

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

Estimation of COVID-19 Impact in Virginia

October 12th, 2022

(data current to September 24th – September 27th)

Biocomplexity Institute Technical report: TR BI-2022-1770



BIOCOMPLEXITY INSTITUTE

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



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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:

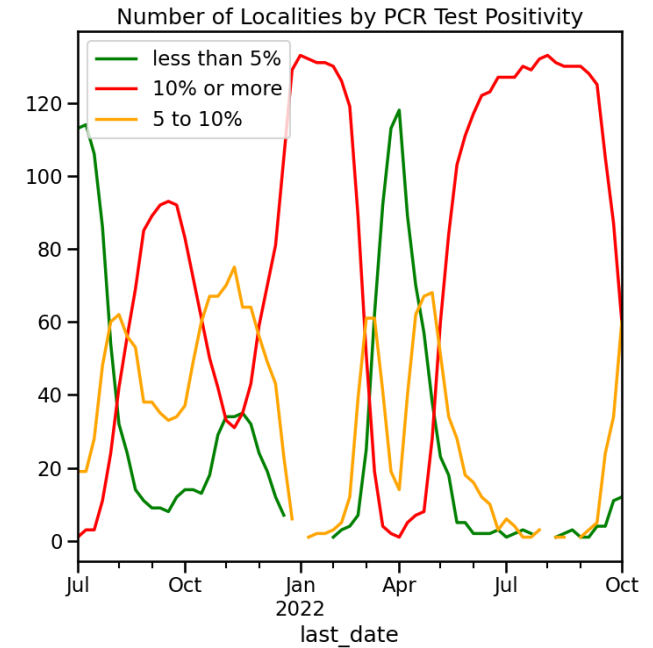
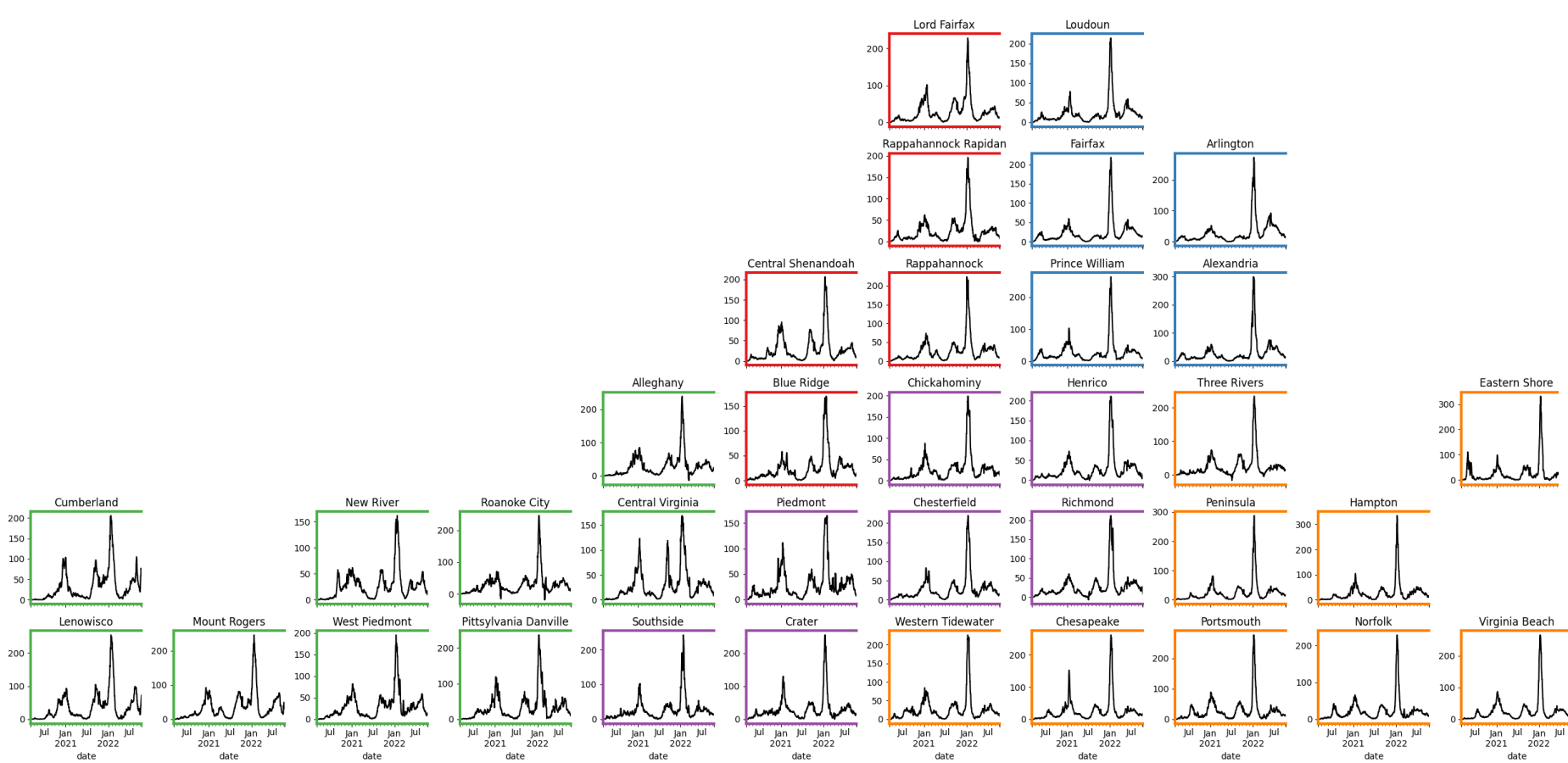
- **Case rates have continued to decline though hospitalizations have shown some recent growth**
- VA weekly case rate is slightly down at 96 per 100K from 99 per 100K
 - US weekly case rate is slightly down to 81 per 100K from 89 per 100K
 - VA hospital occupancy (rolling 7 day mean of 519 slightly down from 545 a week ago) down but experiencing recent activity
- Sub-variant prevalence evolution as expected
- Projections from last week remain largely on target with limited impact of Fall Winter scenario, however hospitalization trajectories have diverged

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

Situation Assessment



Case Rates (per 100k) and Test Positivity

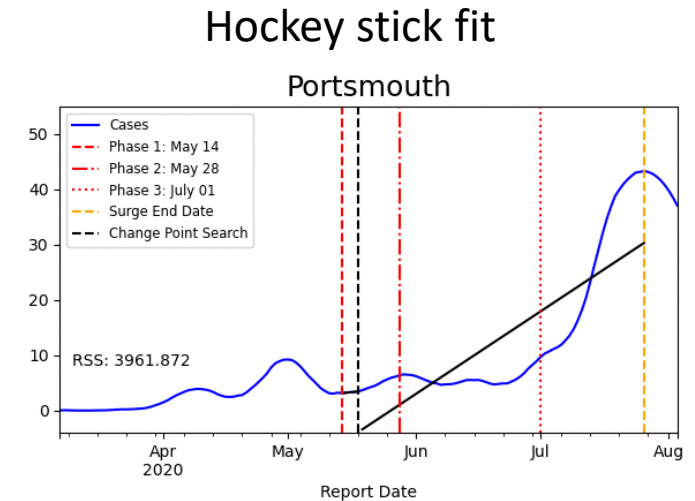


County level RT-PCR test positivity
Green: <5.0% (or <20 tests in past 14 days)
Orange: 5.0%-10.0% (or <500 tests and <2000 tests/100k and >10% positivity over 14 days)
Red: >10.0% (and not "Green" or "Yellow")

District Trajectories

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

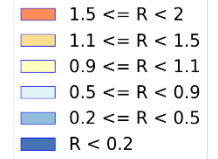
Method: Find recent peak and use hockey stick fit to find inflection point afterwards, then use this period's slope to define the trajectory



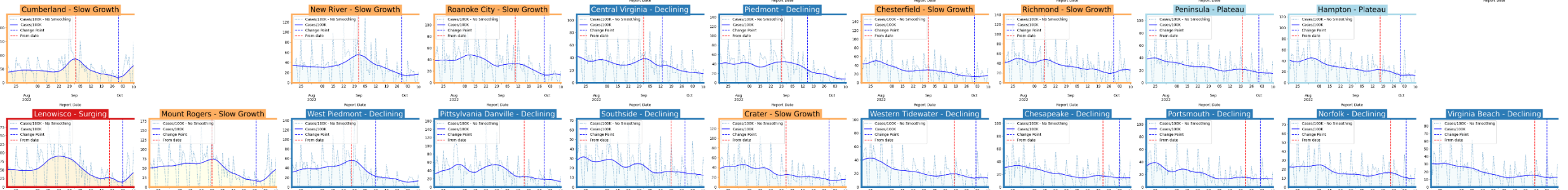
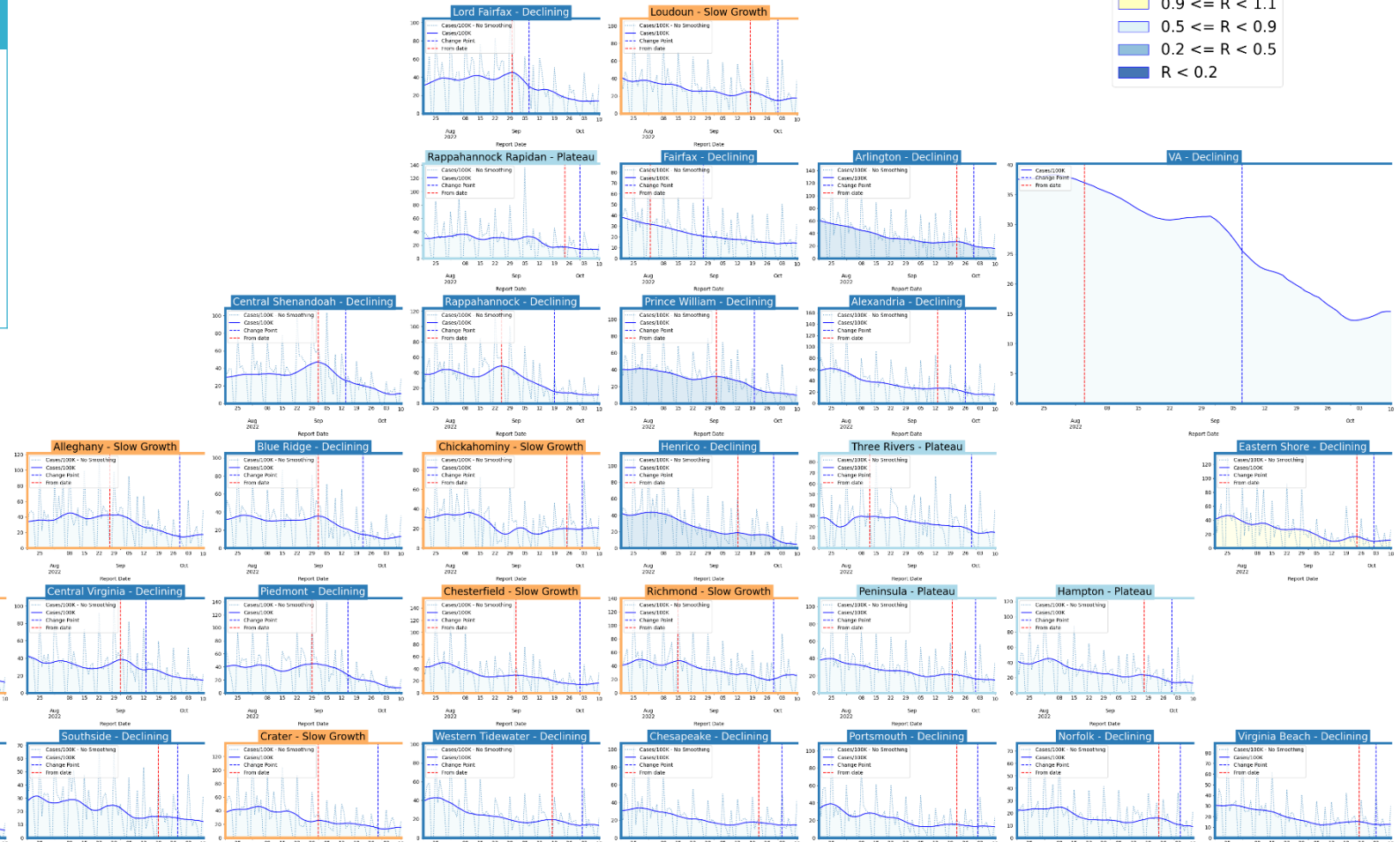
Trajectory	Description	Weekly Case Rate Slope (per 100k)	Weekly Hosp Rate Slope (per 100k)
Declining	Sustained decreases following a recent peak	slope < -0.88/day	slope < -0.07/day
Plateau	Steady level with minimal trend up or down	-0.88/day < slope < 0.42/day	-0.07/day < slope < 0.07/day
Slow Growth	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

Status	Number of Districts	
	Current Week	Last Week
Declining	20	(31)
Plateau	4	(3)
Slow Growth	10	(1)
In Surge	1	(0)



Curve shows smoothed case rate (per 100K)
 Trajectories of states in label & chart box
 Case Rate curve colored by Reproductive number

