Network Systems Science & Advanced Computing Biocomplexity Institute & Initiative University of Virginia

Estimation of COVID-19 Impact in Virginia

October 5th, 2022

(data current to Sept 30th – October 4th) Biocomplexity Institute Technical report: TR BI-2022-1770

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biocomplexity.virginia.edu

About Us

- Biocomplexity Institute at the University of Virginia
 - Using big data and simulations to understand massively interactive systems and solve societal problems
- Over 20 years of crafting and analyzing infectious disease models
 - Pandemic response for Influenza, Ebola, Zika, and others



Points of Contact

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Model Development, Outbreak Analytics, and Delivery Team

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Overview

- Goal: Understand impact of COVID-19 mitigations in Virginia
- Approach:
 - Calibrate explanatory mechanistic model to observed cases
 - Project based on scenarios for next 4 months
 - Consider a range of possible mitigation effects in "what-if" scenarios

• Outcomes:

- Ill, Confirmed, Hospitalized, ICU, Ventilated, Death
- Geographic spread over time, case counts, healthcare burdens



Key Takeaways

Projecting future cases precisely is impossible and unnecessary. Even without perfect projections, we can confidently draw conclusions:

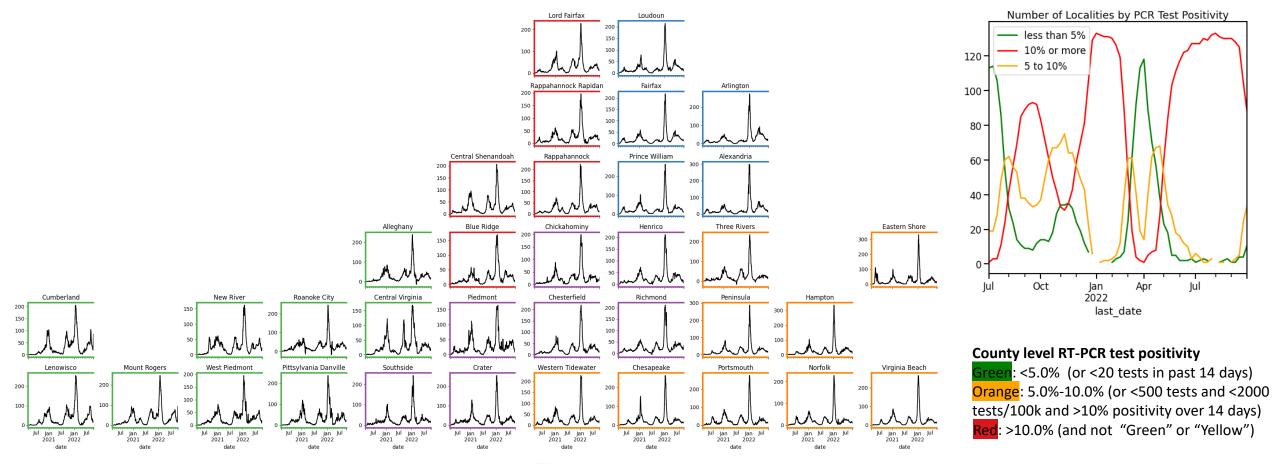
- Case rates continue their decline, hospitalizations continue steady decline
- VA weekly case rate continues decline to 99/100K from 109/100K
 - US weekly case rate is down considerably as well at 89/100K from 105/100K
 - VA hospital occupancy (rolling 7 day mean of 545 down from 599 a week ago) has continued to decline
- Projections anticipate continued declines in cases as well as hospitalizations
- Potential for rebounds due to seasonal forces and/or novel sub-variants in the Fall remains
- Model updates:
 - Maintained Booster Scenarios by slowing down the rate of vaccination adopting the rate of 3rd dose rollout
 - Current monitoring still not finding a definite candidate for Variant X, though BQ.1.1 or BA.2.75.2 remain likely, 50% prevalence pushed back to Nov 15th

The situation continues to change. Models continue to be updated regularly.

Situation Assessment



Case Rates (per 100k) and Test Positivity



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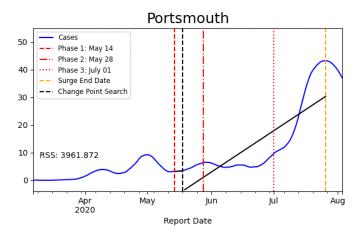
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District Trajectories

Goal: Define epochs of a Health District's COVID-19 incidence to characterize the current trajectory

Method: Find recent peak and use hockey stick fit to find inflection point afterwards, then use this period's slope to define the trajectory

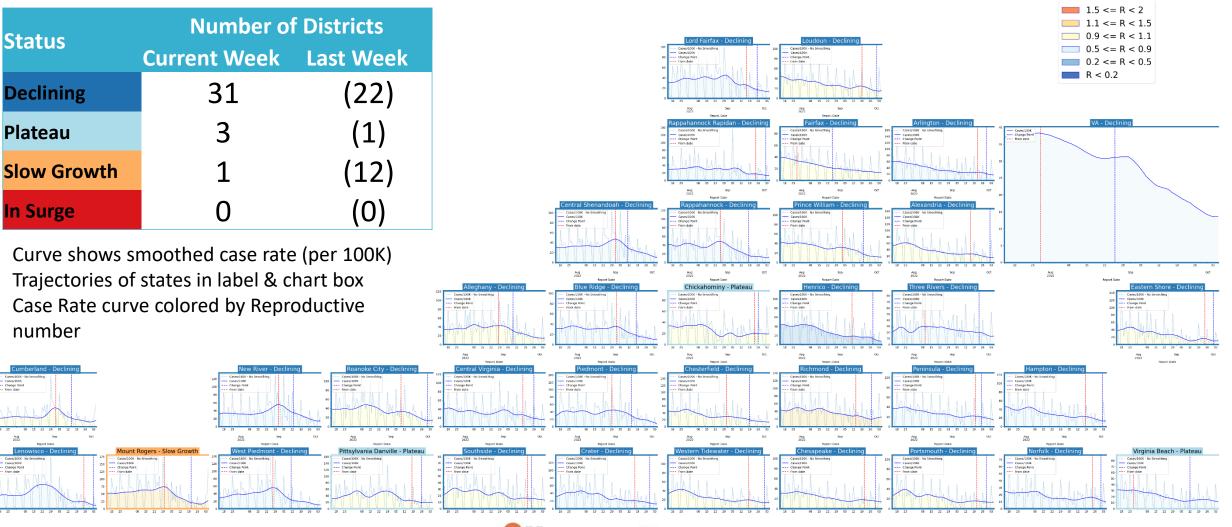
Hockey stick fit



Trajectory	Description	Weekly Case Rate Slope (per 100k)	Weekly Hosp Rate Slope (per 100k)
Declining	Sustained decreases following a recent peak	slope < -0.88/day	slope < -0.07/day
Plateau	Steady level with minimal trend up or down	-0.88/day < slope < 0.42/day	-0.07/day < slope < 0.07/day
Slow Growth	Sustained growth not rapid enough to be considered a Surge	0.42/day < slope < 2.45/day	0.07/day < slope < 0.21/day
In Surge	Currently experiencing sustained rapid and significant growth	2.45/day < slope	0.21/day < slope

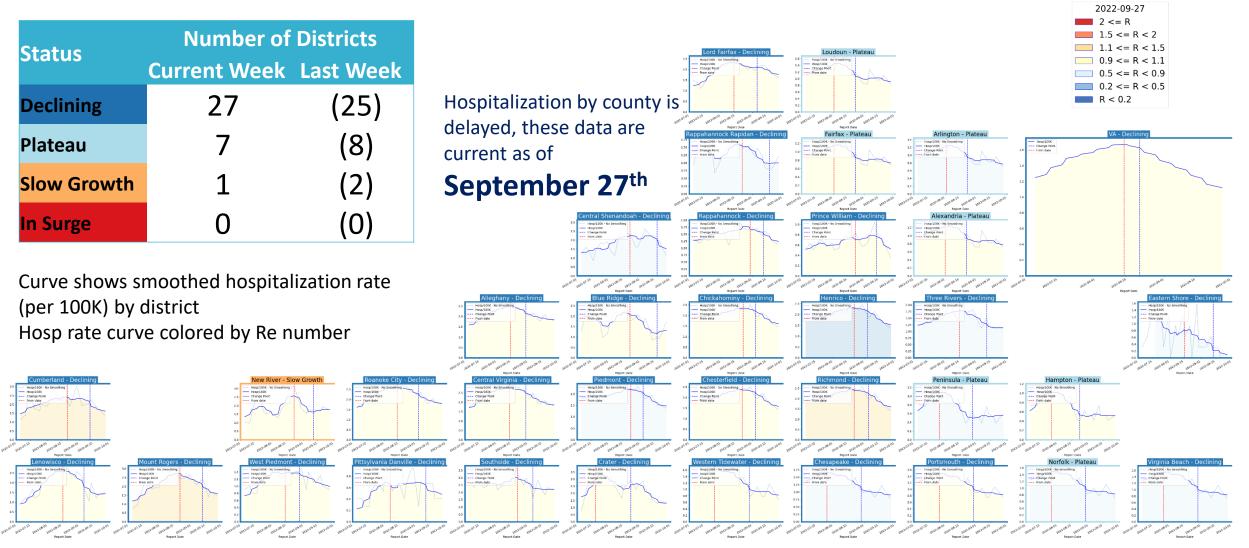


District Case Trajectories – last 10 weeks



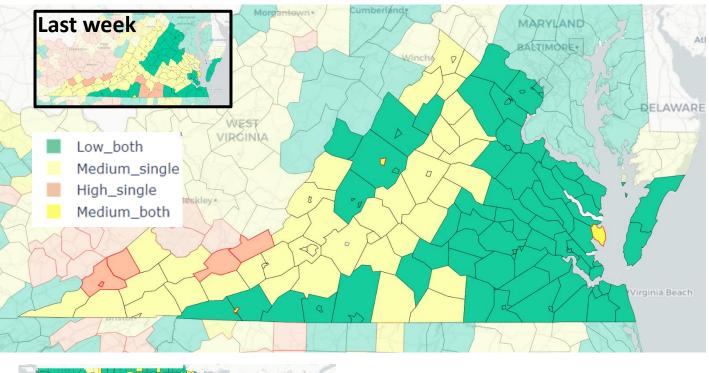
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District Hospital Trajectories – last 10 weeks





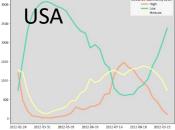
CDC's COVID-19 Community Levels



Last week

7-Oct-22

USA VA 2022-05-05 2022-06-09 2022-07-14 2022-08-18



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Red outline indicates county had 200 or more cases per 100k in last week

Pale color indicates either beds or occupancy set the level for this county

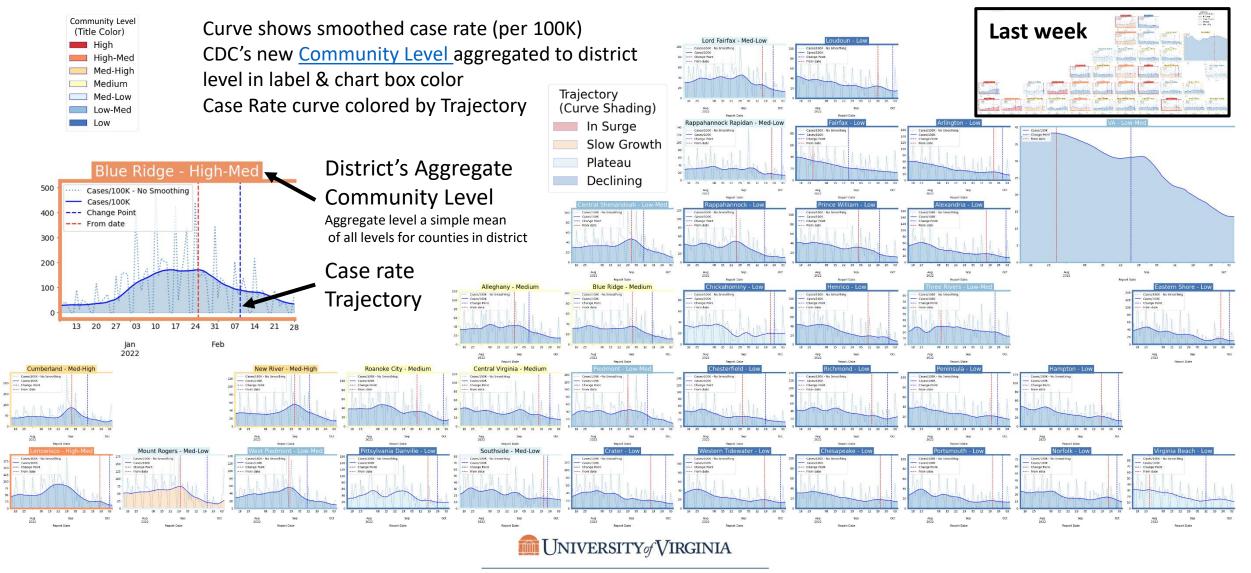
Dark color indicates both beds and occupancy set the level for this county

