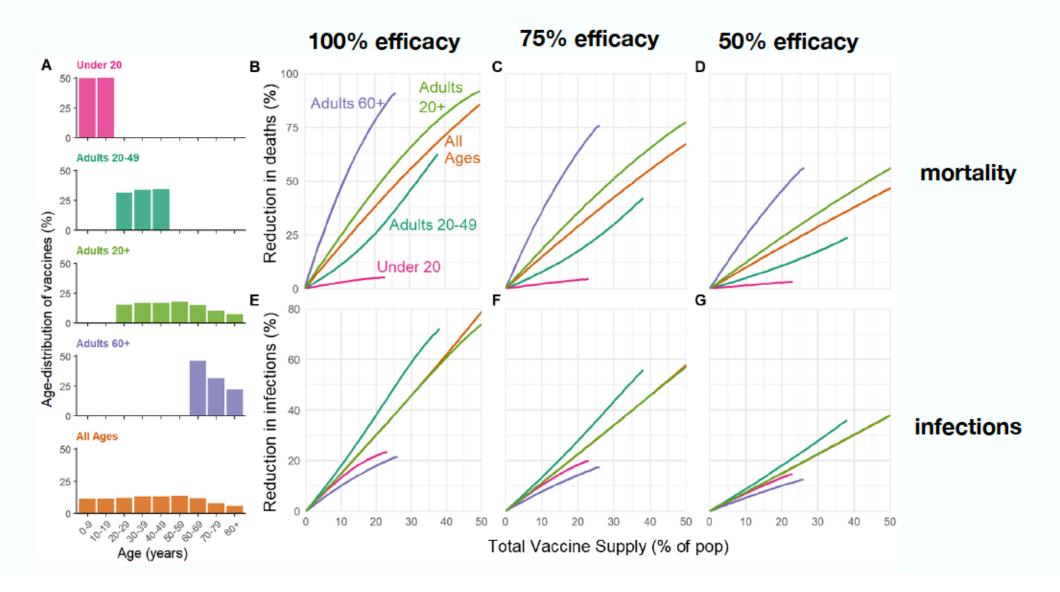
unknowns:

- 1. Safety: who is the vaccine approved for?
- 2. Efficacy: how protective is the vaccine?
- 3. Age-related effects: is the vaccine equally effective across ages?
- 4. Vax properties: transmission blocking?

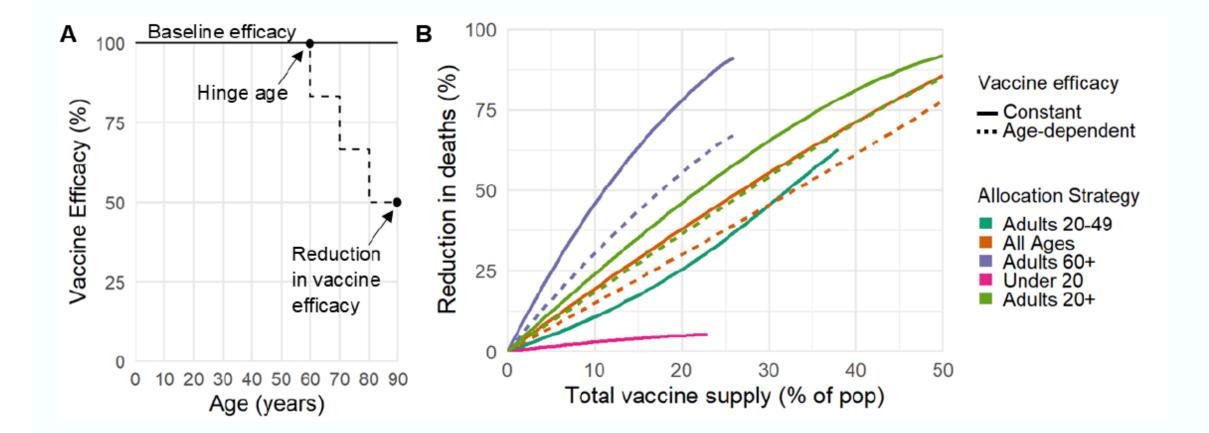
variables:

- 1. **Demographics:** what's the age distribution in the population?
- 2. Age-contact structure: are families multihousehold? Do people of all ages work? Strict retirement age?
- 3. Seroprevalence: what fraction of the population has antibodies already? And, do they correlate with protection?

How do different prioritizations play out?



What about variation in efficacy by age?