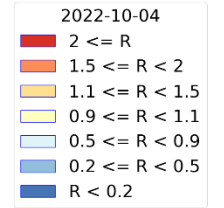


District Hospital Trajectories – last 10 weeks

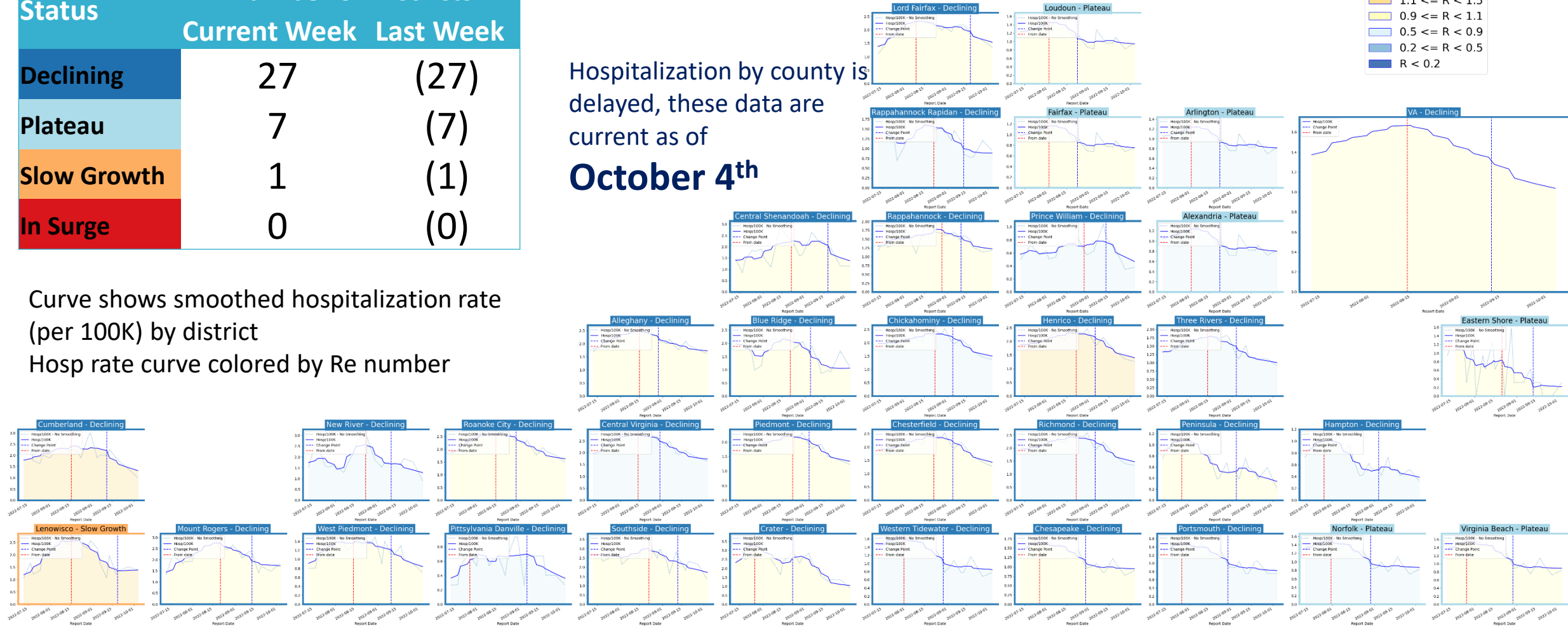
Status	Number of Districts	
	Current Week	Last Week
Declining	27	(27)
Plateau	7	(7)
Slow Growth	1	(1)
In Surge	0	(0)

Hospitalization by county is delayed, these data are current as of

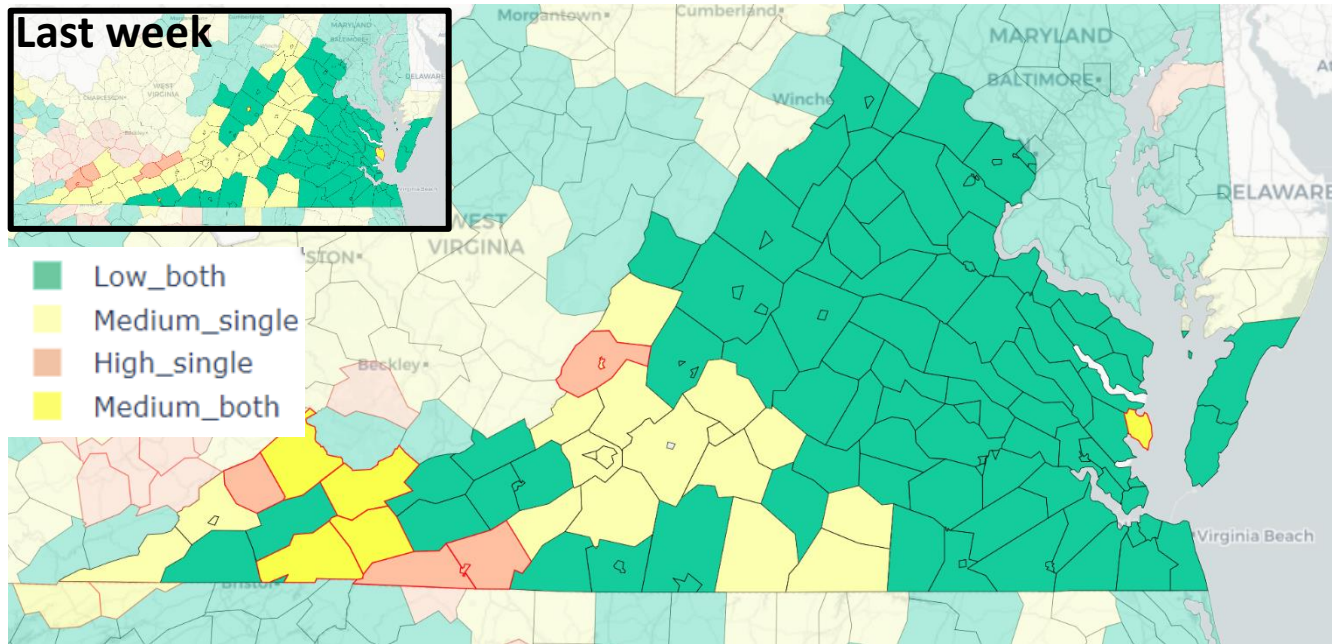
October 4th



Curve shows smoothed hospitalization rate (per 100K) by district
Hosp rate curve colored by Re number



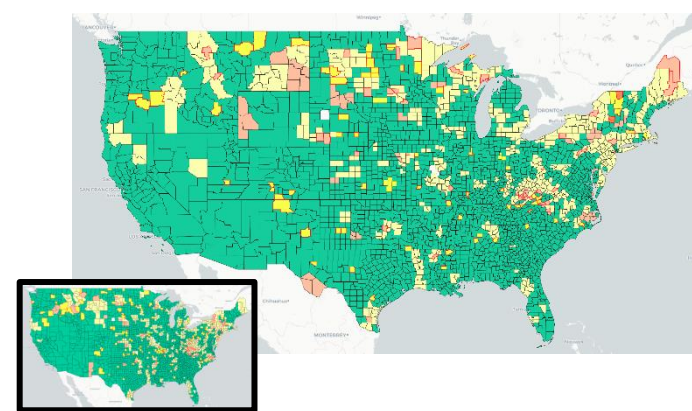
CDC's COVID-19 Community Levels



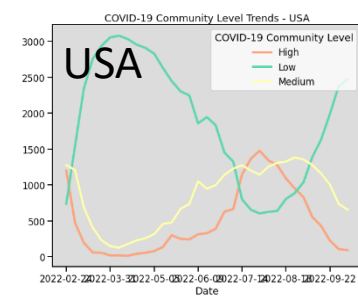
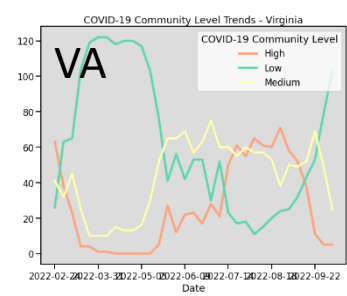
Red outline indicates county had 200 or more cases per 100k in last week

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



14-Oct-22

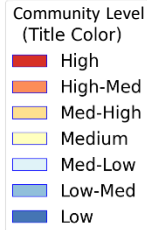


COVID-19 Community Levels - Use the Highest Level that Applies to Your Community				
New COVID-19 Cases Per 100,000 people in the past 7 days	Indicators	Low	Medium	High
Fewer than 200	New COVID-19 admissions per 100,000 population (7-day total)	<10.0	10.0-19.9	≥20.0
	Percent of staffed inpatient beds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

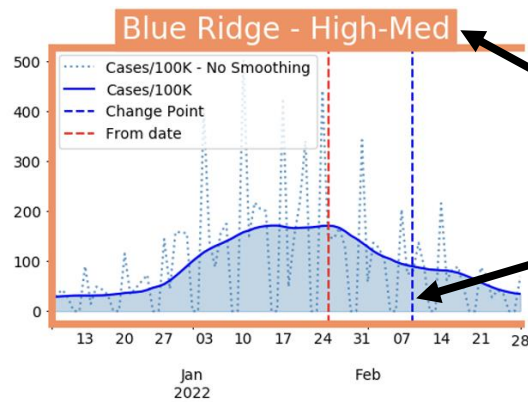
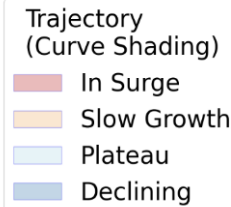
The COVID-19 community level is determined by the higher of the new admissions and inpatient beds metrics, based on the current level of new cases per 100,000 population in the past 7 days



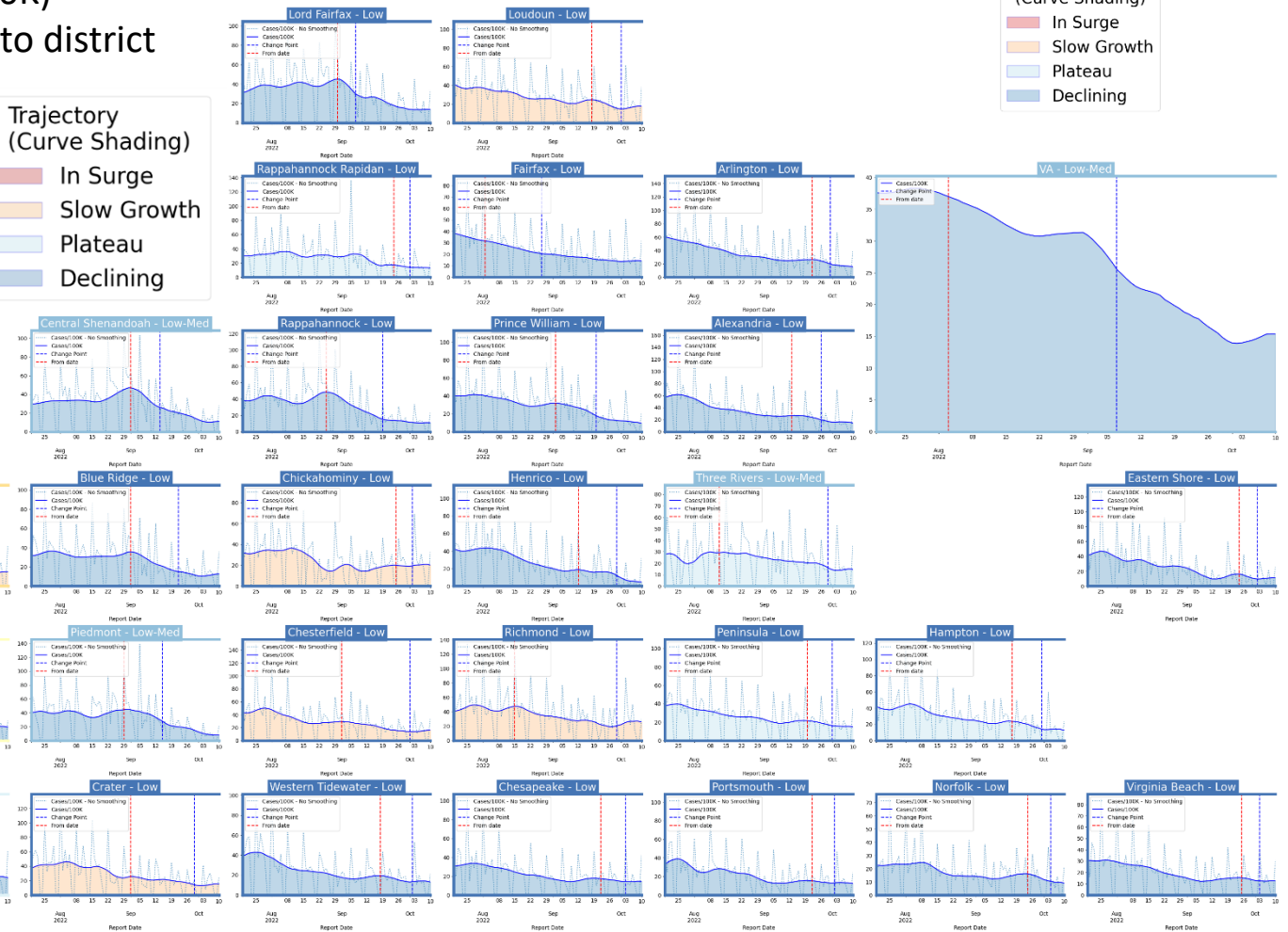
District Trajectories with Community Levels



Curve shows smoothed case rate (per 100K)
 CDC's new [Community Level](#) aggregated to district level in label & chart box color
 Case Rate curve colored by Trajectory



District's Aggregate Community Level
 Aggregate level a simple mean of all levels for counties in district
 Case rate Trajectory



Estimating Daily Reproductive Number – Redistributed gap

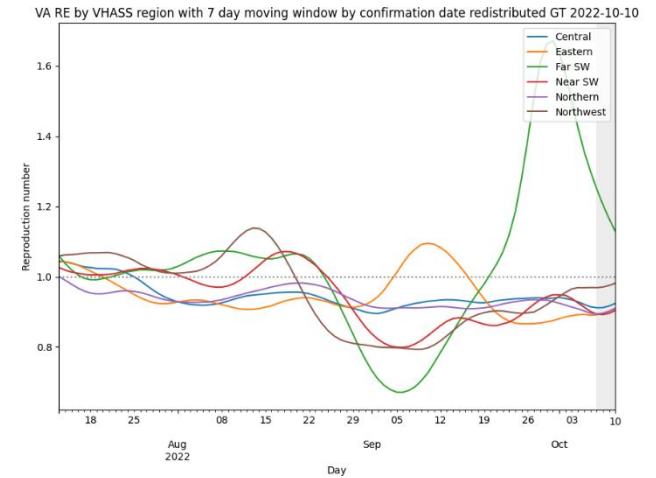
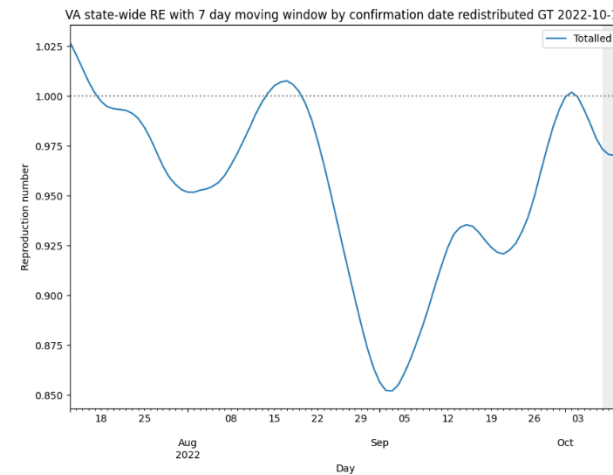
October 10th Estimates