COVID-19 Community Levels - Use the Highest Level that Applies to Your Community					
New COVID-19 Cases Per 100,000 people in the past 7 days	Indicators	Low	Medium	High	
	New COVID-19 admissions per 100,000 population (7-day total)	<10.0	10.0-19.9	≥20.0	
Fewer than 200	Percent of staffed inpatient beds occupied by COVID-19 patients (7-day average)	<10.0%	10.0-14.9%	≥15.0%	
	New COVID-19 admissions per 100,000 population (7-day total)	NA	<10.0	≥10.0	
200 or more	Percent of staffed inpatient beds occupied by COVID-19 patients (7-day average)	NA	<10.0%	≥10.0%	

The COVID-19 community level is determined by the higher of the new admissions and inpatient beds metrics, based on the current level of new cases per 100,000 population in the past 7 days

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District Trajectories with Community Levels



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Estimating Daily Reproductive Number – Redistributed gap

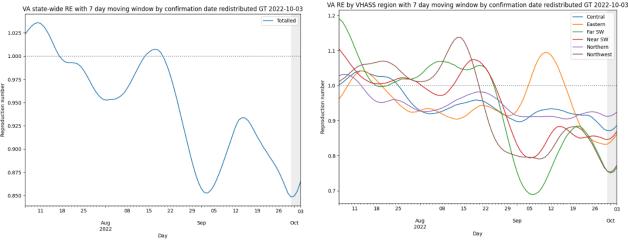
October 3rd Estimates

Region	Date Confirmed R _e	Date Confirmed Diff Last Week
State-wide	0.865	0.006
Central	0.886	0.014
Eastern	0.861	-0.027
Far SW	0.766	-0.121
Near SW	0.868	0.093
Northern	0.924	0.035
Northwest	0.772	-0.025

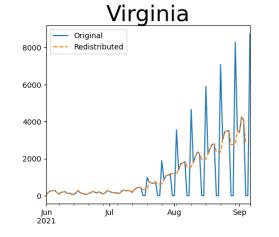
Methodology

- Wallinga-Teunis method (EpiEstim¹) for cases by confirmation date
- Serial interval: updated to discrete distribution from observations (mean=4.3, Flaxman et al, Nature 2020)
- Using Confirmation date since due to increasingly unstable estimates from onset date due to backfill

1. Anne Cori, Neil M. Ferguson, Christophe Fraser, Simon Cauchemez. A New Framework and Software to Estimate Time-Varying Reproduction Numbers During Epidemics. American Journal of Epidemiology, Volume 178, Issue 9, 1 November 2013, Pages 1505–1512, <u>https://doi.org/10.1093/aje/kwt133</u>



Skipping Weekend Reports & holidays biases estimates Redistributed "big" report day to fill in gaps, and then estimate R from "smoothed" time series

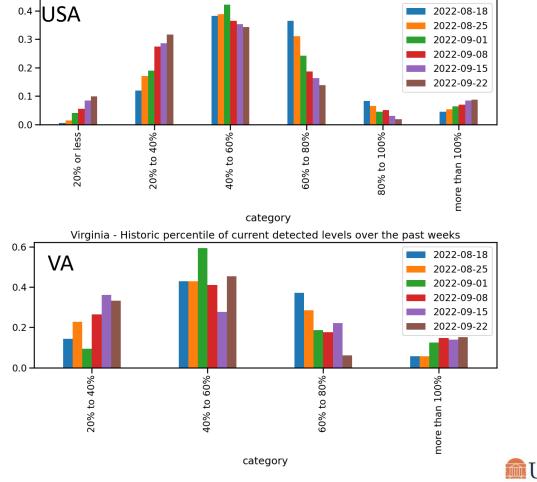


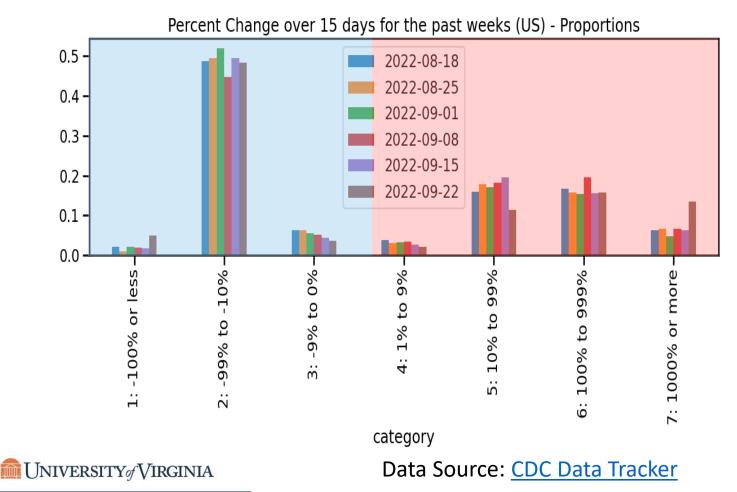
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Wastewater Monitoring

Wastewater provides a coarse early warning of COVID-19 levels in communities

- Overall in the US, there is an increase in sites with increased levels of virus compared to 15 days ago
- Current virus levels are at or exceeding max of previous historical levels, has slowed, though more sites are entering upper quintiles US Historic percentile of current detected levels over the past weeks



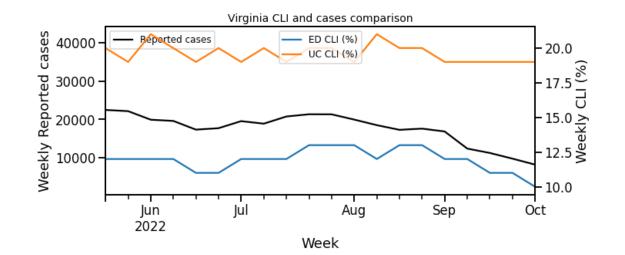


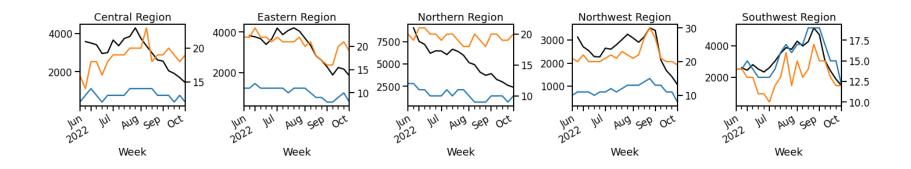
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COVID-like Illness Activity

COVID-like Illness (CLI) gives a measure of COVID transmission in the community

- Emergency Dept (ED) based CLI is more correlated with case reporting
- Urgent Care (UC) is a leading indicator but prone to some false positives
- Current trends in UC CLI have been in a plateau since mid-May state-wide, with some regional fluctuations





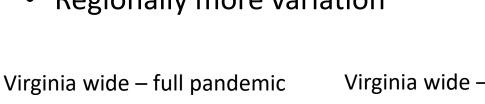


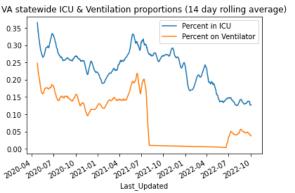
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Hospitalizations and Severe Outcomes Data Source: CDC Data Tracker

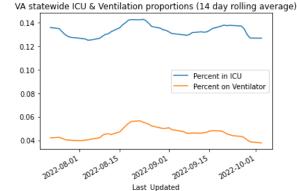
Proportion of most severe outcomes decreasing among those who are hospitalized

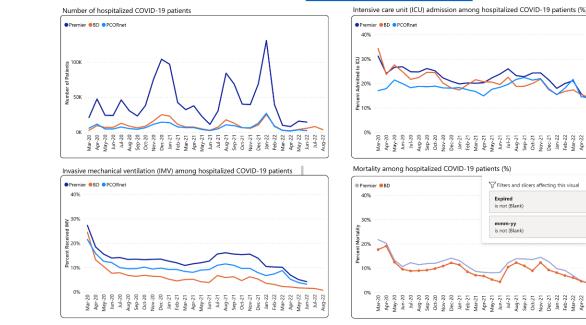
- ICU has declined from ~20% of hospitalized to nearly 10% since initial Omicron wave
- Also seen across all age-groups
- Similar levels of decline seen in VA
- Regionally more variation ullet



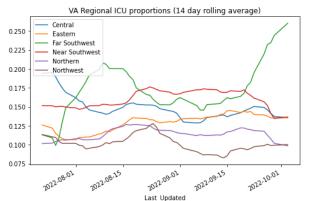


Virginia wide – recent

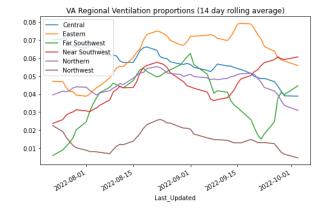




Virginia Regional ICU percent



Virginia Regional Ventilation %



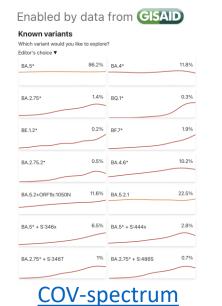
SARS-CoV2 Variants of Concern

Emerging new variants will alter the future trajectories of pandemic and have implications for future control

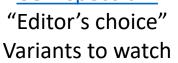
Emerging variants can: Increase transmissibility, increase severity (more hospitalizations and/or deaths), and limit immunity provided by prior infection and vaccinations

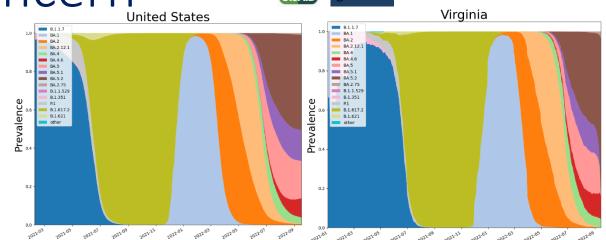
Omicron Updates

- BA.4.6 remains slow growing to about 16% from 15% last week
- BA.5 continues to slowly yield share to other variants, now down to 78% from 80% last week
- BF.7 showing strong growth up to 4.4 from 1.2% last ٠ week
- BQ.1.1 recently seeing growth in England and other countries that mimics past variants of concern that have gone on to dominate
- BA.2.72.2 also shows signs as being a potential candidate as a future variant of concern, has recently been shown to have significant immune escape



COVSPECTRUM





GISAID

outbreak.infc

HHS Region 3: 6/26/2022 - 10/1/2022

HHS Region 3: 9/25/2022 - 10/1/2022 NOWCAST

%Total

77.5%

15.8%

4.4%

0.0%

0.0%

0.0%

74.8-79.99

13.4-18.6%

27-6.9%

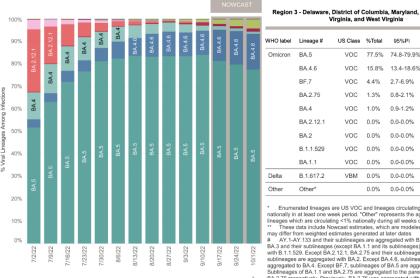
0.8-2.1%

0.0-0.0%

0.0-0.0%

0.0-0.0%

0.0-0.0%



Collection date.