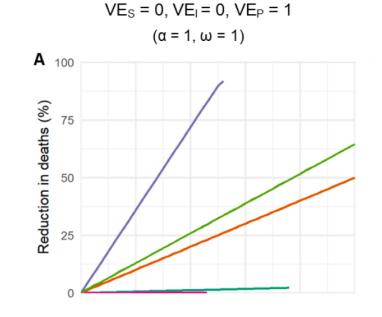


Prioritizing the elderly (similarly, comorbid) to reduce deaths is robust to:

Variations in efficacy by age

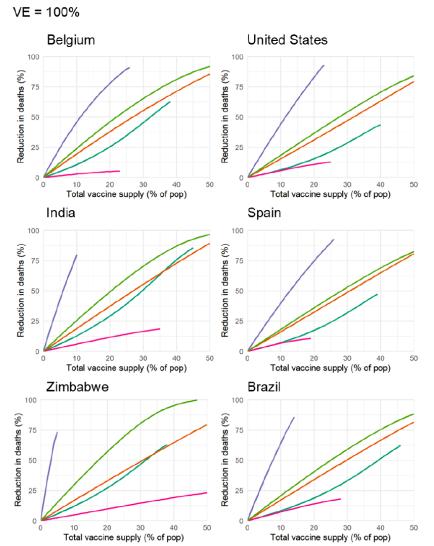
		Assuming an all-or-nothing vaccine.				
Baseline efficacy	Hinge Age	Tipping point when vaccine supply is:				
		5% of pop	15% of pop	25% of pop		
	59	-	-	-		
50%	69	-	-	-		
	79	-	-	-		
	59	-	-	0.8%		
75%	69	-	-	-		
	79	-	-	-		
	59	-	-	3.9%		
100%	69	-	-	-		
	79	-	-	-		

Vaccine that protects only against symptoms/death: no effect on infections



Prioritizing the elderly (similarly, comorbid) to reduce deaths is robust to:

Demography

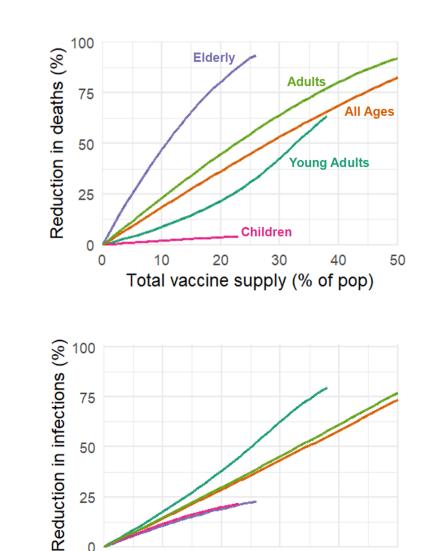


Also to leaky or all-or-nothing vaccine

Current model includes:

- →Contact patterns by age
- → Demographics
- →Susceptibility by age
- →Serology
- →Age-variation in vaccine effectiveness
 →IFR

Framework enables sensitivity testing. Next: vary *R*, the timing of vaccine roll-out



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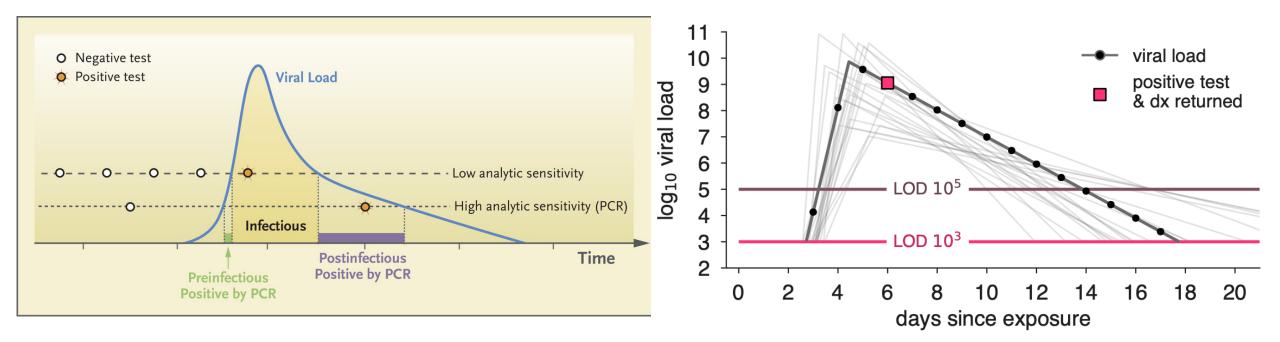
Total vaccine supply (% of pop)