Region	Date Confirmed R_e	Date Confirmed Diff Last Week
State-wide	0.972	0.107
Central	0.924	0.039
Eastern	0.911	0.050
Far SW	1.129	0.362
Near SW	0.904	0.036
Northern	0.910	-0.014
Northwest	0.982	0.210

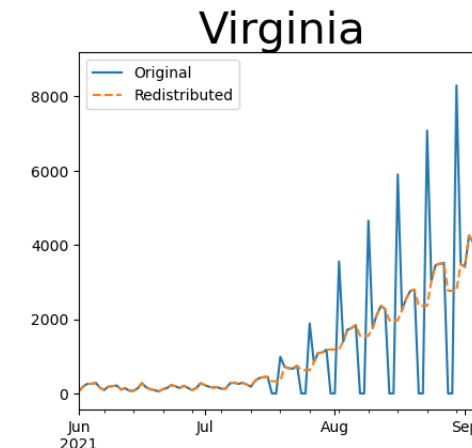
Methodology

- 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, <https://doi.org/10.1093/aje/kwt133>



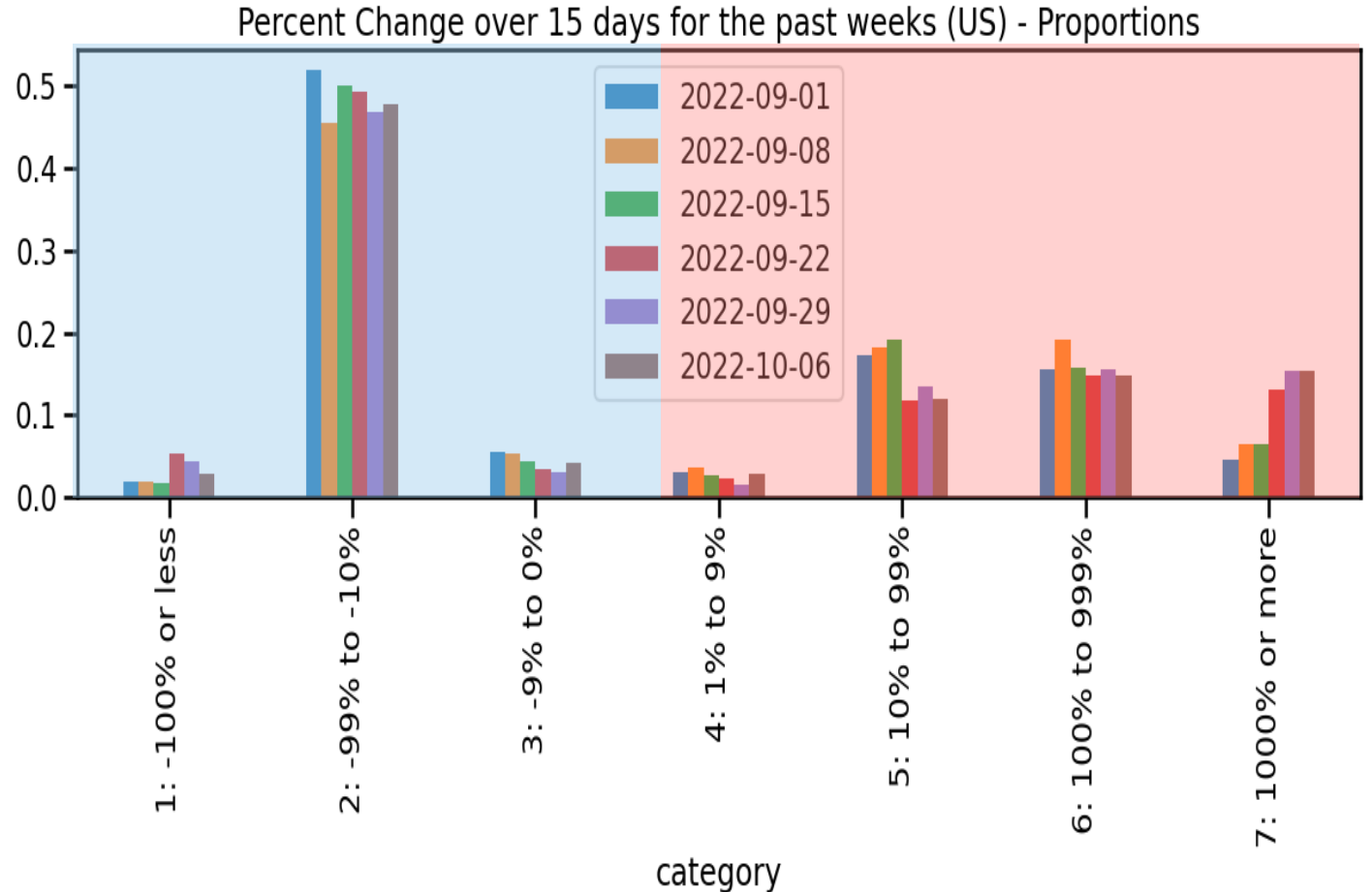
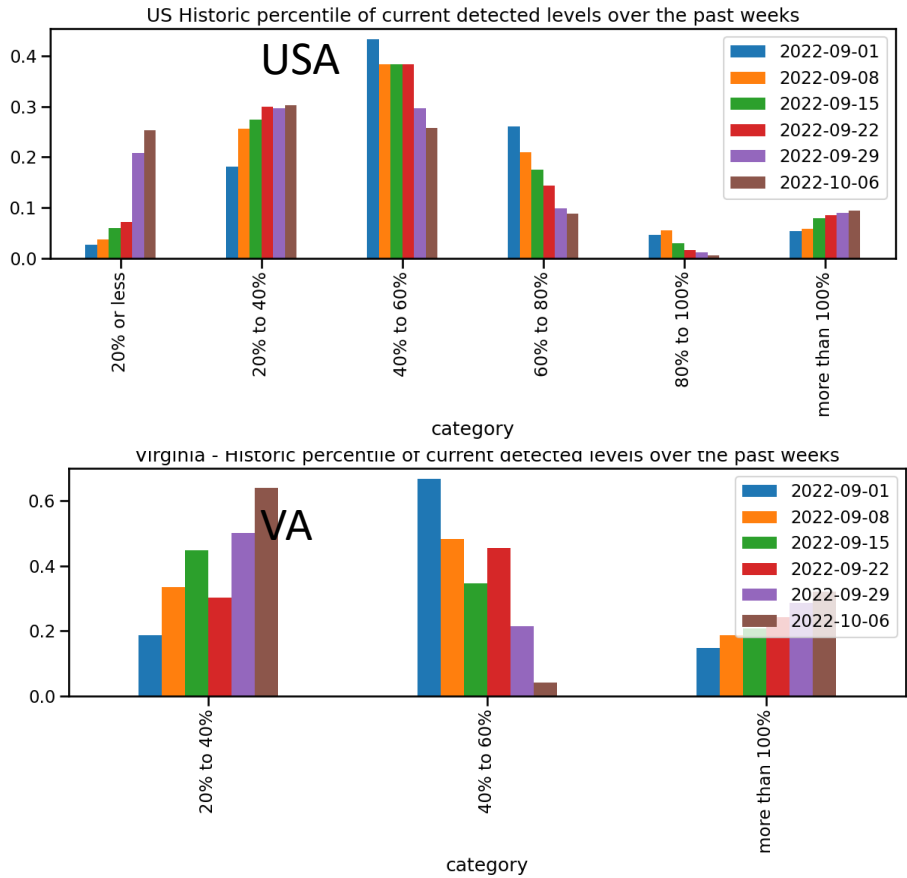
Skipping Weekend Reports & holidays biases estimates
Redistributed “big” report day to fill in gaps, and then estimate R from “smoothed” time series



Wastewater Monitoring

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

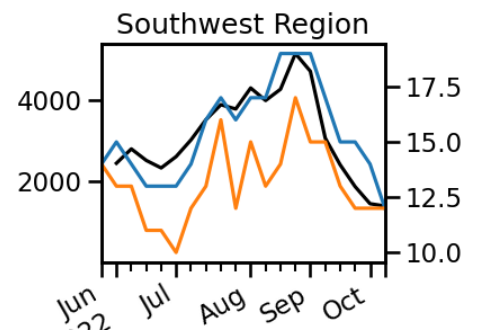
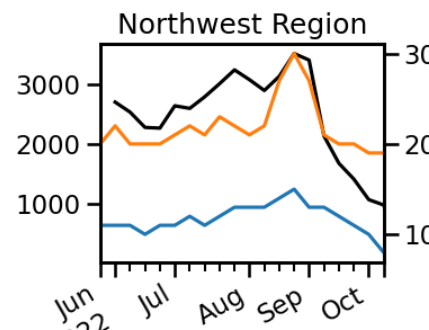
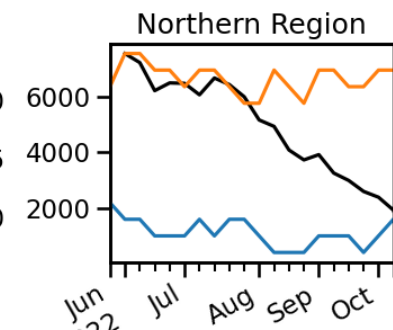
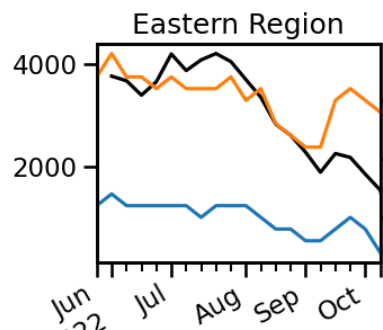
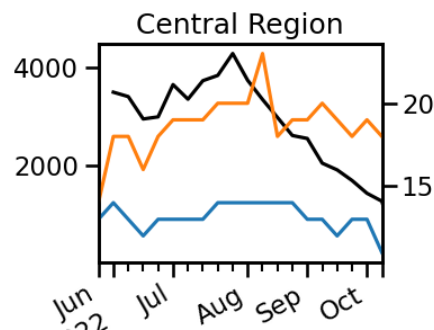
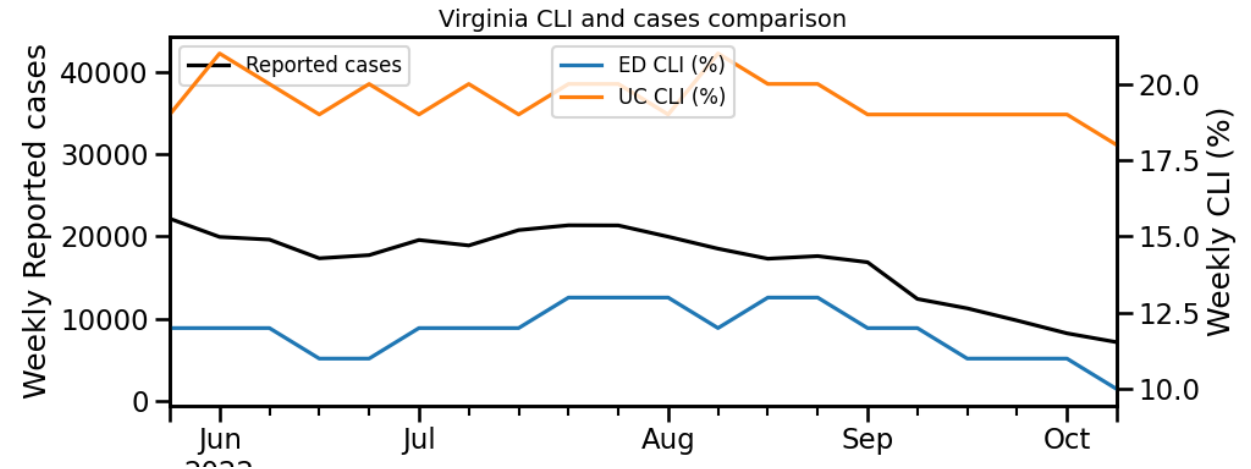
- Overall in the US, there is an increase in sites with increased levels of virus compared to 15 days ago
- Current virus levels are at or exceeding max of previous historical levels, has slowed, though more sites are entering upper quintiles



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 plateaued since May 2022, mixed by region**

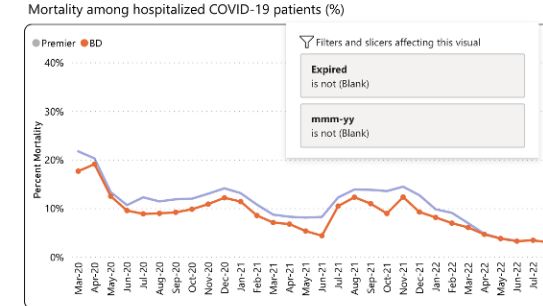
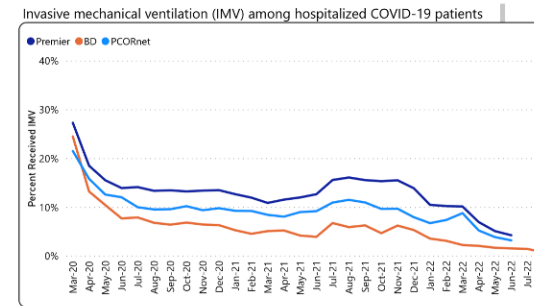
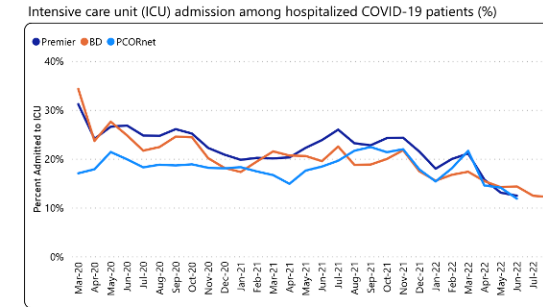
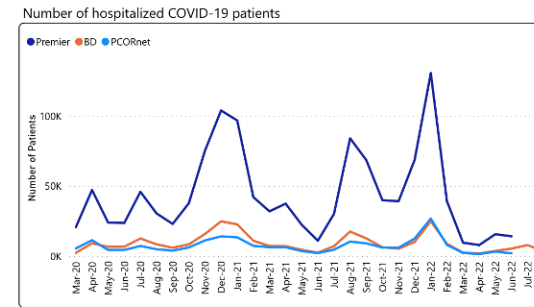