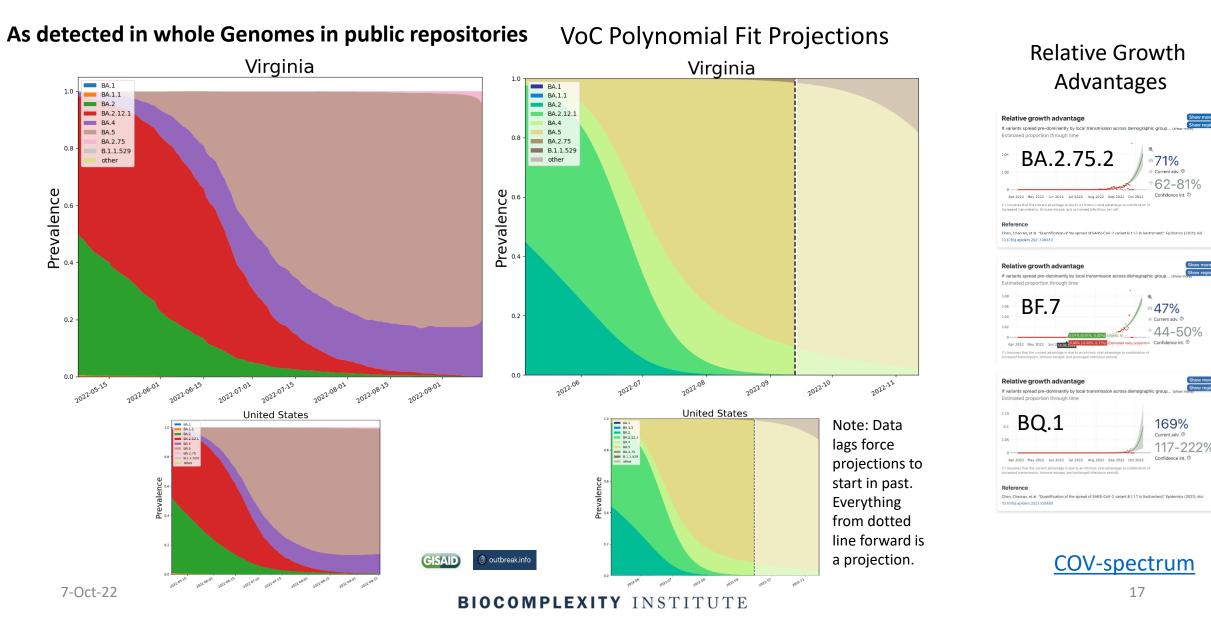
VOC 1.0% 0.9-1.2% VOC 0.0% 0.0-0.0%

0.0% 0.0-0.0% Enumerated lineages are US VOC and lineages circulating above 19 nationally in at least one week period. "Other" represents the aggregation of neages which are circulating <1% nationally during all weeks displayed. nay differ from weighted estimates generated at later dates

AY.1-AY.133 and their sublineages are aggregated with B.1.61 BA.3 and their sublineages (except BA.1.1 and its sublineages) are aggregate with B.1.1.529. Except BA.2.12.1, BA.2.75 and their sublineages, BA.2 sublineages are aggregated with BA.2. Except BA.4.6, sublineages of BA.4 are aggregated to BA.4. Except BF.7, sublineages of BA.5 are aggregated to BA.5 Sublineages of BA.1.1 and BA.2.75 are aggregated to the parental BA.1.1 and BA.2.75 respectively. Previously, BA.2.75 was aggregated with BA.2, and BF.7 vas aggregated with BA.5. Lineages BA.4.6, BF.7, and many BA.2.75 contain the snike substitution R346T



SARS-CoV2 Omicron and Sub-Variants



Pandemic Pubs

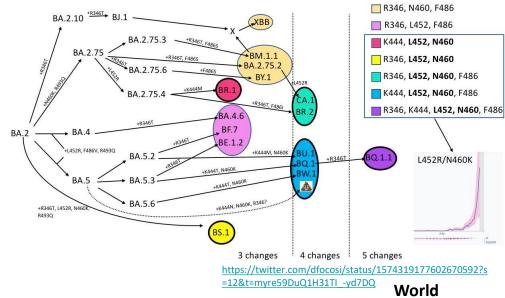
1. BQ.1.1, a descendant of BA.5, shows strong growth advantage in recent sequence surveillance.

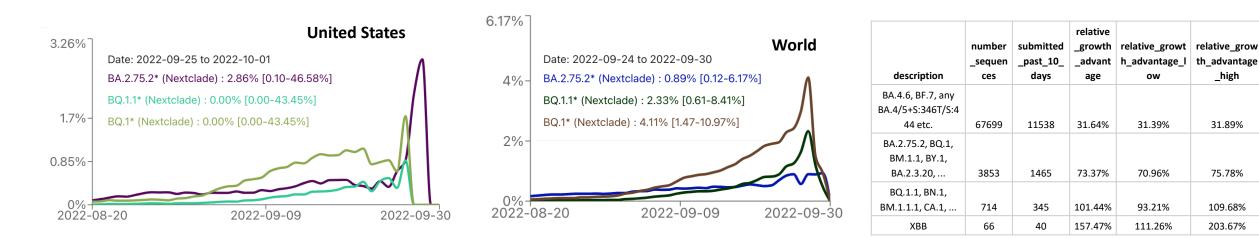
Current Bivalent booster doses target BA.5 making it a well-timed and a worthwhile intervention.



(*) Assumes that the current advantage is due to an intrinsic viral advantage (a combination of increased transmission, immune escape, and prolonged infectious period).

Omicron lineages with changes at 346, 444, 452, 460, and 486



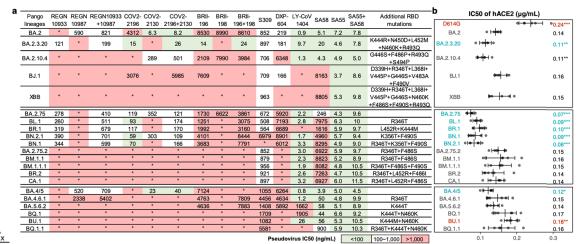


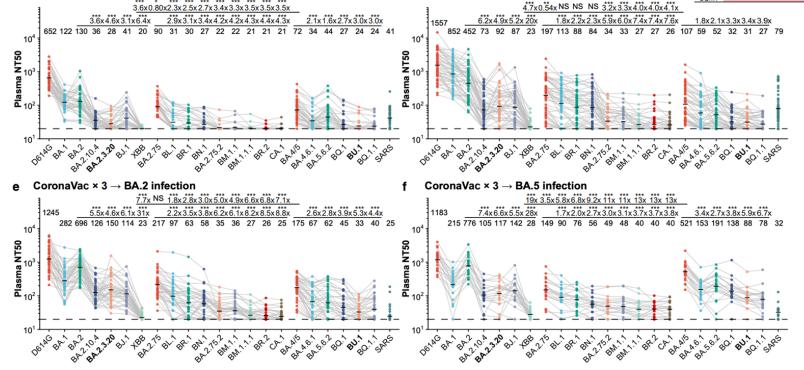
Estimates from surveillance using mutational profiles approximating BQ.1, BQ.1.1, and BA.2.75.2 <u>https://cov-spectrum.org/explore/World/AllSamples/from=2022-08-17&to=2022-10-</u> <u>04/variants?nextcladePangoLineage=BA.2.75.2*&nextcladePangoLineage1=bq.1.1*&nextcladePangoLineage2=bq.1&analysisM</u> ode=CompareEquals& https://cov-spectrum.org/collections/54

Pandemic Pubs

c CoronaVac × 3

2. Both BQ.1.1 and XBB show substantial immune escape and significant negation of many pharmaceutical interventions. XBB is significantly more immune evasive than BA.2.75.2 and BQ.1.1 against plasma from all breakthrough infections, comparable to or even exceeding the level of escape displayed by SARS-CoV-1 against SARS-CoV-2 convalescent plasma.



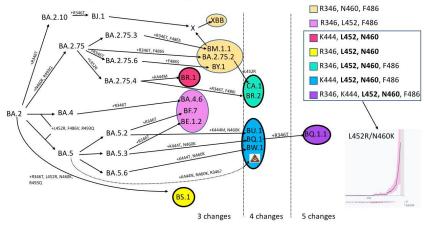


Researchers in Beijing characterize multiple concerning immune escape variants using plasma from breakthrough infections in those vaccinated with three doses of CoronaVac. Similar to BQ.1.1, XBB also escapes Evusheld and Bebtelovimab. BU.1, BR.2,

BM.1.1.1, CA.1, and XBB all displayed moderate hACE2 binding capability relative to previously widely circulating variants of

CoronaVac \times 3 \rightarrow BA.1 infection

Omicron lineages with changes at 346, 444, 452, 460, and 486



https://twitter.com/dfocosi/status/1574319177602670592?s =12&t=myre59DuQ1H31TI_-yd7DQ

https://www.biorxiv.org/content/10.1101/2022.09.15.507787v3 https://twitter.com/yunlong_cao/status/1577343549120872448

BA.2 and BA.5.

Pandemic Pubs (last week

1. Representative survey estimates 7.3% of the US adult population (~18.5 million) reported having long COVID by July

2. A propensity matched cohort from the UK found that 2+ doses of vaccine decreased adjusted long COVID risk by 41% (95% CI 31%-50%)

3. BA.2.75.2 neutralised, on average, at titers ~6.5times lower than BA.5, making BA.2.75.2 the most neutralisation-resistant variant evaluated to date 4. Retrospective cohort study of 6,245,282 older adults (age ≥65 years) who had medical encounters between 2/2020–5/2021, indicates that people with COVID-19 were at significantly increased risk for a new diagnosis of Alzheimer's disease within 360 days after the initial COVID-19 diagnosis

Populatior			HR (95% CI)
Overall (>=6		¦ ⊢•I	1.69 (1.53–1.72)
65-74		⊢ •−−1	1.59 (1.37–1.85)
75-84		⊢ ⊷-1	1.69 (1.54–1.85)
>=85		►•	1.89 (1.73–2.07)
Women			1.82 (1.69–1.97)
Men		⊢ 1	1.50 (1.37–1.65)
Black		⊢ •−−1	1.62 (1.36–1.93)
White		+++	1.61 (1.51–1.72)
Hispanic	· · · ·	⊭ ' '	1.25 (0.97–1.61)
	0 0.5 Haz	1 1.5 2 ard Ratio	