30

40

50

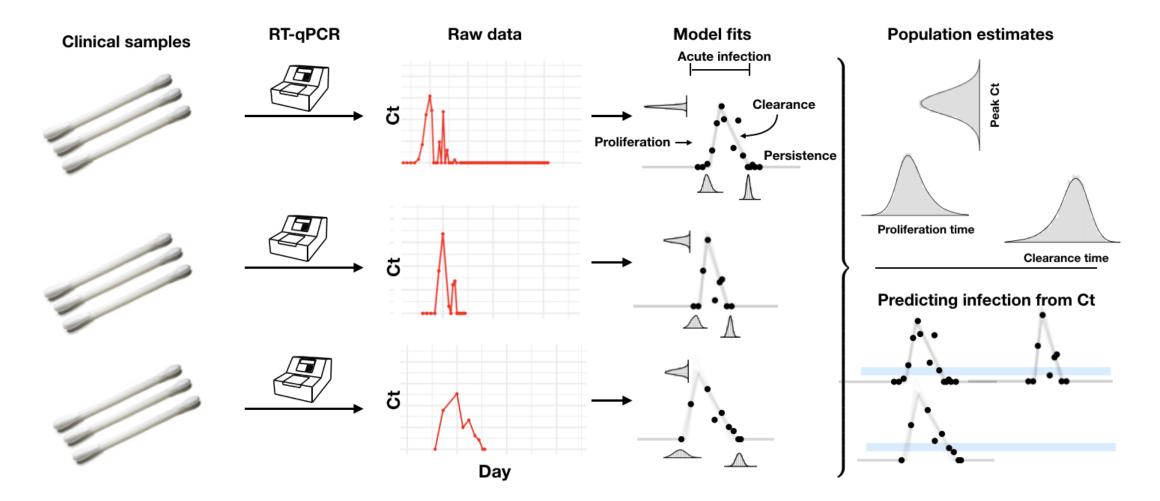
Charting the course of SARS-CoV-2 infection



Mina, Parker, Larremore. *NEJM*. 2020

Larremore et al., Science Advances. In press.

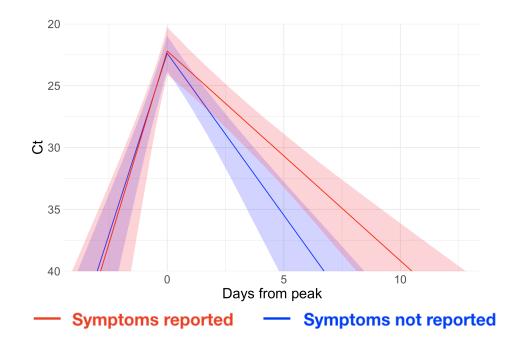
The quantitative prospective longitudinal NBA testing enables estimation of the full viral trajectory

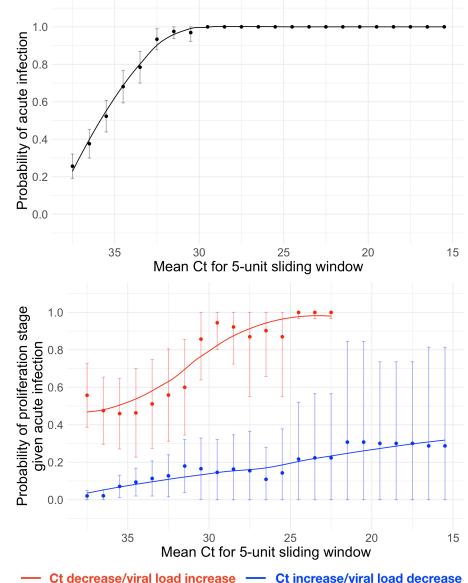


Data from 68 individuals, 46 with acute infections, 13 of whom reported symptoms

Kissler et al., *medRxiv* 2020

Quantitative tests can inform clinical and public health decision-making





Kissler et al., medRxiv 2020

Acknowledgements and citations

Bubar KM, Kissler SM, Lipsitch M, Cobey S, Grad YH, Larremore DB. <u>Model-informed COVID-19 vaccine</u> prioritization strategies by age and serostatus. *medRxiv*. 2020.

Kissler SM, Tedijanto C, Goldstein E, Grad YH*, Lipsitch M*. <u>Projecting the transmission dynamics of SARS-CoV-2</u> <u>through the post-pandemic period</u>. *Science*. 2020 May 22;368(6493):860-868. *co-senior authors

Kissler SM, Kishore N, Prabhu M, Goffman D, Beilin Y, Landau R, Gyamfi-Bannerman C, Bateman BT, Katz D, Gal J, Bianco A, Stone J, Larremore D, Buckee CO, Grad YH. <u>Reductions in commuting mobility correlate with geographic</u> <u>differences in SARS-CoV-2 prevalence in New York City</u>. *Nature Communications*. 2020. 11:4674 doi.org/10.1038/s41467-020-18271

Kissler SM, Fauver JR, Mack C, Tai C, Shiue KY, Kalinich CC, Jednak S, Ott IM, Vogels CBF, Wohlgemuth J, Weisberger J, DiFiori J, Anderson DJ, Mancell J, Ho DD, Grubaugh ND*, Grad YH*. <u>Viral dynamics of SARS-CoV-2 infection and</u> <u>the predictive value of repeat testing</u>. *medRxiv*. 2020. *co-senior authors.