Hospitalizations and Severe Outcomes

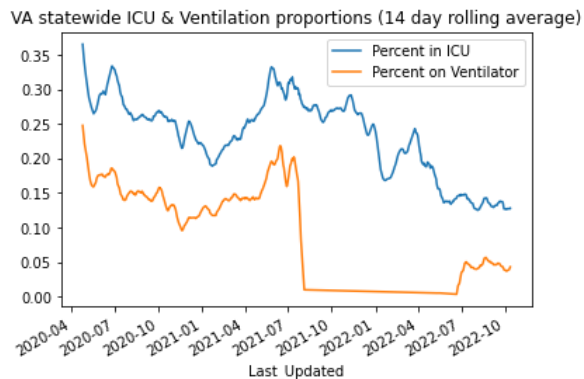
Data Source: [CDC Data Tracker](#)

Proportion of most severe outcomes decreasing among those who are hospitalized

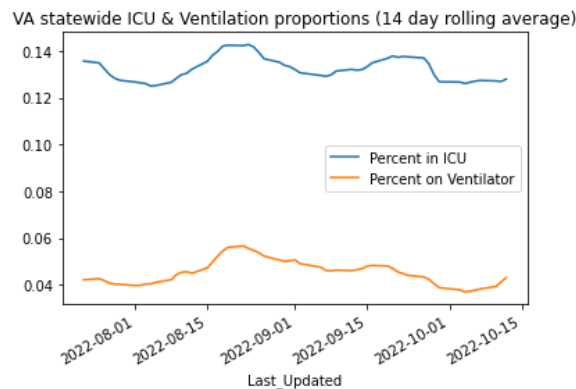
- ICU has declined from ~20% of hospitalized to nearly 10% since initial Omicron wave
- Also seen across all age-groups
- Similar levels of decline seen in VA
- Regionally more variation



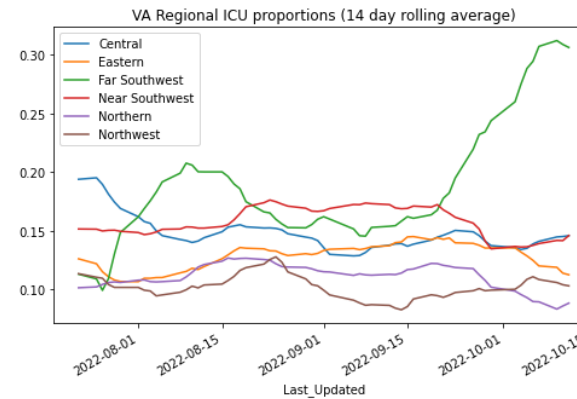
Virginia wide – full pandemic



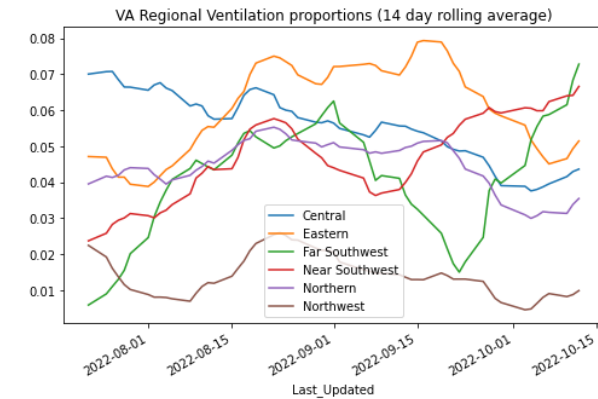
Virginia wide – recent



Virginia Regional ICU percent



Virginia Regional Ventilation %



SARS-CoV2 Variants of Concern

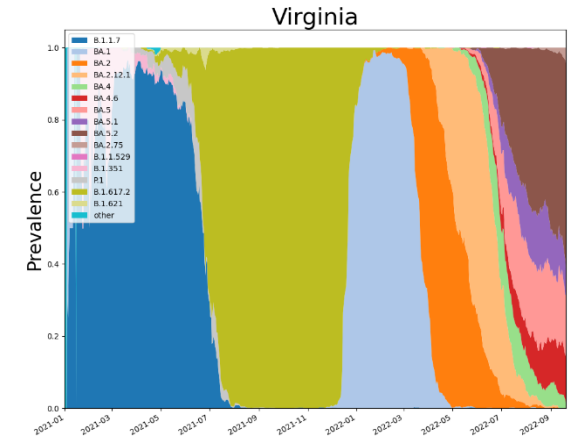
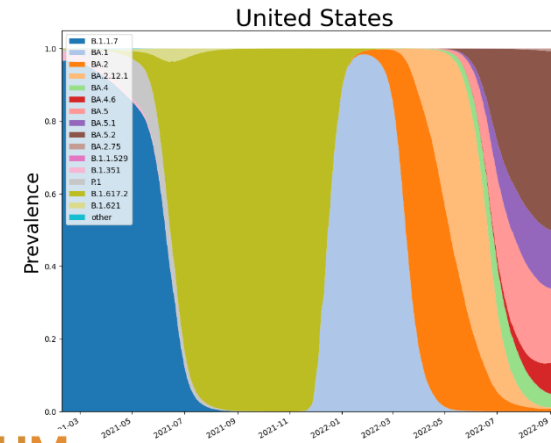


Emerging new variants will alter the future trajectories of pandemic and have implications for future control

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

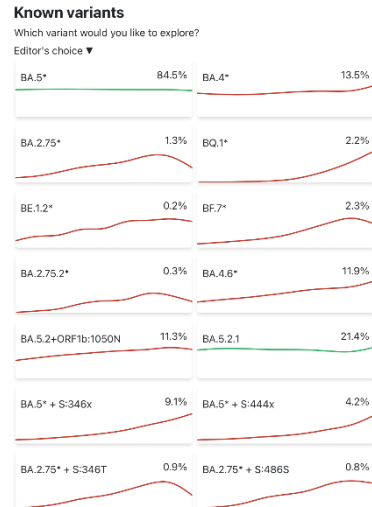
Omicron Updates

- 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 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

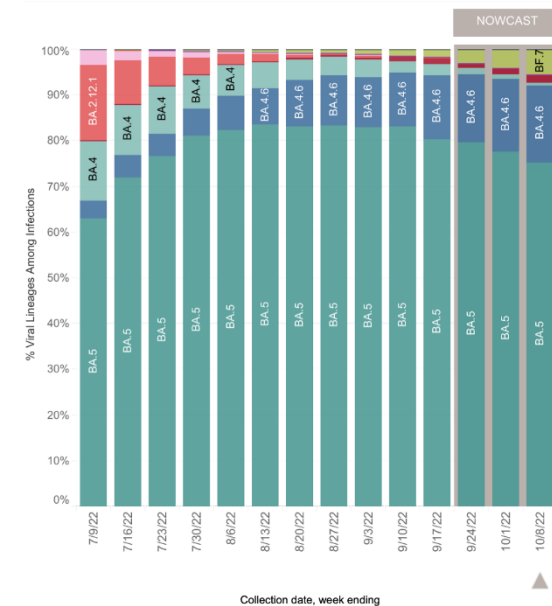
Enabled by data from GISAID



COV-spectrum

“Editor’s choice”
Variants to watch

HHS Region 3: 7/3/2022 – 10/8/2022



HHS Region 3: 10/2/2022 – 10/8/2022 NOWCAST

WHO label	Lineage #	US Class	%Total	95%PI
Omicron	BA.5	VOC	75.2%	72.8-77.5%
	BA.4.6	VOC	16.7%	14.1-19.7%
	BF.7	VOC	5.6%	4.1-7.5%
	BA.2.75	VOC	1.7%	1.1-2.5%
	BA.4	VOC	0.7%	0.6-0.9%
	BA.2.12.1	VOC	0.0%	0.0-0.0%
	BA.2	VOC	0.0%	0.0-0.0%
	B.1.1.529	VOC	0.0%	0.0-0.0%
	BA.1.1	VOC	0.0%	0.0-0.0%
Delta	B.1.617.2	VBM	0.0%	0.0-0.0%
Other	Other*		0.0%	0.0-0.0%

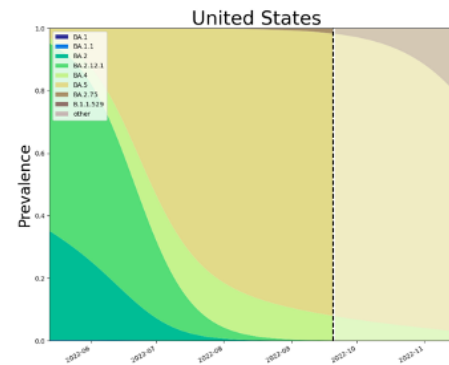
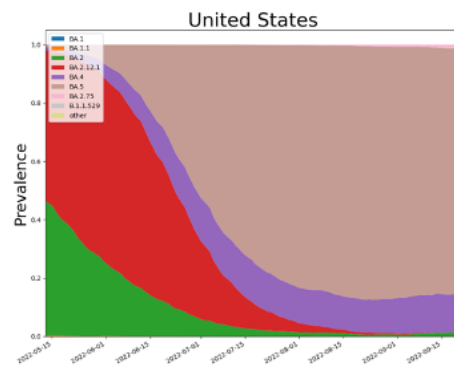
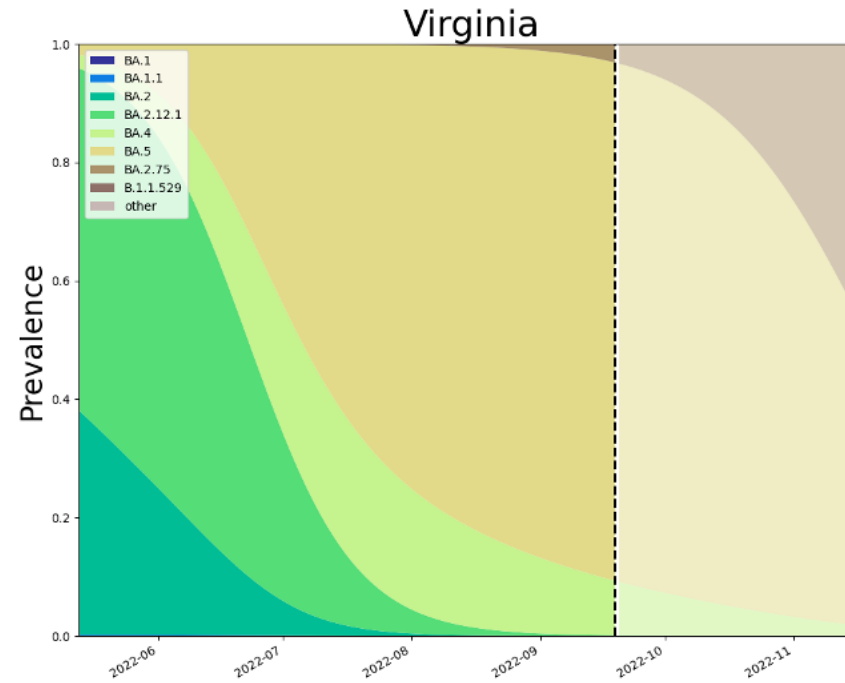
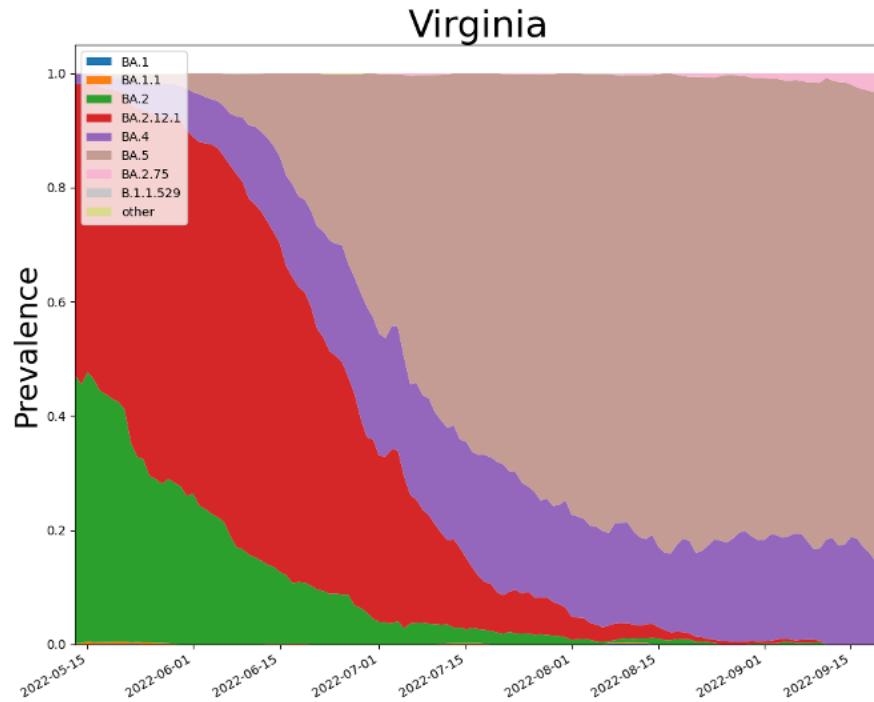
* Enumerated lineages are US VOC and lineages circulating above 1% nationally in at least one week period. "Other" represents the aggregation of lineages which are circulating <1% nationally during all weeks displayed.
** These data include Nowcast estimates, which are modeled projections that may differ from weighted estimates generated at later dates.
AY.1-AY.133 and their sublineages are aggregated with B.1.617.2. BA.1, BA.3 and their sublineages (except BA.1.1 and its sublineages) are aggregated 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 was aggregated with BA.5. Lineages BA.4.6, BF.7, and many BA.2.75 contain the spike substitution R346T.