ek)	Total	Long COVID	Crude Prevalence of Long COVID % (95% CI)	Age and sex direct- standardized prevalence of long COVID*	Crude prevalence ratio (PR)	Adjusted prevalence ratio (aPR)**	Estimated Number with Long COVID
	Weighted N (%)	Weighted N (%)		% (95% CI)	PR (95% CI)	aPR (95% CI)	
Fotal	3,042 (100.0)	222 (100.0)	7.3 (6.1, 8.5)				18,533,864
ge							
8-24	365 (12.0)	17 (7.8)	4.8 (0.9, 8.6)	4.4 (1.8 - 10.0)	0.45 (0.27, 0.76)	0.50 (0.30, 0.84)	1,445,641
5-34	547 (18.0)	58 (26.1)	10.6 (6.6, 14.6)	10.0 (6.8 - 14.6)	Ref	Ref	4,837,339
5-44	495 (16.3)	43 (19.1)	8.6 (5.3, 11.9)	9.0 (6.1 - 13.1)	0.81 (0.56, 1.18)	0.87 (0.60, 1.26)	3,539,968
5-54	498 (16.4)	37 (16.5)	7.4 (4.6, 10.2)	7.4 (5.1 - 10.7)	0.70 (0.47, 1.03)	0.72 (0.49, 1.07)	3,058,088
5-64	508 (16.7)	41 (18.2)	8.0 (5.5, 10.5)	8.3 (6.0 - 11.2)	0.75 (0.51, 1.10)	0.80 (0.55, 1.18)	3,373,163
5+	629 (20.7)	27 (12.2)	4.3 (3.0, 5.6)	4.2 (3.0 - 5.8)	0.41 (0.26, 0.63)	0.43 (0.27, 0.66)	2,261,13
Sender							
lale	1,443 (47.4)	72 (32.4)	5.0 (3.6, 6.3)	5.0 (3.8 - 6.5)	Ref	Ref	6,004,97
emele	1 516 (40.9)	144 (64.8)	9.5 (7.5, 11.5)	9.4 (7.7 - 11.6)	1.90 (1.45, 2.50)	1.84 (1.40, 2.42)	12,009,94
T emale		11 68 83	П	205 554 296 4	169 573 86 E	837 659 433 4	74 560 99
<u>3</u>		11 68 83 p<0.0001 p<0.0001	П	205 554 296 4 p<0.00	169 573 86 E	837 659 433 4 p<0.000	74 560 99 11 .0001
3	621 134 1	11 68 83 p<0.0001 p<0.0001 p<0.0001	₃₁ D ₂₂	205 554 296 4 p<0.00	69 573 86 E	837 659 433 4 p<0.000	74 560 99 01 .0001 p=0.0002
100000	621 134 1	11 68 83 p<0.0001 p<0.0001 p<0.0001 p=0.0001	• 31 D 22	205 554 296 4 p<0.00	169 573 86 E	837 659 433 4 p<0.000	74 560 99 11 .0001
100000	621 134 1	11 68 83 p<0.0001 p<0.0001 p<0.0001 p=0.0001	₃₁ D ₂₂	205 554 296 4 p<0.00	69 573 86 E	837 659 433 4 p<0.000	74 560 99 01 .0001 p=0.0002
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100000		11 68 83 p=0.0001 p=0.0001 p=0.0001 p=0.0001 p=0.0001 p=0.0001		205 554 296 4 	69 573 86 E ³¹ ⊳<0.0001 p<0.0001	837 659 433 4 	74 560 99 11 .0001 p=0.0002 p<0.0001
Neutralisation titre (ID ₅₀)		11 68 83 p=0.0001 p=0.0001 p=0.0001 p=0.0001 p=0.0001 p=0.0001	• 31 D 22	205 554 296 4 	69 573 86 E	837 659 433 4 	74 560 99 11 .0001 p=0.0002 p<0.0001
Neutralisation titre (ID ₅₀) 	621 134 1	11 68 83 p=0.0001 p=0.0001 p=0.0001 p=0.0001 p=0.0001 p=0.0001	a	205 554 296 4 p=000 p=0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	69 573 86 E 31 p=0.0000	837 659 433 4 p=0.000 p=0 0 0 0 0 0 0 0 0 0 0 0 0 0	74 560 99 11 0001 p=0.0002 p=0.0001 p=0.0001 p=0.0001 p=0.0001

In 18 random blood donor samples in Stockholm, sampled recently BA.2.75.2 was neutralised, on average, five-fold less potently than BA.5. These are recent samples in a city that has good vaccine coverage and likely relatively high prior infection rates. Researchers also report the sensitivity of emerging omicron sublineages BA.2.75.2, BA.4.6, and BA.2.10.4 to neutralisation by a panel of clinically relevant and pre-clinical monoclonal antibodies, as well as by serum from blood donated in Stockholm, Sweden.

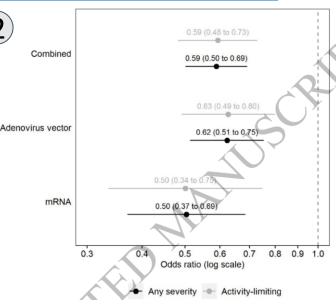
https://www.biorxiv.org/content/10.1101/2022.09.16.508299v2

https://twitter.com/benjmurrell/status/1570862185819303937?s=12&t=sL45iDMQ7GGmC2KakXPNZg

NY researchers conducted a population-representative survey, June 30-July 2, 2022, of a random sample of 3,042 United States adults. Using questions developed by the United Kingdom's Office of National Statistics, we estimated the prevalence by sociodemographics, adjusting for gender and age. An estimated 7.3% (95% Cl: 6.1-8.5%) of all respondents reported long COVID, approximately 18,533,864 adults. One-quarter (25.3% [18.2-32.4%]) of respondents with long COVID reported their day-to-day activities were impacted 'a lot' and 28.9% had SARS-CoV-2 infection >12 months ago.

https://www.medrxiv.org/content/10.1101/2022.09.12.22279862v1

2

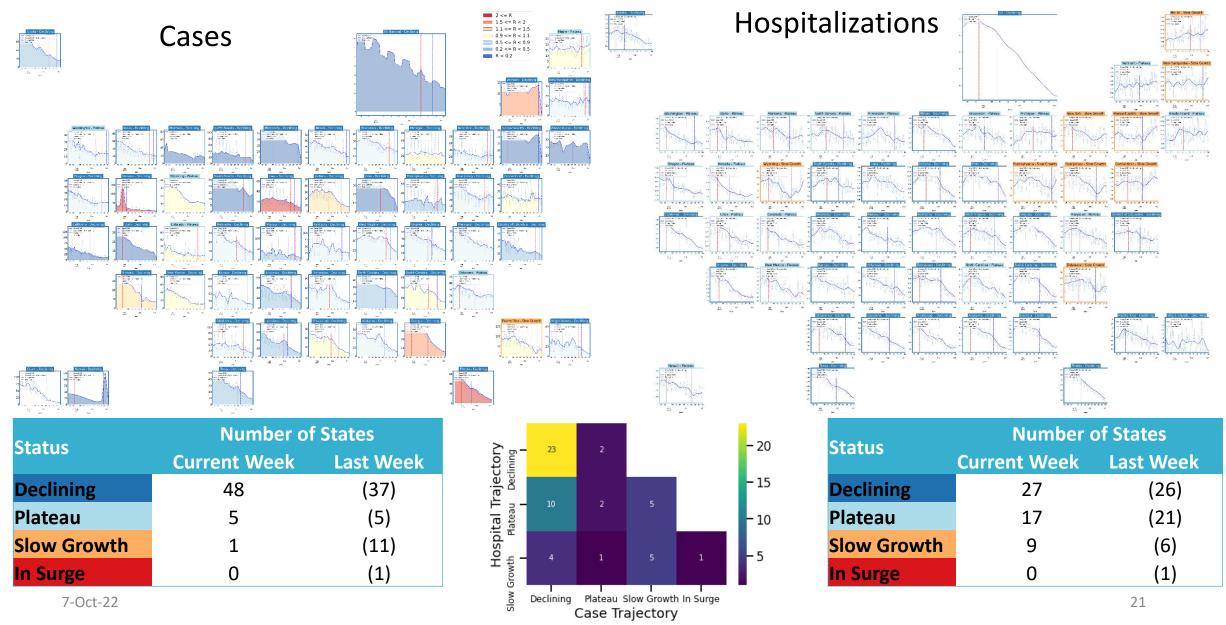


Oxford researchers investigated Long Covid incidence by vaccination status in a random sample of UK adults from April 2020 to November 2021. Persistent symptoms were reported by 9.5% of 3,090 breakthrough SARS-CoV-2 infections and 14.6% of unvaccinated controls (adjusted odds ratio 0.59, 95% CI: 0.50-0.69), emphasising the need for public health initiatives to increase population-level vaccine uptake. Matched study participants who were double-vaccinated at time of infection to control participants who were unvaccinated when infected and remained so at their first follow-up visit ≥12 weeks later. Most double-vaccinated participants (3,057, 98.9%) were infected after 17 May 2021, when the Delta variant dominated in the UK, while nearly all unvaccinated participants (3,082, 14 99.7%) were infected before this date.

https://academic.oup.com/ofid/advance-article/doi/10.1093/ofid/ofac464/6696170?login=false https://twitter.com/DFisman/status/1570901408211402752

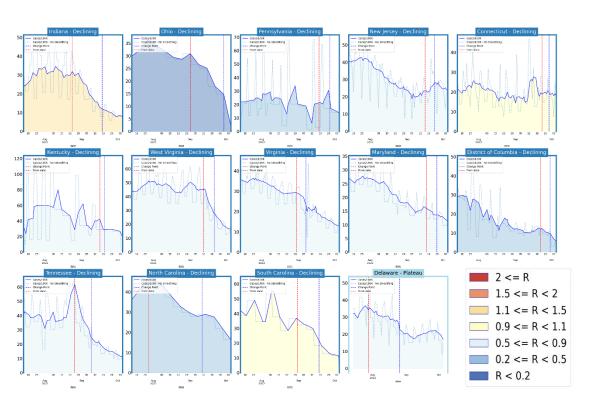
https://content.iospress.com/articles/journal-of-alzheimers-disease/jad220717

United States Case & Hospitalizations

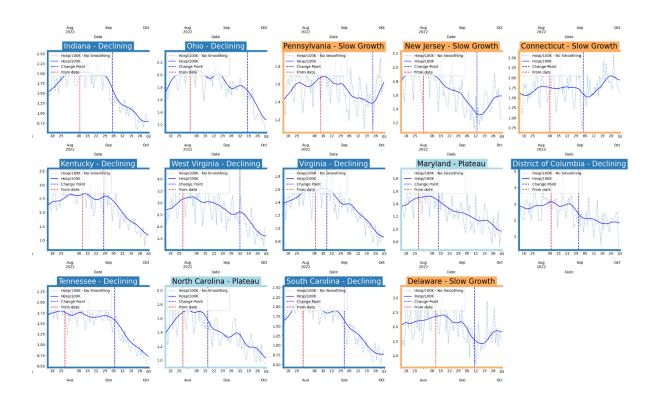


Virginia and Her Neighbors

Cases



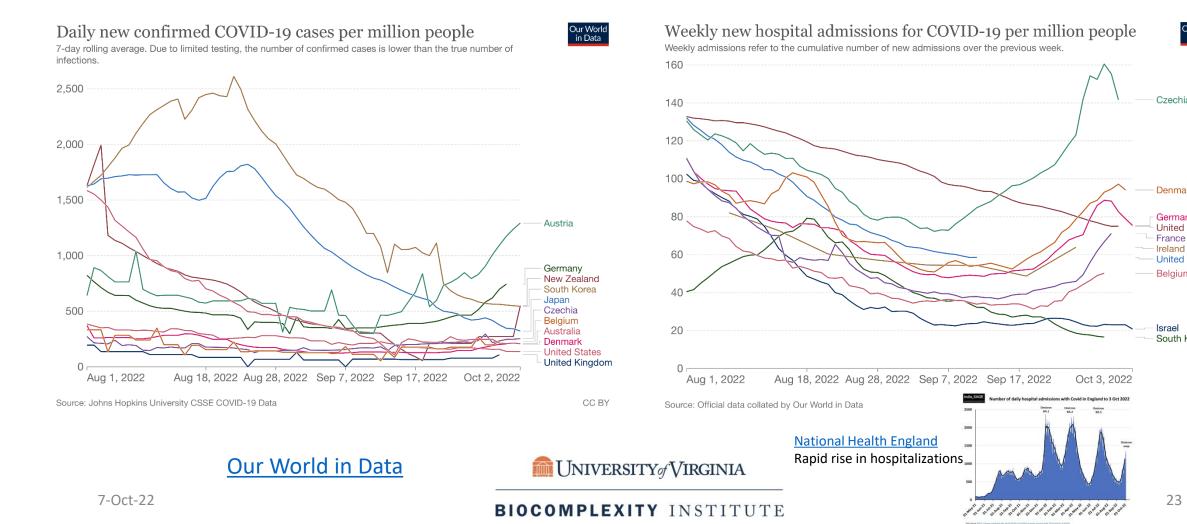
Hospitalizations





Around the World – Various trajectories

Confirmed cases



Hospitalizations

23

Our World in Data

Czechia

Denmark

Germany

Ireland

Belgium

Israel

South Korea

CC BY

United States

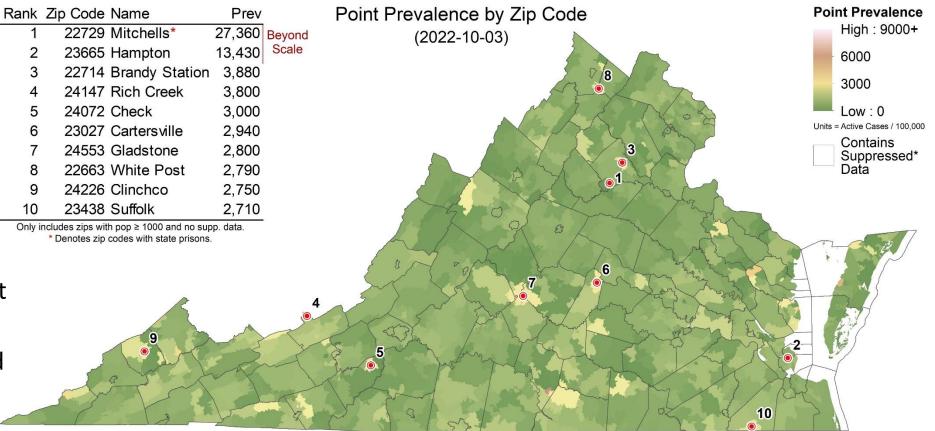
United Kingdom

Zip code level weekly Case Rate (per 100K)

Case Rates in the last week by zip code

- Statewide rates have fallen significantly. We will adjust the color ramp if trends continue.
- High prevalence areas are spread randomly across Virginia (no spatial autocorrelation). -
- Mitchells, VA has by far the highest prevalence. It is home to a prison.
- Some counts are low and suppressed to protect anonymity. Those are shown with a dark red outline.