SARS-CoV2 Omicron and Sub-Variants

As detected in whole Genomes in public repositories

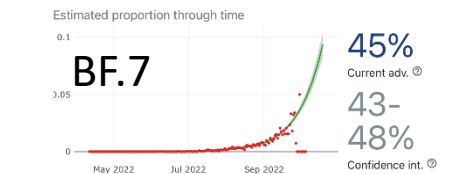
VoC Polynomial Fit Projections



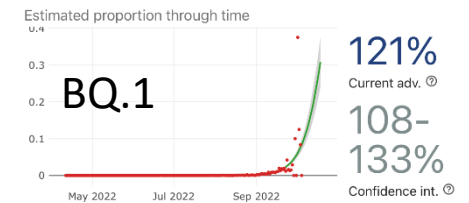
Relative Growth Advantages



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



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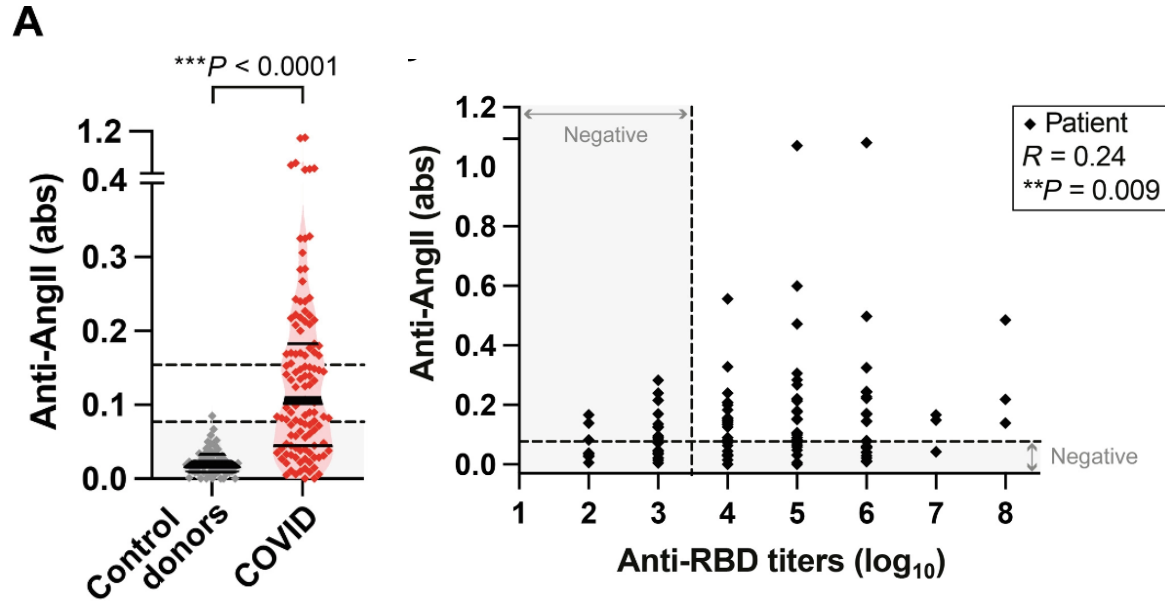
Note: Data lags force projections to start in past. Everything from dotted line forward is a projection.



14-Oct-22

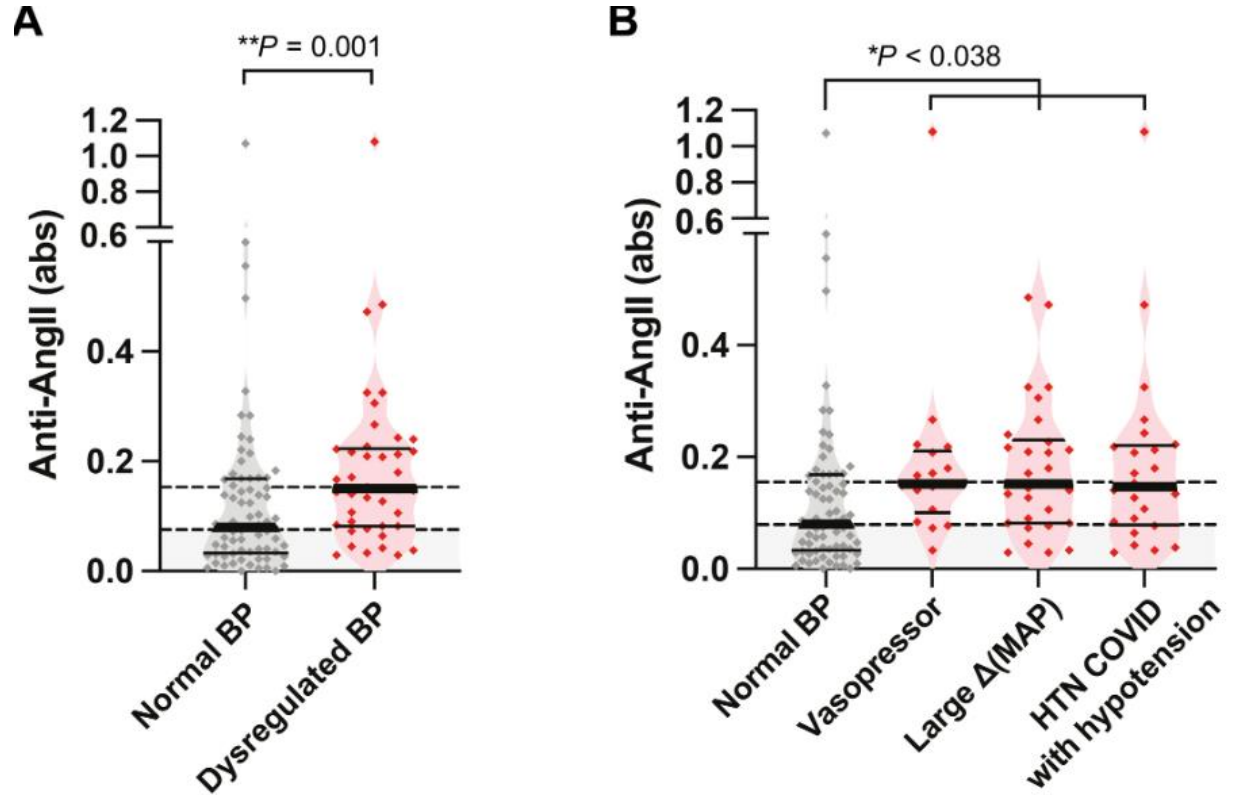
Pandemic Pubs

1. A significant proportion of hospitalized patients with COVID-19 developed autoantibodies against vasoconstrictive peptide in recent study



Chicago researchers show anti-angII antibodies can develop upon specific immune reaction to the SARS-CoV-2 proteins Spike or receptor-binding domain (RBD), to which they can cross-bind, suggesting some epitope mimicry between AngII and Spike/RBD. Angiotensin-converting enzyme 2 (ACE2), which is the main entry receptor of SARS-CoV-2 also tightly regulates blood pressure by converting the vasoconstrictive peptide angiotensin II (AngII) to a vasopressor peptide.

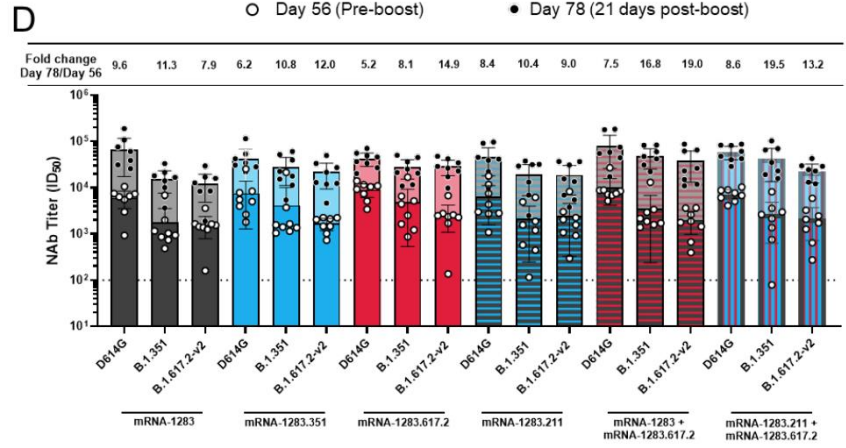
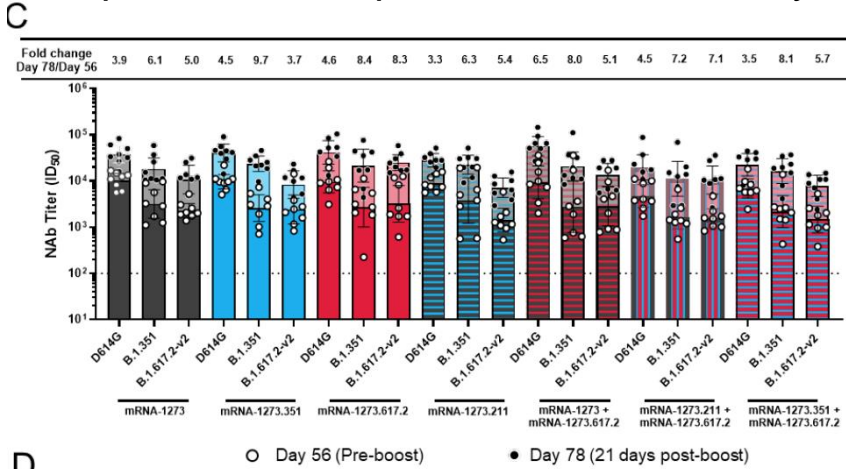
<https://www.science.org/doi/10.1126/sciadv.abn3777>



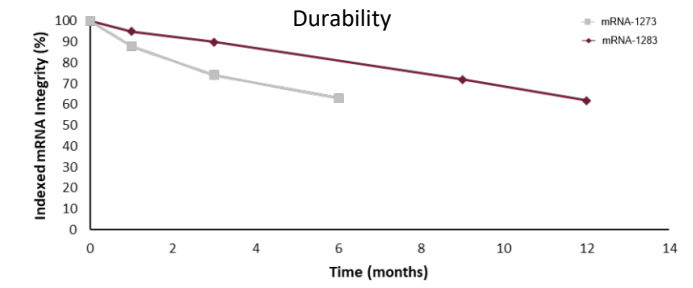
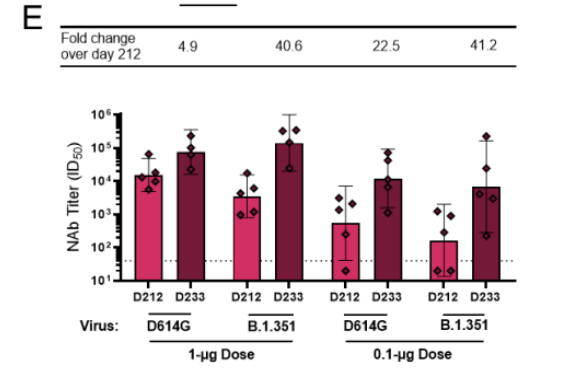
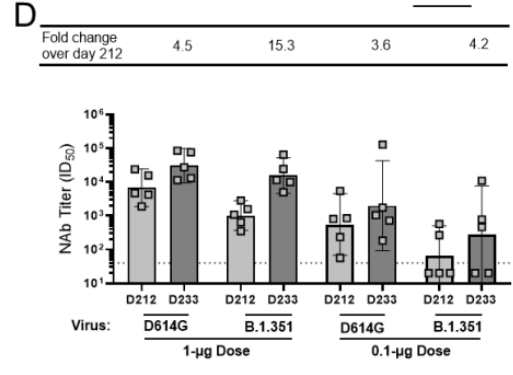
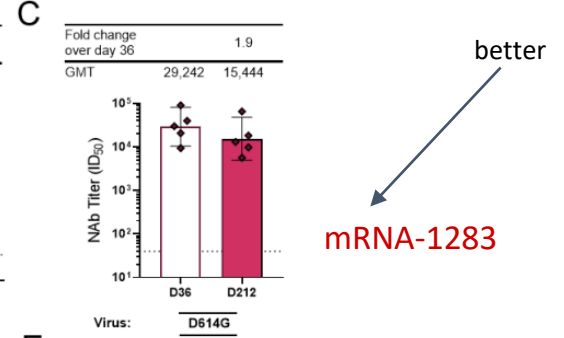
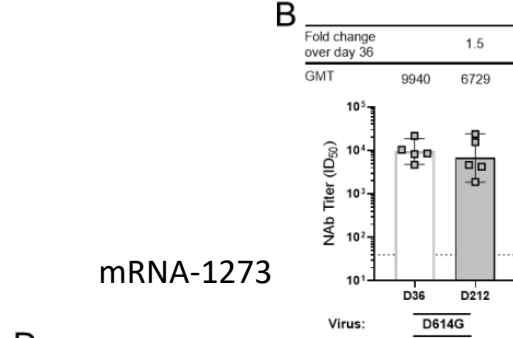
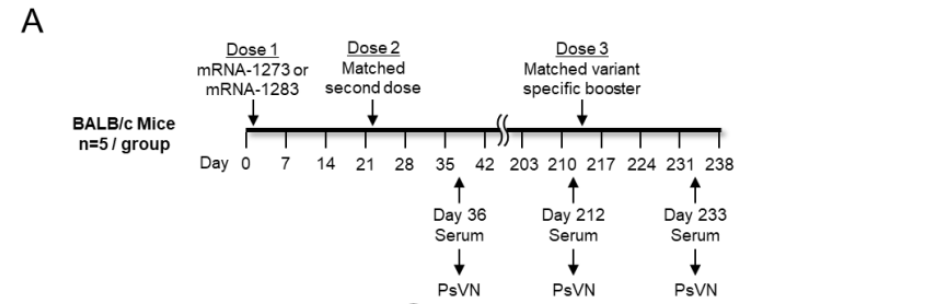
Levels of anti-AngII antibodies in COVID patients with normal BP as compared to patients with dysregulated BP, categorized as being under vasopressive drugs, exhibiting a large mean arterial pressure variability (large Δ MAP), or having experienced severe acute hypotensive episodes despite their known preexisting HTN condition (these categories are nonexclusive; Kruskal-Wallis test with Dunn's posttest comparison to normal BP)

Pandemic Pubs

2. An NTD-RBD linked candidate vaccine, mRNA-1283, showed improved antigen expression, antibody responses, and stability at refrigerated temperatures compared with the clinically available mRNA-1273



Immunogenicity in mRNA-1273- or mRNA-1283-primed mice receiving a matched B.1.351-specific booster. (A) Dosing schedule and sample collection. nAb titers in mice immunized with a 2-dose mRNA-1273 (1 µg; B) or mRNA-1283 (1 µg; C) primar series and against D614G and B.1.351 following mRNA-1273.351 (D) and mRNA-1283.351 (E) booster; N=5.



Researchers from Moderna present results that support clinical assessment of mRNA-1283 a domain-based mRNA vaccines encoding the wild-type spike-protein receptor-binding (RBD) and N-terminal domains (NTD). In mice administered mRNA-1283 as a primary series, booster, or variant-specific booster, similar or greater immune responses and protection from viral challenge were observed against wild-type, beta, delta, or omicron (BA.1) compared with mRNA-1273 immunized mice, especially at lower vaccine dosages.

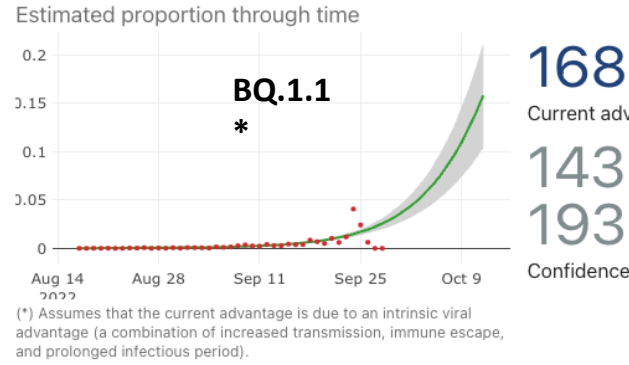
<https://www.biorxiv.org/content/10.1101/2022.10.07.511319v1>

Pandemic Pubs

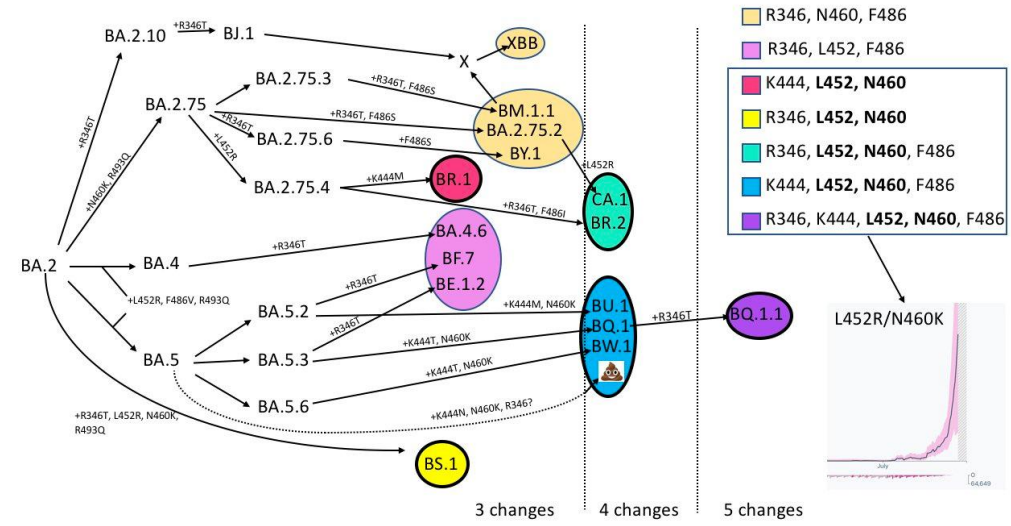
(last week)

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.

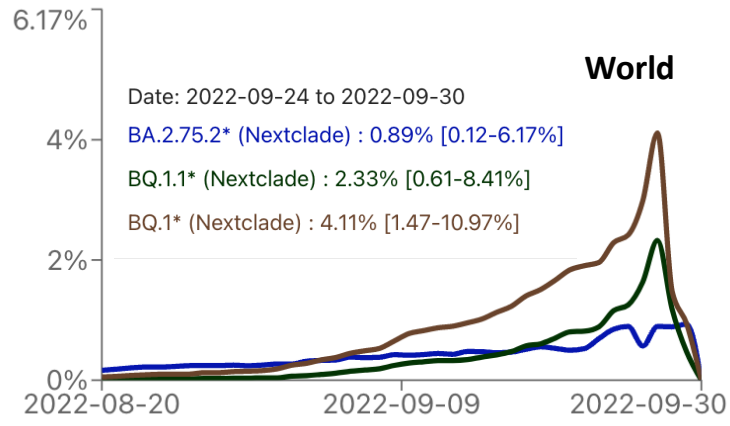
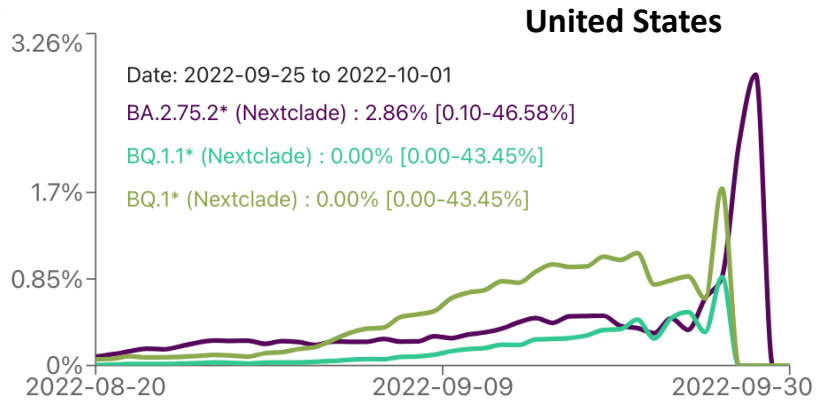


Omicron lineages with changes at 346, 444, 452, 460, and 486



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

World



description	number _sequen ces	submitted _past_10_ days	relative _growth _advant age	relative_growt h_ advantage_l ow	relative_grow th_ advantage _high
BA.4.6, BF.7, any BA.4/5+S:346T/S:44 etc.	67699	11538	31.64%	31.39%	31.89%
BA.2.75.2, BQ.1, BM.1.1, BY.1, BA.2.3.20, ...	3853	1465	73.37%	70.96%	75.78%
BQ.1.1, BN.1, BM.1.1.1, CA.1, ...	714	345	101.44%	93.21%	109.68%
XBB	66	40	157.47%	111.26%	203.67%

Estimates from surveillance using mutational profiles approximating BQ.1, BQ.1.1, and BA.2.75.2

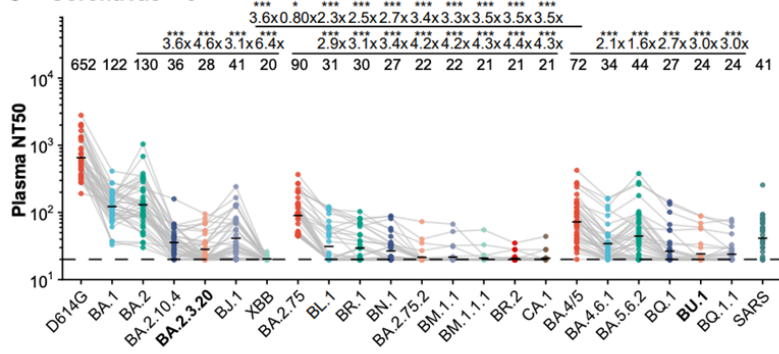
https://cov-spectrum.org/explore/World/AllSamples/from=2022-08-17&to=2022-10-04/variants?nextcladePangoLineage=BA.2.75.2*&nextcladePangoLineage1=bq.1.1*&nextcladePangoLineage2=bq.1&analysisMode=CompareEquals&

<https://cov-spectrum.org/collections/54>

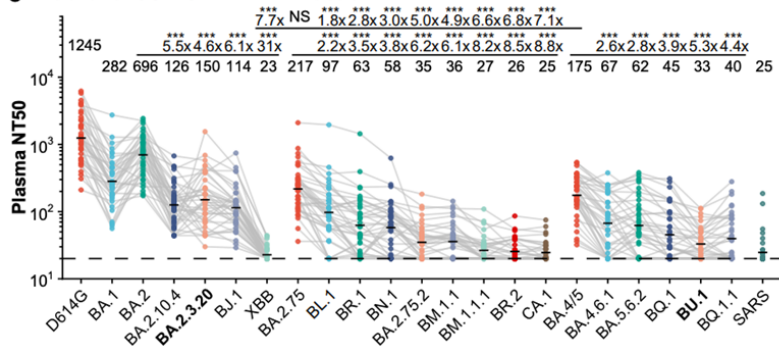
Pandemic Pubs (last week)

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.

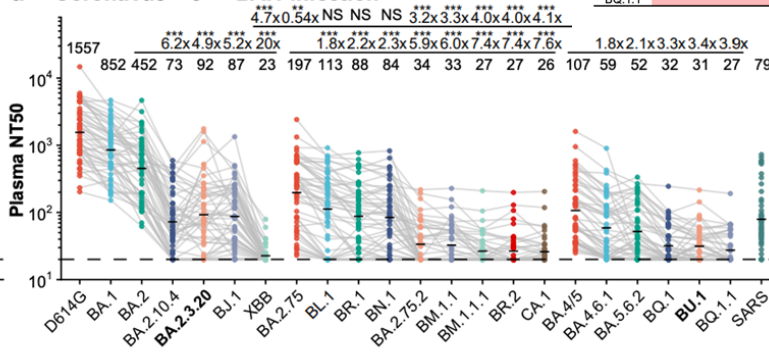
c CoronaVac × 3



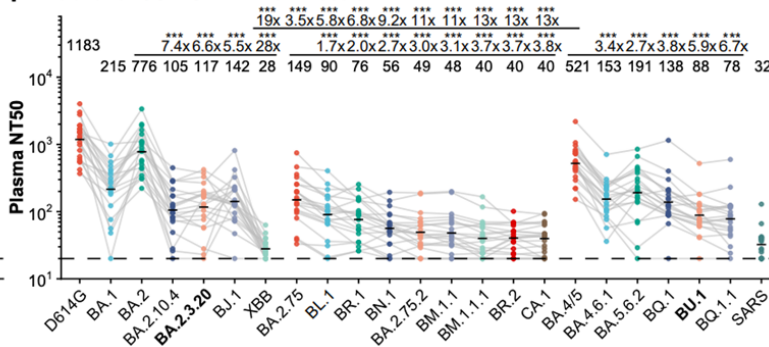
e CoronaVac × 3 → BA.2 infection



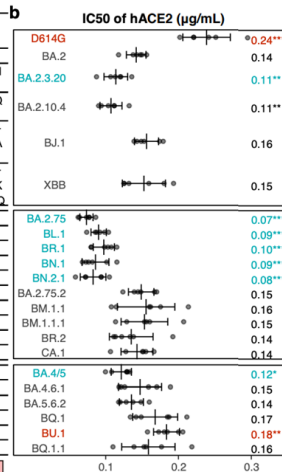
d CoronaVac × 3 → BA.1 infection



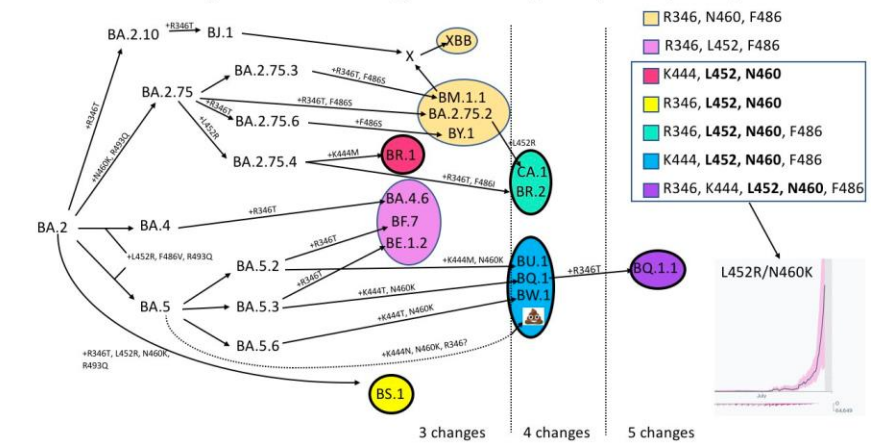
f CoronaVac × 3 → BA.5 infection



Pango lineages	REGN 10933	REGN 10987	REGN10933 +10987	COV2-2196	COV2-2130	COV2-2196+2130	BR1-196	BR1-198	BR1-196+198	S309	DXP-604	LY-CoV 1404	SA58	SA55	SA55+SA58	Additional RBD mutations	
BA.2	*	590	821	4312	2130	6.3	8.2	8530	8990	8610	852	219	0.9	5.1	7.2	7.8	
BA.2.3.20	121	*	199	15	*	26	14	*	24	*	897	181	9.7	20	4.6	7.8	K444R+N450D+L452M+N460K+R493Q
BA.2.10.4	*	*	*	289	501	2109	7990	3984	706	6348	1.3	4.3	4.9	5.0			G446S+F486P+R493Q+S494P
BJ.1	*	*	*	3076	5985	7609	*	*	*	709	166	*	8163	3.7	8.6		D399H+R346T+L388I+V445P+G446S+N460K+F486S
XBB	*	*	*	*	*	*	*	*	*	963	*	*	8805	5.3	9.8		D399H+R346T+L388I+V445P+G446S+N460K+F486S+F490S+R493Q
BA.2.75	278	*	410	119	352	121	1730	6622	3861	672	5920	2.2	246	4.3	9.6		
BL.1	260	*	511	93	*	174	1251	*	3075	508	7193	2.8	7975	6.3	10		R346T
BR.1	319	*	679	117	*	170	1992	*	3160	564	6689	*	1616	5.9	9.7		L452R+K444M
BN.1	390	*	701	59	303	109	4101	*	8444	6979	8901	1.7	4960	5.7	9.4		K356T+F490S
BN.1	344	*	599	70	*	166	3683	*	7791	*	6012	3.3	8295	4.9	9.0		R346T+K356T+F490S
BA.2.75.2	*	*	*	*	*	*	*	*	*	852	*	3.0	8922	5.9	9.7		R346T+F486S
BM.1.1	*	*	*	*	*	*	*	*	*	879	*	2.3	8823	5.2	8.9		R346T+F486S
BM.1.1.1	*	*	*	*	*	*	*	*	*	956	*	1.9	8082	4.8	10.5		R346T+F486S+F490S
BR.2	*	*	*	*	*	*	*	*	*	921	*	2.6	7263	4.7	10.5		R346T+L452R+F486I
CA.1	*	*	*	*	*	*	*	*	*	897	*	3.2	6927	6.0	11.5		R346T+L452R+F486S
BA.4/5	520	709	*	23	40	7124	*	*	1055	6264	0.8	3.9	5.0	4.5			
BA.4.6.1	2338	5402	*	*	*	4763	*	*	7809	4456	4634	1.2	50	4.8	9.9		R346T
BA.5.6.2	*	*	*	*	*	4636	*	*	7883	1408	5892	1662	58	5.1	8.9		K444T
BQ.1	*	*	*	*	*	*	*	*	1709	1905	44	6.6	9.2				K444T+N460K
BU.1	*	*	*	*	*	*	*	*	1082	26	5.3	10.5					K444M+N460K
BQ.1.1	*	*	*	*	*	*	*	*	5581	*	900	5.9	10.3				R346T+K444T+N460K



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



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 BA.2 and BA.5.