7-Oct-22



Based on Spatial Empirical Bayes smoothed point prevalence, with an 8:1 ascertainment ratio, for week ending 2022-10-03.

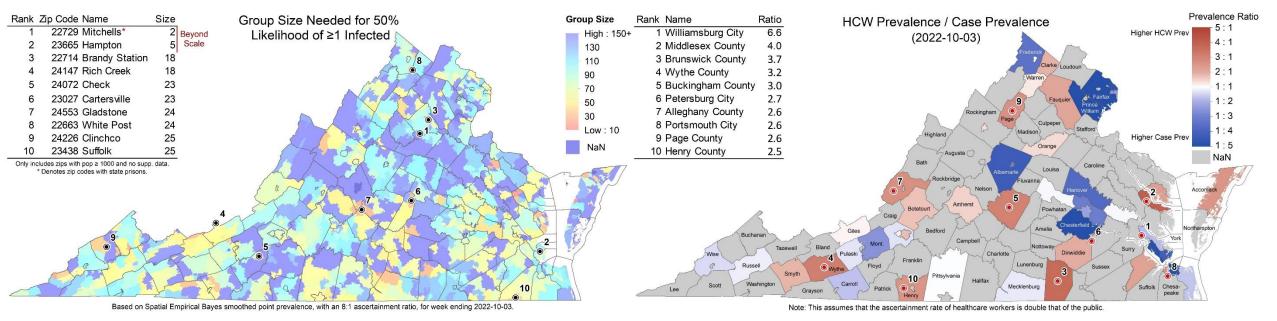
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Risk of Exposure by Group Size and HCW prevalence

Case Prevalence in the last week by zip code used to calculate risk of encountering someone infected in a gathering of randomly selected people

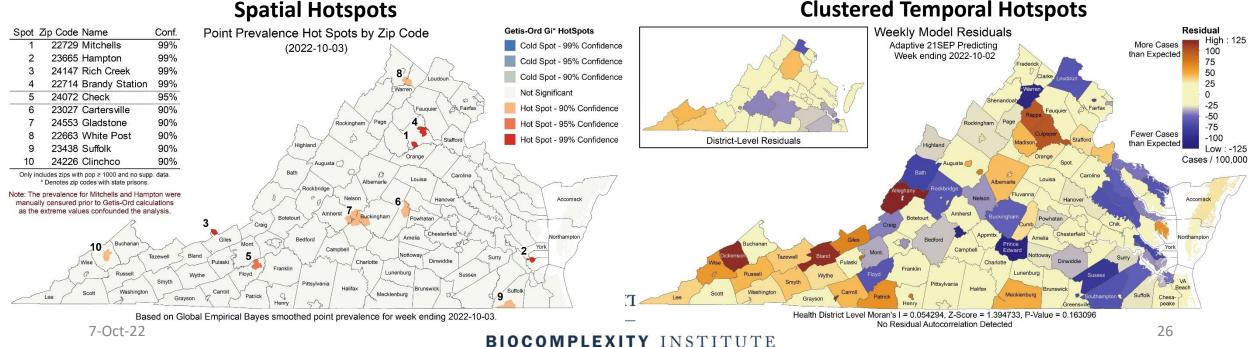
- **Group Size**: Assumes **8 undetected infections** per confirmed case (ascertainment rate from recent seroprevalence survey) and shows minimum size of a group with a 50% chance an individual is infected by zip code (e.g., in a group 2 in Mitchells, there is a 50% chance someone will be infected).
- HCW ratio: Case rate among health care workers (HCW) in the last week using patient facing health care workers as the denominator / general population's case prevalence. Note Williamsburg City.



Current Hot-Spots

Case rates that are significantly different from neighboring areas or model projections

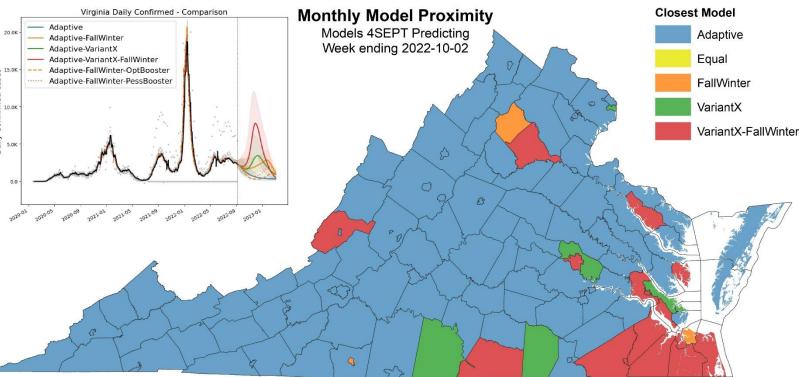
- Spatial: Getis-Ord Gi* based hot spots compare clusters of zip codes with weekly case prevalence higher than nearby zip codes to identify larger areas with statistically significant deviations
- Temporal: The weekly case rate (per 100K) projected last week compared to observed by county, which highlights temporal fluctuations that differ from the model's projections. Note new color ramp (down 75%).
- Spatial hot spots are appearing sporadically across the Commonwealth. Models slightly underpredicted the Far Southwest, and slightly overpredicted Central Virginia, Northern Virginia, and the Tidewater.



Scenario Trajectory Tracking

Which scenario from a month ago did the projection for each county track closest?

- One month out separates the projections more and reveals larger overall patterns.
- Booster scenarios not included as vaccination efforts have only recently begun.
- Overall state level models were on track, largely tracking the Adaptive scenario over Variant X This is expected, as no novel variants supplanted BA.4/5 in the time since the modeling run.
- Currently only a handful of counties, mostly in the Southeast, are tracking FallWinter scenarios more closely than the Adaptive scenario.





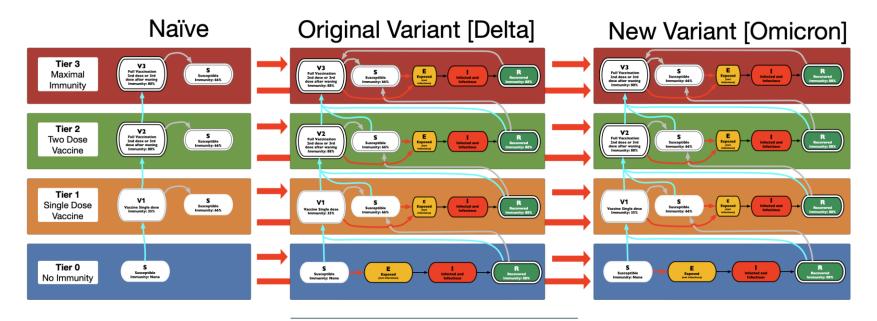
Model Update – Adaptive Fitting



Model Structure Extended for more sub-variants

Omicron sub-variants escape immunity induced by previous sub-variants

- Multiple strain support allows representation of differential protection based on immunological history (BA.1, BA.2, BA.2.12.1, BA.4/5, and future variants (VariantX))
- Each sub-variant has differing levels of immune escape to previous sub-variants, the prevalences are based on observations for fitting purposes, and projections use estimated future prevalences
- Adaptive fitting approach continues to use simulation to generate the full distribution of immune states across the population



Adaptive Fitting Approach

Each county fit precisely, with recent trends used for future projection

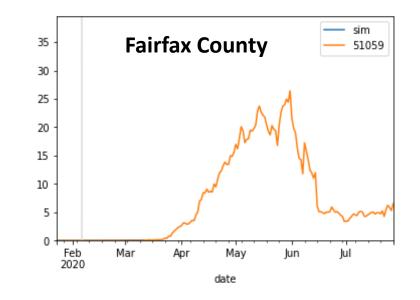
 Allows history to be precisely captured, and used to guide bounds on projections

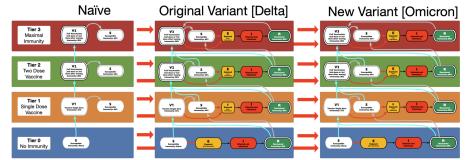
Model: An alternative use of the same meta-population model, PatchSim with multiple tiers of immunity

- Allows for future "what-if" Scenarios to be layered on top of calibrated model
- Allows for waning of immunity and for partial immunity against different outcomes (eg lower protection for infection than death)

External Seeding: Steady low-level importation

 Widespread pandemic eliminates sensitivity to initial conditions, we use steady 1 case per 10M population per day external seeding







Using Ensemble Model to Guide Projections

Ensemble methodology that combines the Adaptive with machine learning and statistical models such as:

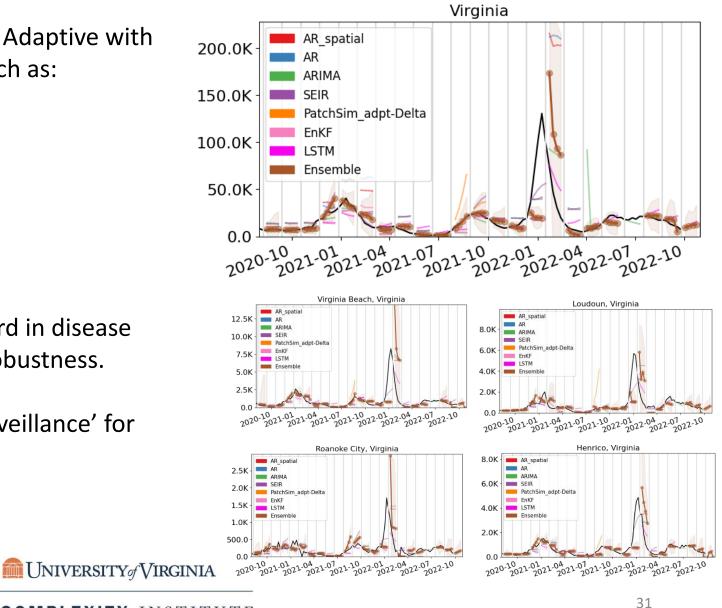
- Autoregressive (AR, ARIMA)
- Neural networks (LSTM)
- Kalman filtering (EnKF)

Weekly forecasts done at county level.

Models chosen because of their track record in disease forecasting and to increase diversity and robustness.

Ensemble forecast provides additional 'surveillance' for making scenario-based projections.

Also submitted to CDC Forecast Hub.



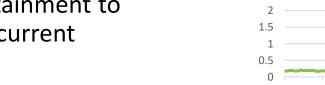
Seroprevalence updates to model design

Several seroprevalence studies have stopped

 CDC Nationwide Commercial Laboratory Seroprevalence Survey is no longer reporting updates; pre-Omicron this data estimated ascertainment ratio of ~4-6x

Testing Behavior has changed, fewer cases are reported

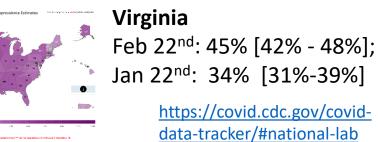
- Home testing, reduced symptoms due to breakthrough / reinfection, and elimination of public health leave
- Outbreaks Near Me from Boston Children's Hospital and Momentive collects reports of home testing
- Wastewater data is consistent with case ascertainment being significantly lower than during the Omicron BA.1 wave
- Accounting for home testing, changes case ascertainment to be 2 times more than pre-Omicron resulting in a current rate of 16 infections to one case

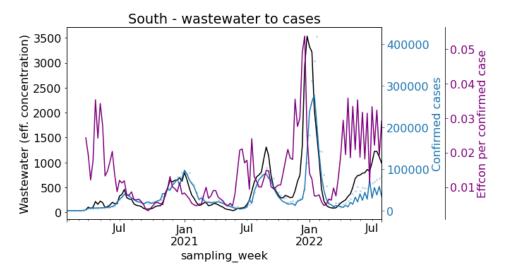


57.7%

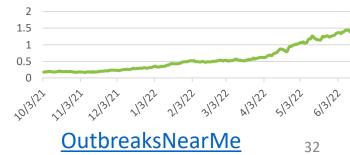
Number of Extension Infortions 1986,893,000 PAR Or Falser Income INVERTIGATION

tol N. moor of Cool Broad the Find of 2/28/2822 78:357:352





Smoothed_Ratio_Home_Test_to_Not_Home_Test



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Calibration Approach

- Data:
 - County level case counts by date of onset (from VDH)
 - Confirmed cases for model fitting
- Calibration: fit model to observed data and ensemble's forecast
 - Tune transmissibility across ranges of:
 - Duration of incubation (5-9 days), infectiousness (3-7 days)
 - Undocumented case rate (1x to 7x) guided by seroprevalence studies
 - Detection delay: exposure to confirmation (4-12 days)
 - Approach captures uncertainty, but allows model to precisely track the full trajectory of the outbreak
- **Project:** future cases and outcomes generated using the collection of fit models run into the future
 - Mean trend from last 7 days of observed cases and first week of ensemble's forecast used
 - Outliers removed based on variances in the previous 3 weeks
 - 2 week interpolation to smooth transitions in rapidly changing trajectories
- **Outcomes**: Data driven by shift and ratio that has least error in last month of observations
 - Hospitalizations: 3 days from confirmation, 6.8% of cases hospitalized
 - Deaths: 11 days from confirmation, 1.45% of cases die



COVID-19 in Virginia:





Dashboard Updated: 10/5/2022 Data entered by 5:00 PM the prior day.

Cases, Hospitalizations and Deaths					
Total 0 2,093		Total H Admiss		Tot Dea	
(New Case	,	56,	344	21,9	919
Confirmed† 1,482,278	Probable† 611,115	Confirmed† 52,871	Probable† 3,473	Confirmed† 18,208	Probable† 3,711

* Includes both people with a positive test (Confirmed), and symptomatic with a known exposure to COVID-19 (Probable).
** Hospitalization of a case is captured at the time VDH performs case investigation. This underrepresents the total number of hospitalizations in Virginia.

^New cases represent the number of confirmed and probable cases reported to VDH in the past 24 hours.

† VDH adopted the updated CDC COVID-19 confirmed and probable surveillance case definitions on August 27, 2020. Found

here: https://wwwn.cdc.gov/nndss/conditions/coronavirus-disease-2019-covid-19/case-definition/2020/08/05/

٥	utbreaks
Total Outbreaks*	Outbreak Associated Cases
9,912	161,992

* At least two (2) lab confirmed cases are required to classify an outbreak.

Testing (PCR Only)			
Testing Encounters PCR Only*	Current 7-Day Positivity Rate PCR Only**		
15,035,997	10.5%		

* PCR" refers to "Reverse transcriptase polymerase chain reaction laboratory testing."
** Lab reports may not have been received yet. Percent positivity is not calculated for days with incomplete data

-	m Inflammatory ne in Children
Total Cases*	Total Deaths
181	1

*Cases defined by CDC HAN case definition: https://emergency.cdc.gov/han/2020/han00432.asp

Accessed 9:00am October 5, 2022 https://www.vdh.virginia.gov/coronavirus/



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Scenarios – Transmission Conditions

- Variety of factors continue to drive transmission rates
 - Seasonal impact of weather patterns, travel and gatherings, fatigue and premature relaxation of infection control practices
- Waning Immunity: Omicron waning with a mean of 4 months

• Projection Condition Ingredients:

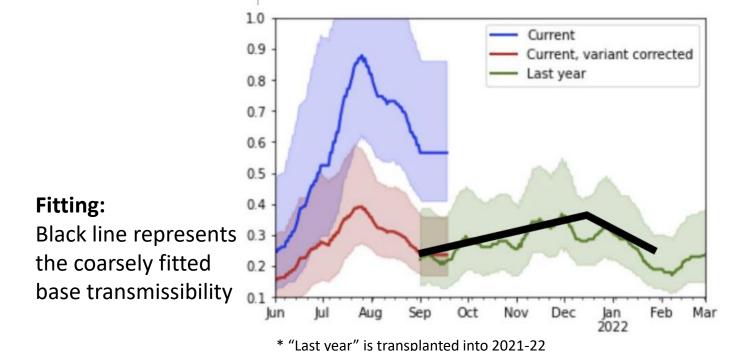
- Adaptive: Controls remain as currently experienced into the future with NO influence from other conditions (eg seasonal, variants, etc.)
- Seasonal (Fall-Winter boosting): Controls remain the same, however, seasonal forcing similar to past Fall-Winter waves is added from Sept-Feb
- Vaccine Booster Campaign (Booster): Reformulated booster available this fall provides improved immunity against Omicron sub-variants
- New Variants (VariantX): As of yet unidentified novel sub-variant with similar immune escape but no transmission advantage emerges 4 months after the last significant sub-variant and grows at a similar rate



Scenarios – FallWinter

September – February saw strong waves of transmission for both years

- Based on analyses of the past 2 seasons we generate a "coarse baseline transmission boost"
 - In 2021 the distribution of fitted model transmissibility was nearly identical between these periods when corrected for Delta's increased transmissibility
- FallWinter captures these "transmission drivers" from the past and uses them as if they were to occur again this season

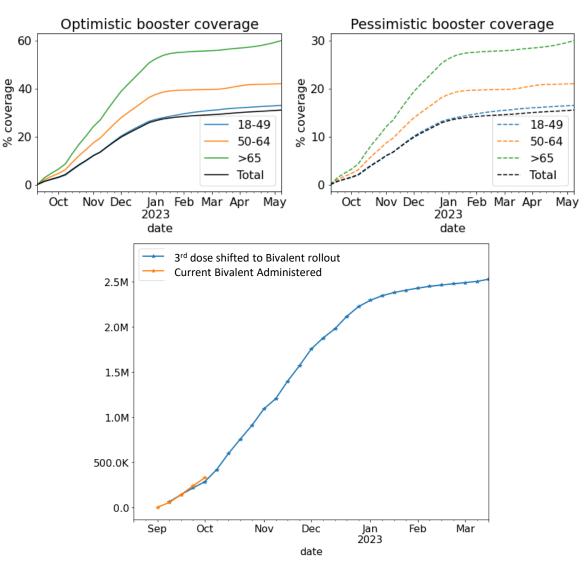


Scenarios – Optimistic vs. Pessimistic Booster Coverage

Reformulated Boosters available now

- Assuming Vax efficacy for BA.4/5 and previous variants is 80% against symptomatic illness
- Campaign starts in mid-September following the ongoing rollout
- Pace of vaccination follows that of 3rd booster (adjusted after initial uptake did not keep pace with seasonal influenza vax uptake)
- Variant X has same immune escape to these vaccines as against BA.5 (40%)

Optimistic coverage: 33% of pop (same as 3rd dose) **Pessimistic coverage**: 16% of pop (half of 3rd dose)

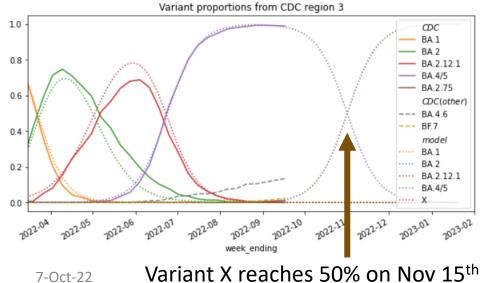


Scenarios – Variant X

Omicron sub-variants seem to be emerging and then dominating with some regularity

- ECDC currently monitoring BA.2 + L452X and BA.2.75 as VOI and XAK as well as other BA.4 and BA.5 strains as Variants under Monitoring
- BA.2.75.2 remains main sub-variant characterized enough to suggest future dominance, but has yet to demonstrate rapid growth in US
- Hypothetical future sub-variant, **VariantX**, may continue the pattern. Assumes similar growth and level of immune escape against previous sub-variants as BA.4/5 (same transmissibility and 40% immune escape against BA.4/5, higher for other sub-variants).



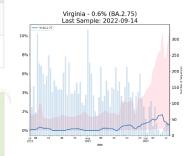


Variants of Interest

WHO label	Lineage + additional mutations	Country first detected (community)	Spike mutations of interest	Year and month first detected	Impact on transmissibility	Impact on immunity	Impact on severity	Transmission in EU/EEA		
Omicron	BA.2 + L452X (x)	n/a	L452X	n/a	No evidence	Increased (7)	No evidence	Detected (a)		
Omicron	BA.2.75	India	(y)	May 2022	No evidence	Increased (9, 10)	No evidence	Detected (a)		
Variants under Monitoring										
WHO	Lineage + additional mutations	Country first detected	Spike mutations	Year and month first	Impact on	Impact on	Impact on	Transmission		
10.001	mutations	(community)	of interest	detected	transmissibility	immunity	severity	in EU/EEA		
Omicron	XAK	(community) Germany	(y)	June 2022	No evidence	immunity No evidence	No evidence	in EU/EEA Detected (a)		
						No	No			
Omicron	<u>ХАК</u> <u>ВА.4</u> +	Germany	(y)	June 2022	No evidence	No evidence No	No evidence No	Detected (a)		

BA.2.75 detected in VA

(very limited samples)



Projection Scenarios – Combined Conditions

Name	Txm	Variant	Description			
Adaptive	С	SQ	Likely trajectory based on conditions remaining similar to the current experience, includes immune escape due to Omicron			
Adaptive-FallWinter	FallWinter	SQ	Like Adaptive, with seasonal forcing of FallWinter added on			
Adaptive-FallWinter-OptBooster	FallWinter	SQ	Like Adaptive-Fall Winter but with Optimistic Booster (90% of seasonal influenza coverage)			
Adaptive-FallWinter-PessBooster	FallWinter	SQ	Like Adaptive-Fall Winter but with Pessimistic Booster (45% of seasonal influenza coverage)			
Adaptive-VariantX	С	Х	Like Adaptive, with emergence of a speculative unknown variant 4 months after BA.4/5 with similar level of immune escape and equal transmissibility			
Adaptive-VariantX-FallWinter	FallWinter	Х	Like Adaptive-VariantX but with the seasonal force of FallWinter added on			
FallWinter = Tra added as a sea	C = Current levels persist into the future FallWinter = Transmission rates learned from Sept through February of past seasons are estimated and added as a seasonal boosting to baseline transmission rates SQ = Status quo of current transmission driver from BA.5 remains the same (eg already significantly past					

SQ = Status quo of current transmission driver from BA.5 remains the same (eg already significantly past dominance, thus no significant major driving of transmission anticipated)

X = Speculative novel sub-variant scenario, emerges 4 months after current variant, similar levels of escape

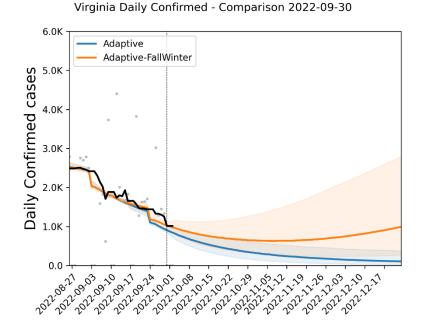
Model Results



Previous projections comparison - Cases

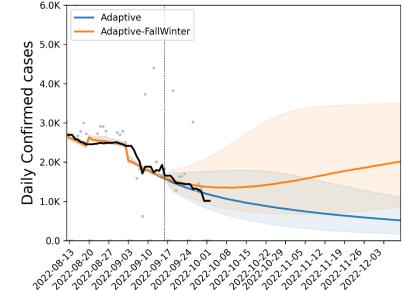
- Previous projections continue to track observed cases
- Projection from 2 weeks ago projected continued decline but cases plateaued
- Projection from 4 weeks ago projected slower decline better capturing recent plateau