<https://www.biorxiv.org/content/10.1101/2022.09.15.507787v3>

https://twitter.com/yunlong_cao/status/1577343549120872448

<https://twitter.com/dfocosi/status/1574319177602670592?s=12&t=myre59DuQ1H31TI-yd7DQ>

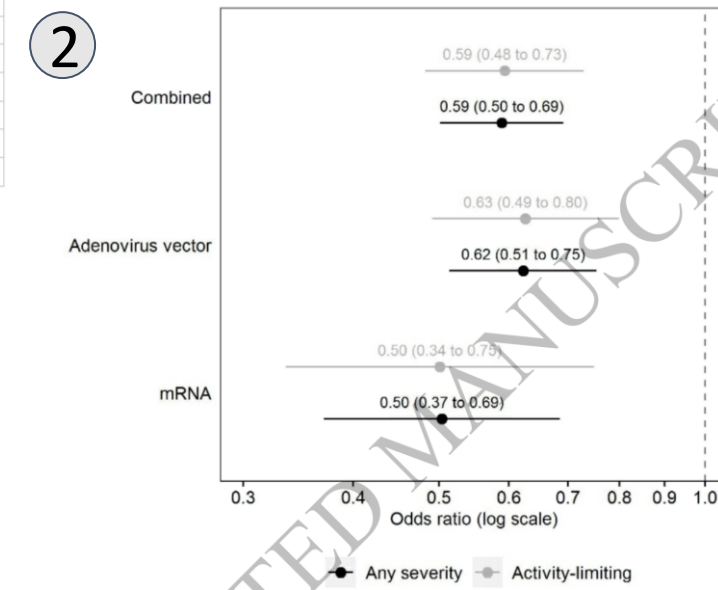
Pandemic Pubs

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 ~6.5-times 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

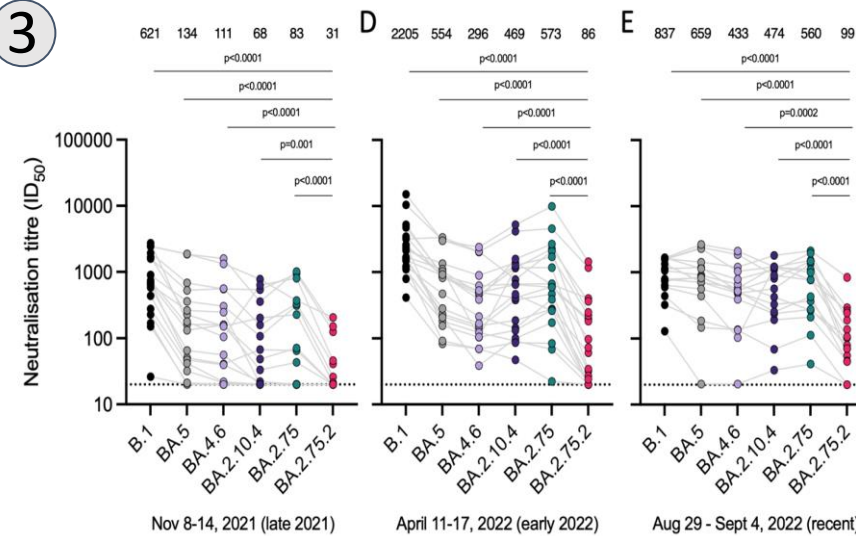
1

	Prevalence of Long COVID Among All Respondent (with or without COVID)						
	Total	Long COVID	Crude Prevalence of Long COVID % (95% CI)	Age and sex direct-standardized prevalence of long COVID* % (95% CI)	Crude prevalence ratio (PR) PR (95% CI)	Adjusted prevalence ratio (aPR)** aPR (95% CI)	Estimated Number with Long COVID
	Weighted N (%)	Weighted N (%)					
Total	3,042 (100.0)	222 (100.0)	7.3 (6.1, 8.5)				18,533,864
Age							
18-24	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
25-34	547 (18.0)	58 (26.1)	10.6 (6.6, 14.6)	10.0 (6.8 - 14.6)	Ref	Ref	4,837,339
35-44	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
45-54	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
55-64	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
65+	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)	2,261,131
Gender							
Male	1,443 (47.4)	72 (32.4)	5.0 (3.6, 6.3)	5.0 (3.8 - 6.5)	Ref	Ref	6,004,972
Female	1,516 (49.8)	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)	12,009,944

2



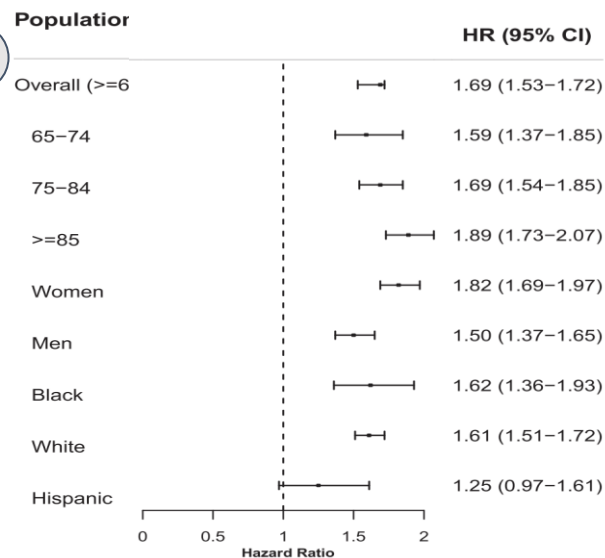
3



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=sl45IDMQ7G6mC2KakXPNZg>

4



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

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% CI: 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.

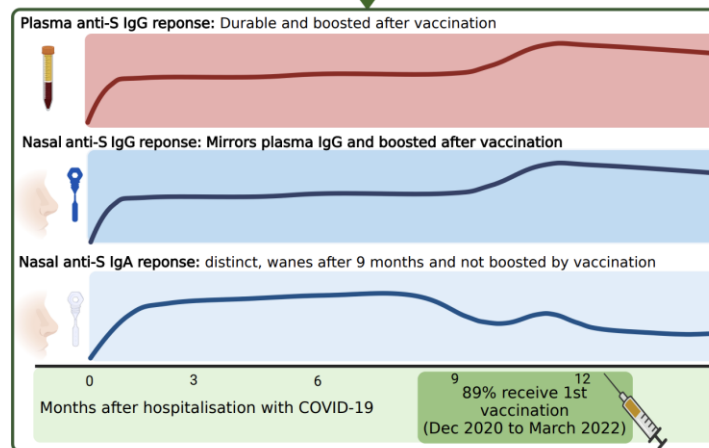
<https://www.medrxiv.org/content/10.1101/2022.09.12.22279862v1>

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>

Pandemic Pubs (last week) 1

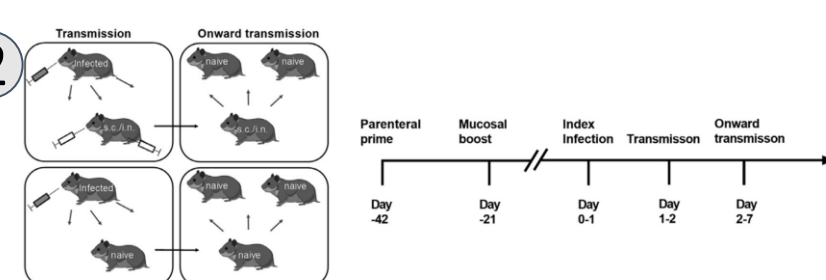
446 patients hospitalised with COVID-19 (Feb 2020 to March 2021)



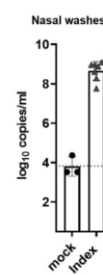
Plasma and nasosorption samples were prospectively collected from 446 adults hospitalised for COVID-19 between February 2020 and March 2021. IgA and IgG responses to NP and S of ancestral SARS-CoV-2, Delta and Omicron (BA.1) variants were measured by electrochemiluminescence and compared with plasma neutralisation data. **Nasal antibody induced by infection with pre-Omicron variants, bind Omicron virus in vitro better than plasma antibody. Although nasal and plasma IgG responses were enhanced by vaccination, Omicron binding responses did not reach levels equivalent to responses for ancestral SARS-CoV-2.** Results show that nasal IgA declines and has a minimal response to vaccination while plasma antibody responses to S antigen are well maintained and boosted by vaccination. Authors highlight the need to develop vaccines that enhance nasal immunity

<https://www.medrxiv.org/content/10.1101/2022.09.09.22279759v1>

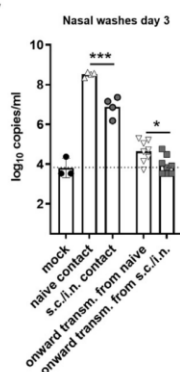
2



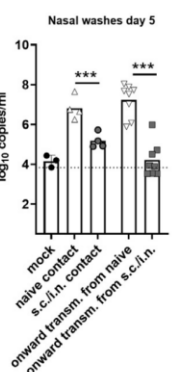
b



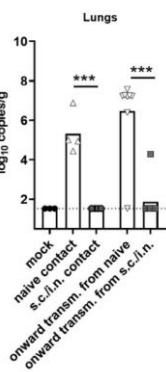
c



d



e



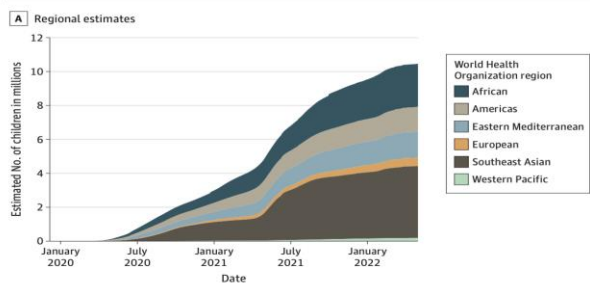
Center for Vaccine Research, Denmark: Study suggests mucosal booster strategy using protein-based subunit vaccines may be an effective means to protect against transmission of SARS-CoV-2 and potentially other respiratory viruses despite incomplete clearance of virus from the upper respiratory tract.

[https://www.thelancet.com/journals/ebiom/article/PIIS2352-3964\(22\)00430-3/fulltext](https://www.thelancet.com/journals/ebiom/article/PIIS2352-3964(22)00430-3/fulltext)

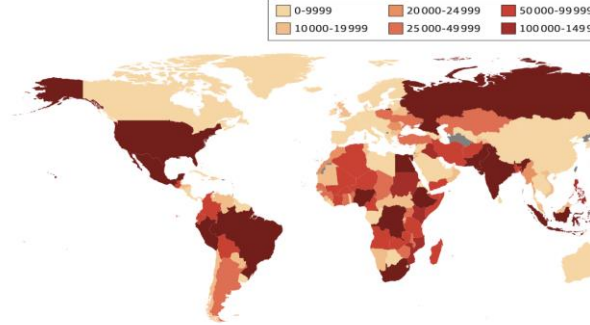
1. Nasal immune response to Omicron enhanced but not completely restored to ancestral levels when boosted using single strain booster.
2. Hamsters: intranasal boost elicited high-magnitude serum neutralizing antibody responses and IgA responses in the upper respiratory tract. Protected against virus in the lower airways and against onward SARS-CoV-2 transmission.
3. Impaired diffusion from air to red blood cells in 39% patients (6 months after infection) and 31% (12 months).
4. Estimated 10 million children orphaned due to primary and secondary caregiver deaths from COVID worldwide.