This week's projection



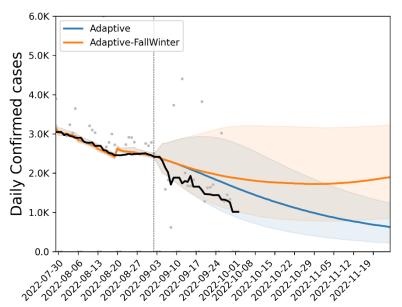
Projection from 2 weeks ago

Virginia Daily Confirmed - Comparison 2022-09-16



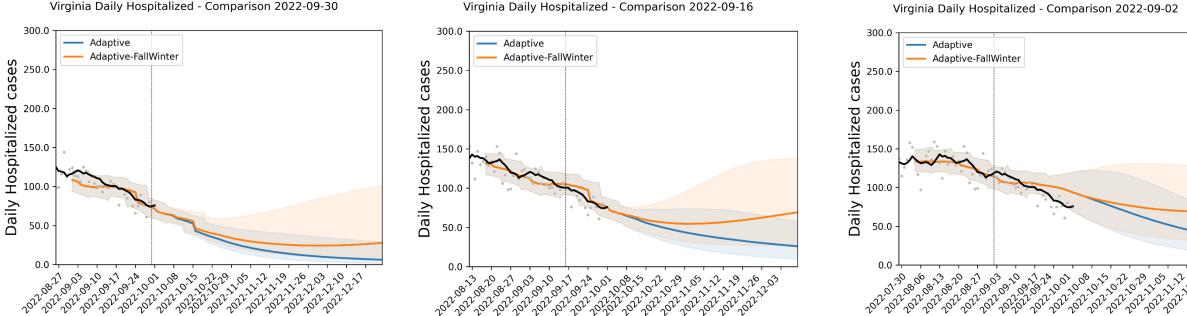
Projection from 4 weeks ago

Virginia Daily Confirmed - Comparison 2022-09-02



Previous projections comparison - Hospitalizations

- Previous projections have tracked observed hospitalizations well
- Projection from 2 weeks ago projected continued decline
- Projection from late July anticipated a plateau giving way to gentle decline



Projection from 2 weeks ago

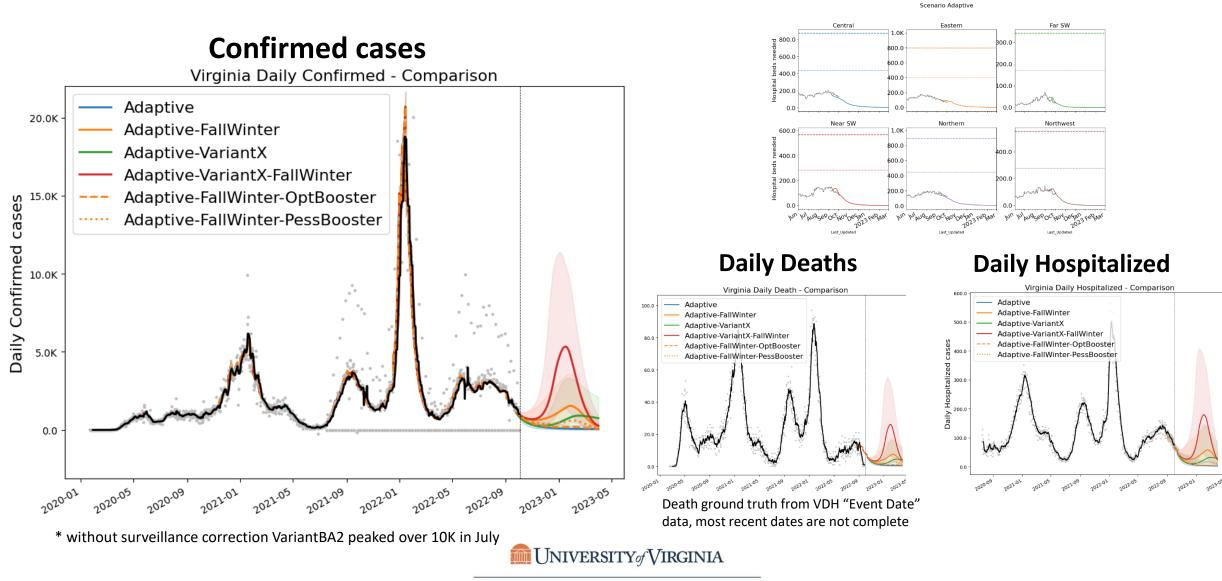
Virginia Daily Hospitalized - Comparison 2022-09-16

This week's projection

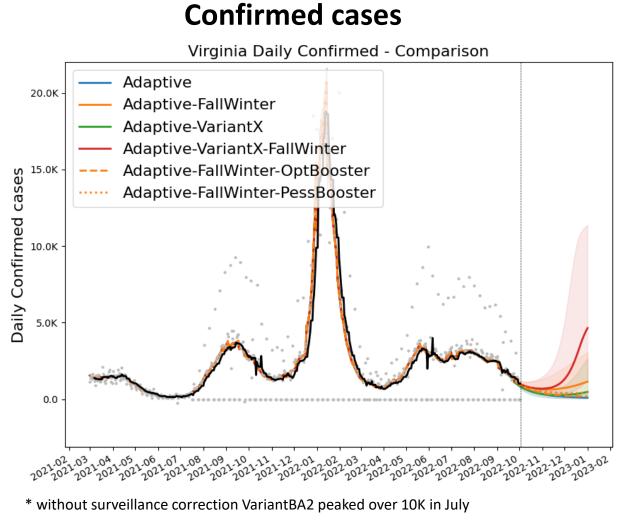
Projection from 4 weeks ago

Outcome Projections

Estimated Hospital Occupancy

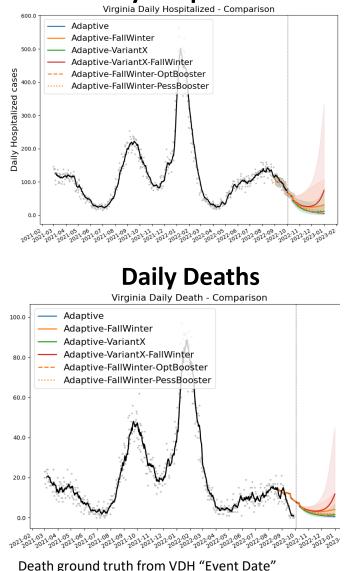


Outcome Projections – Closer Look



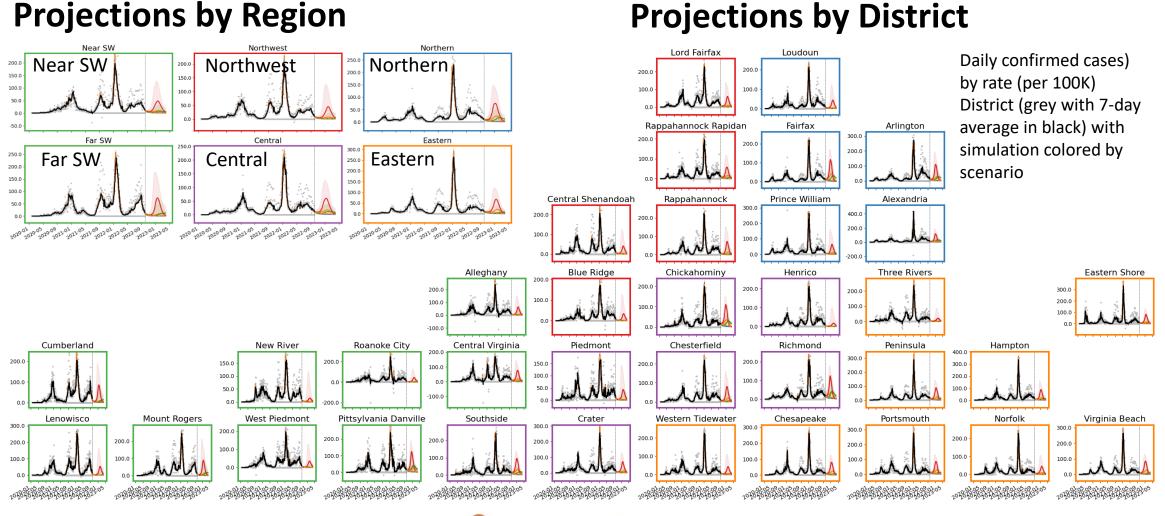






data, most recent dates are not complete

Detailed Projections: All Scenarios



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Detailed Projections: All Scenarios - Closer Look

Projections by Region Projections by District Northern Lord Fairfax Daily confirmed cases by Northwest Northern NeanSW 40.0 40.0 30.0 30.0 rate (per 100K) District 20.0 20.0 20.0 10.0 20.0 (grey with 7-day average 10.0 10.0 in black) with simulation Rappahannock Rapidan Fairfax Arlington 40.0 Far SW Central Eastern 30.0 colored by scenario 100.0 40.0 Eastern 20.0 20 (Central 80.0 Far SW 10.0 60.0 40.0 20.0 Central Shenandoah Rappahannock Prince William Alexandria 10.0 20.0 40 C 40.0 20²²⁻⁰¹ 20²²⁻⁰⁸ 20²²⁻⁰⁹ 20²²⁻¹⁰ 20²⁻¹¹ 02209 02220 00222 00222 002302 002302 -22.12 223.01 223.02 20.0 20.0 20.0 Blue Ridge Chickahominy Henrico Three Rivers Eastern Shore Alleghan 40.0 · 30.0 30.0 40 C 20.0 20.0 20.0 20.0 20 10.0 20.0 10.0 Central Virginia Cumberland Piedmont Chesterfield Richmond Peninsula New River Roanoke City Hampton 100.0 40.0 40.0 40.0 m 40.0 40.0 40.0 40.0 50.0 20.0 20.0 20.0 20.0 20.0 20.0 Lenowisco Mount Rogers West Piedmont Pittsylvania Danville Southside Crater Western Tidewater Chesapeake Portsmouth Norfolk Virginia Beach 30.0 100.0 60.0 75.0 60.0 40.0 30.0 30.0 40.0 20.0 40.0 40.0 50.0 20.0 20.0 50.0 10.0 10.0 10.0 25.0 20.0 0.0 2022022020202020202020202020202 201201202202202202202202202 02202202202202202202202302 202202202202202202202202202202 201201202202202202202202202302 20220220202020202020202020202 201201202202202202202202202302 022022020202020202020202302302 02202202202202202202202302302 022020202020202020202302302 20220220220220220220220220230

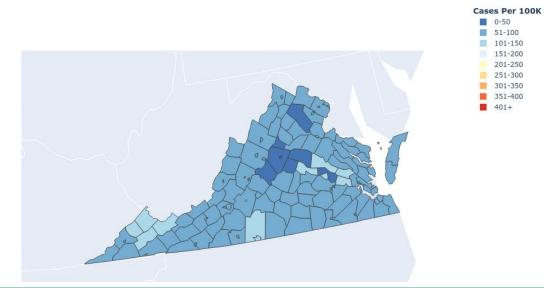
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Adaptive

Weekly Projections (Adaptive) 28-Sep-2022



Weekly Projections (Adaptive-VariantX) 28-Sep-2022



Cases Per 100K

0-50

51-100

101-150

151-200

201-250

251-300

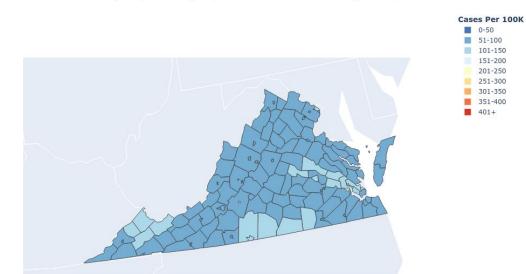
301-350

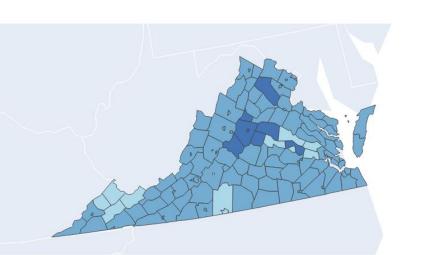
351-400

201-250

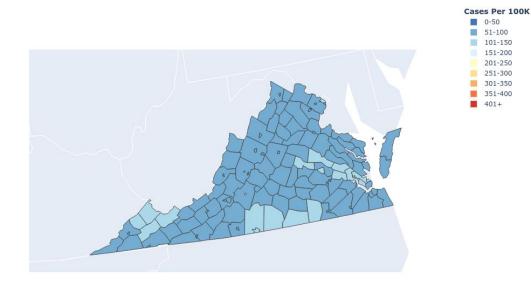
401+

Weekly Projections (Adaptive-VariantX-FallWinter) 28-Sep-2022



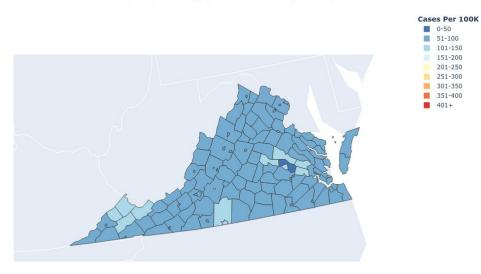


Weekly Projections (Adaptive-FallWinter) 28-Sep-2022



Impact of Optimistic vs. Pessimistic Booster Distribution

Weekly Projections (Optimistic Booster) 28-Sep-2022

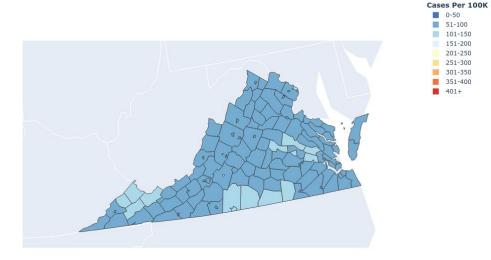


Weekly Projections (Pessimistic Booster) 28-Sep-2022

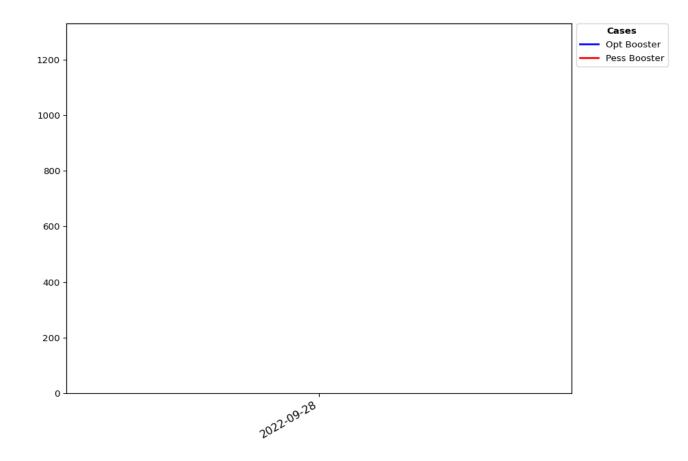
0-50

51-100 101-150 151-200 201-250 251-300 301-350

351-400 401+



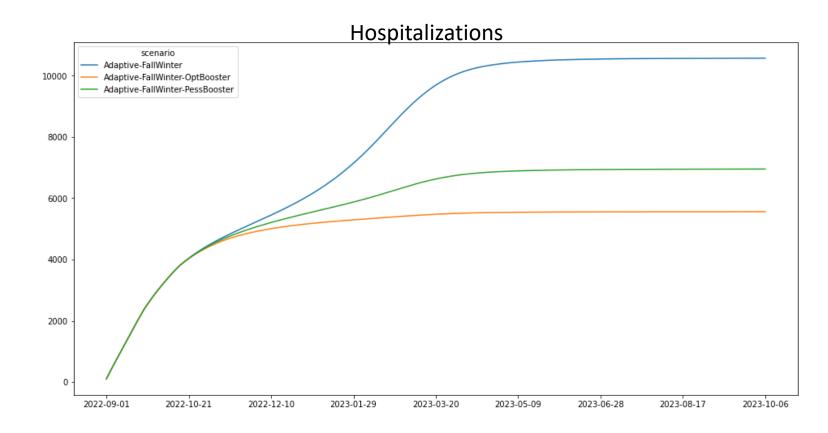
Cases for Optimistic vs. Pessimistic Boosters 28-Sep-2022



Booster Campaign Coverage has impact on future hospitalizations

Booster Campaign can significantly limit future hospitalizations and severe outcomes

 Reduction in future hospitalizations (3600 to 5K) through Spring 2023



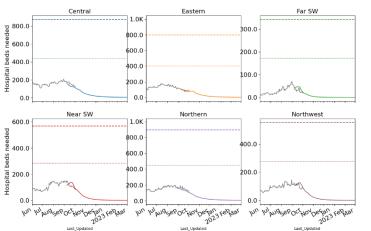


Hospital Demand and Bed Capacity by Region

Capacities by Region

COVID-19 capacity ranges from 80% (dots) to 120% (dash) of total beds

Adaptive



Adaptive – VariantX & Fall Winter

Northerr

In Julaugsepoct Nov Degan 3 Feb Mar

200.0

100.0

400.0

Northwes

600.0

400.0

00.0

0.0

1.0K

500.0

400.0

Length of Stay more variable with Omicron, occupancy projections may vary as a result, ad-hoc estimation performed per region Length of Stay Estimates

Central	8
Eastern	6
Far SW	4
Near SW	9
Northern	5
Northwestern	9

Estimated LOS shortened slightly to better fit observed data

Projections show continued declines and with expanded capacities and adjusted length of stay, no capacities exceeded



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Jun Jul Augsepoct Nov Degan Feb Mar

0.008 eeee

8 400.0

200.0

0.0

600.0

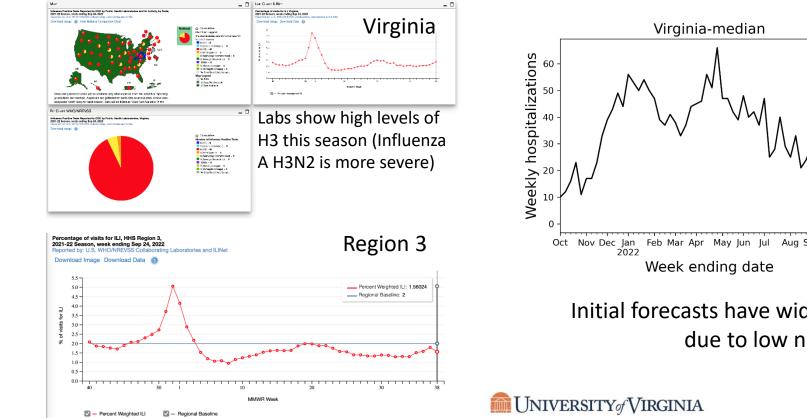
400.0

R 200.0

Current Influenza Hospitalization Forecast

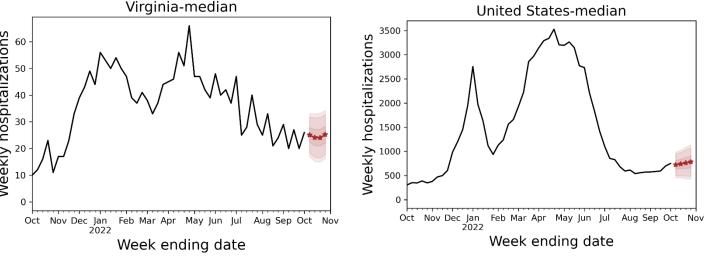
Statistical models for submitting to CDC FluSight forecasting challenge

• Similar to COVID-19 case forecasts, uses a variety of statistical and ML approaches to forecast weekly hospital admissions for the next 4 weeks for all states in the US



Influenza-like Illness Activity

Hospital Admissions for Influenza and Forecast for next 4 weeks (UVA ensemble)



Initial forecasts have wide uncertainty due to noisiness in data due to low numbers of hospitalizations

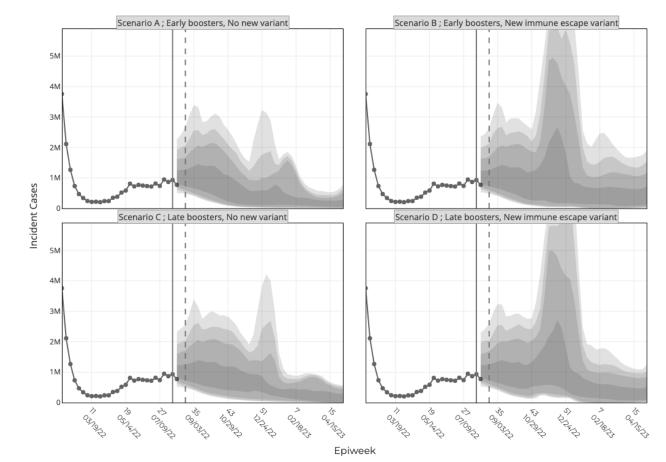
Scenario Modeling Hub – COVID-19 (Round 15)

Collaboration of multiple academic teams to provide national and stateby-state level projections for 4 aligned scenarios

- Round 15 results published
 - Scenarios: Test benefits of reformulated fall boosters w/ and w/out a new variant
 - Timing of reformulated boosters is one of the axes

https://covid19scenariomodelinghub.org/viz.html

Projected Incident Cases by Epidemiological Week and by Scenario for Round 15 - US (- Projection Epiweek; -- Current Week)



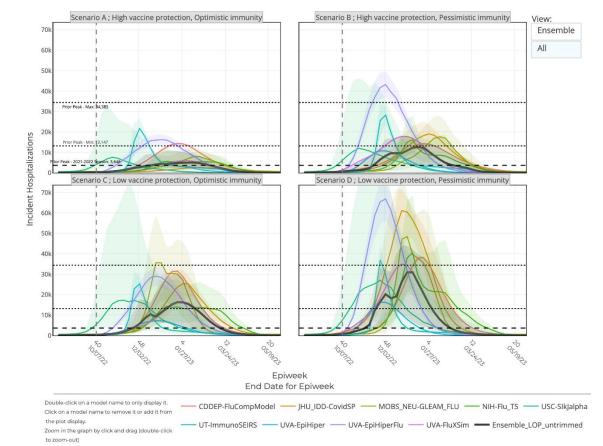
Scenario Modeling Hub – Influenza (Round 1)

Collaboration of multiple academic teams to provide national and state-bystate level projections for 4 aligned scenarios

- Round 1 results recently published
 - Impact of missed flu seasons on preseason immunity
 - Testing different seasonal vaccine coverage and efficacy
 - Projected from Aug 14th 2022
- High degree of uncertainty as previous 2 seasons have been irregular and there is still limited data for this season available
- Demonstrates importance of good vaccine coverage especially if previous immunity is weak

https://fluscenariomodelinghub.org/viz.html





Key Takeaways

Projecting future cases precisely is impossible and unnecessary. Even without perfect projections, we can confidently draw conclusions:

- Case rates continue their decline, hospitalizations continue steady decline
- VA weekly case rate continues decline to 99/100K from 109/100K
 - US weekly case rate is down considerably as well at 89/100K from 105/100K
 - VA hospital occupancy (rolling 7 day mean of 545 down from 599 a week ago) has continued to decline
- Projections anticipate continued declines in cases as well as hospitalizations
- Potential for rebounds due to seasonal forces and/or novel sub-variants in the Fall remains
- Model updates:
 - Maintained Booster Scenarios by slowing down the rate of vaccination adopting the rate of 3rd dose rollout
 - Current monitoring still not finding a definite candidate for Variant X, though BQ.1.1 or BA.2.75.2 remain likely, 50% prevalence pushed back to Nov 15th

The situation continues to change. Models continue to be updated regularly.

References

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NSSAC. PatchSim: Code for simulating the metapopulation SEIR model. <u>https://github.com/NSSAC/PatchSim</u>

Virginia Department of Health. COVID-19 in Virginia. <u>http://www.vdh.virginia.gov/coronavirus/</u>

Biocomplexity Institute. COVID-19 Surveillance Dashboard. <u>https://nssac.bii.virginia.edu/covid-19/dashboard/</u>

Google. COVID-19 community mobility reports. <u>https://www.google.com/covid19/mobility/</u>

Biocomplexity page for data and other resources related to COVID-19: <u>https://covid19.biocomplexity.virginia.edu/</u>



Questions?

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