4

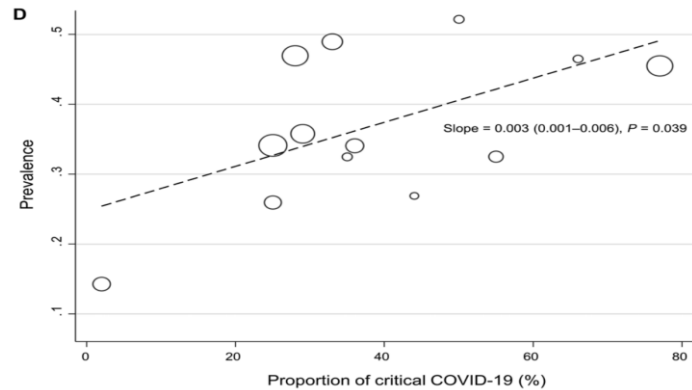
Figure. World Health Organization (WHO) Regional and National Estimates of Orphanhood and Primary and/or Secondary Caregiver Loss From January 1, 2020, Through May 1, 2022



National estimates



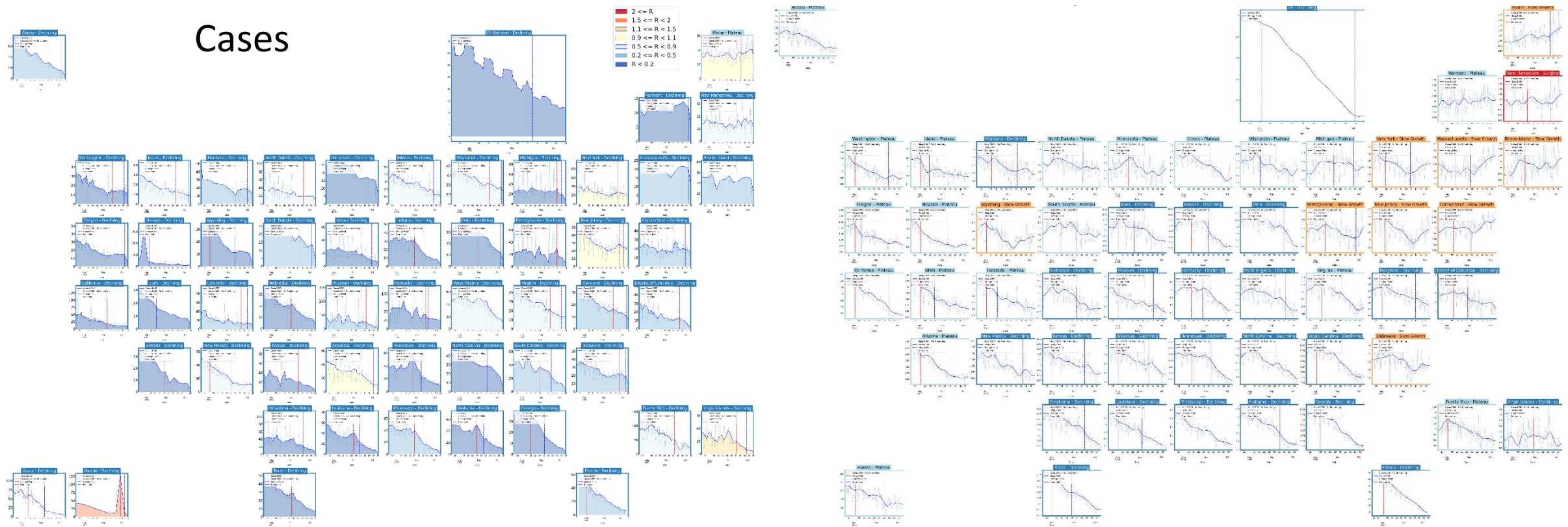
3



Systematic review and meta-analysis using a random-effects model to estimate the pooled prevalence of the pulmonary sequelae of COVID-19, as demonstrated by pulmonary function testing (PFT) and chest computed tomography (CT) performed at least 6 months after initial infection. A substantial number of COVID-19 survivors displayed pulmonary sequelae as part of PACS. Except for restrictive pulmonary dysfunction, the prevalence of these sequelae did not decrease until 1 year after initial infection. Of the 18,062 studies identified, 30 met eligibility criteria. Among these studies, 25 and 22 had follow-up PFT and chest CT data, respectively. **The follow-up durations were approximately 6 and 12 months in 18 and 12 studies, respectively. Impaired diffusion capacity was the most common abnormality on PFT (pooled prevalence 35%, 95% confidence interval [CI] 30–41%) with a prevalence of 39% (95% CI 34–45%) and 31% (95% CI 21–40%) in the 6-month and 12-month follow-up studies, respectively (P = 0.115)**

<https://respiratory-research.biomedcentral.com/articles/10.1186/s12931-022-02163-x>

United States Case & Hospitalizations



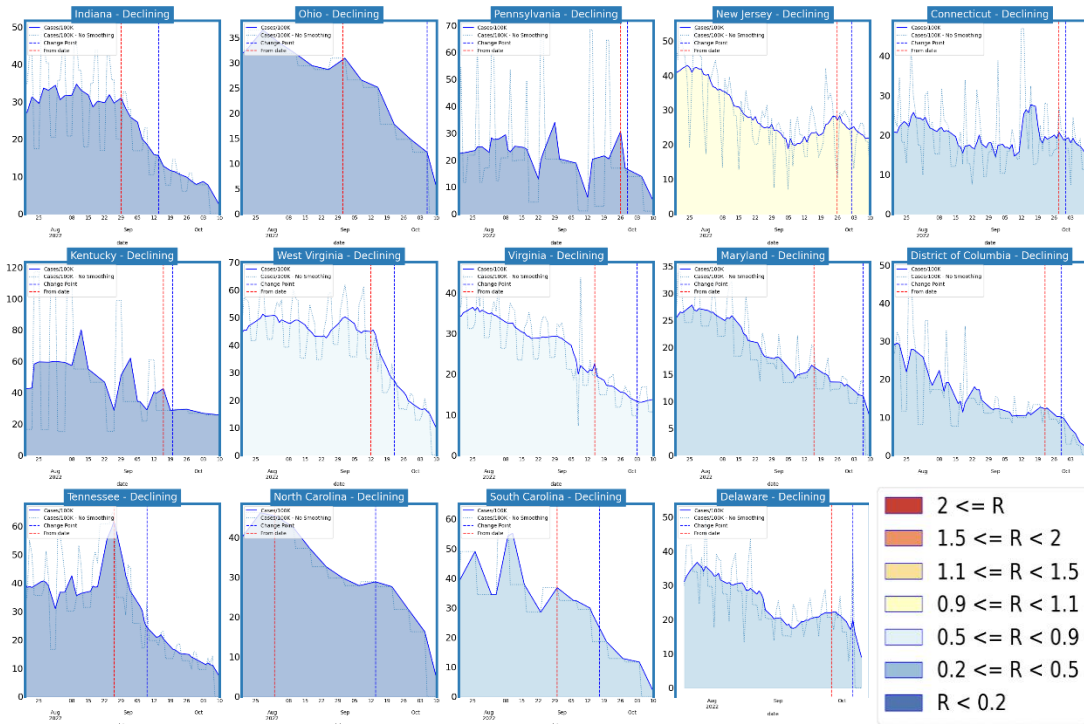
Status	Number of States	
	Current Week	Last Week
Declining	53	(48)
Plateau	1	(5)
Slow Growth	0	(1)
In Surge	0	(0)



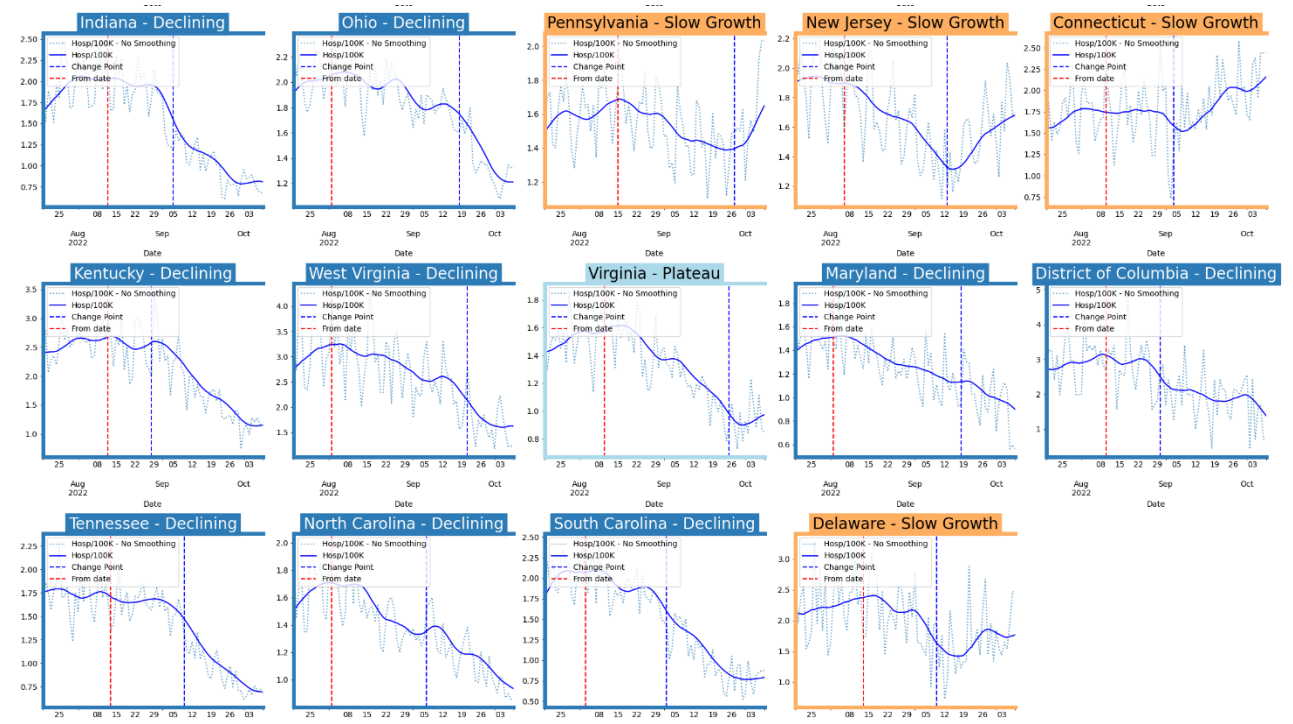
Status	Number of States	
	Current Week	Last Week
Declining	24	(27)
Plateau	19	(17)
Slow Growth	9	(9)
In Surge	1	(0)

Virginia and Her Neighbors

Cases



Hospitalizations

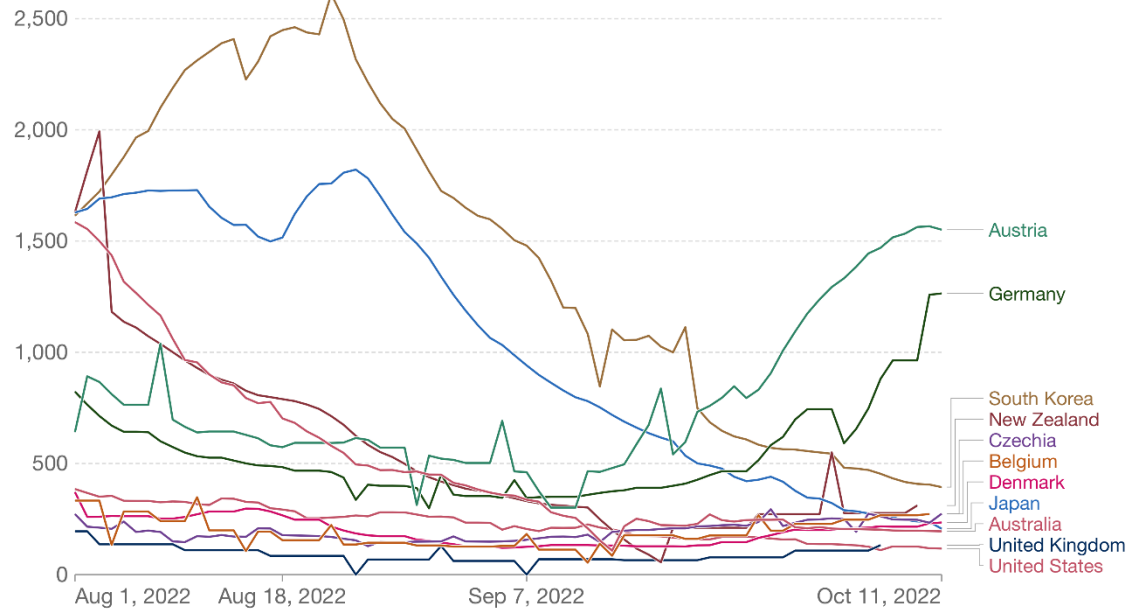


Around the World – Various trajectories

Confirmed cases

Daily new confirmed COVID-19 cases per million people

7-day rolling average. Due to limited testing, the number of confirmed cases is lower than the true number of infections.



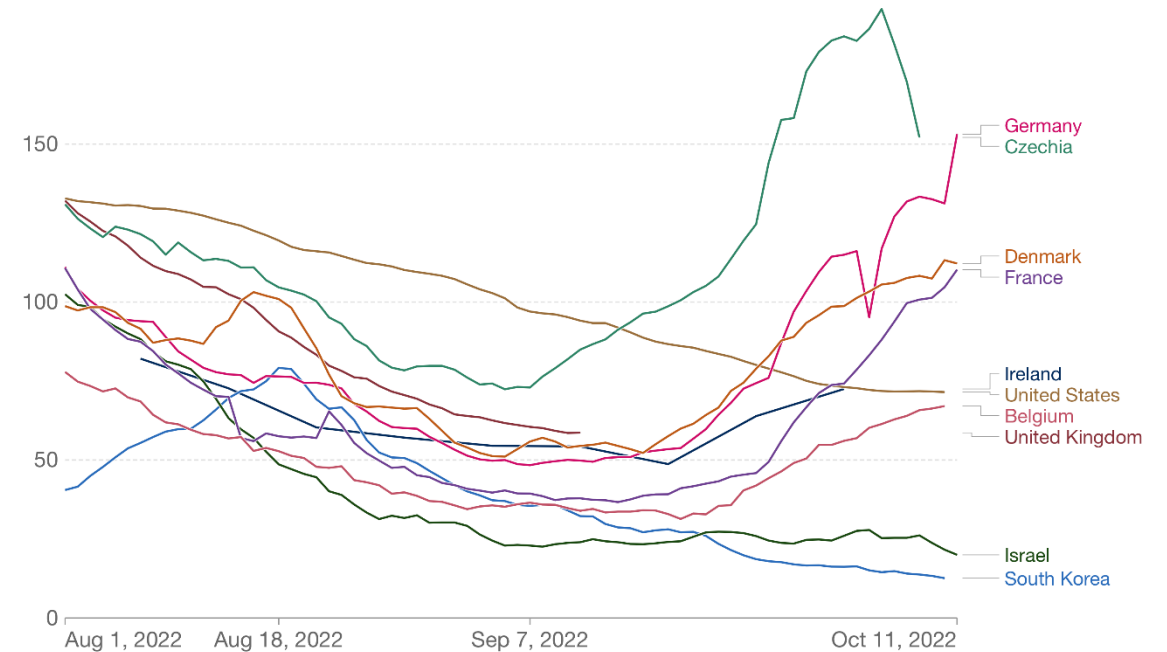
Source: Johns Hopkins University CSSE COVID-19 Data

CC BY

Hospitalizations

Weekly new hospital admissions for COVID-19 per million people

Weekly admissions refer to the cumulative number of new admissions over the previous week.



Source: Official data collated by Our World in Data

CC BY

Using Ensemble Model to Guide Projections

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

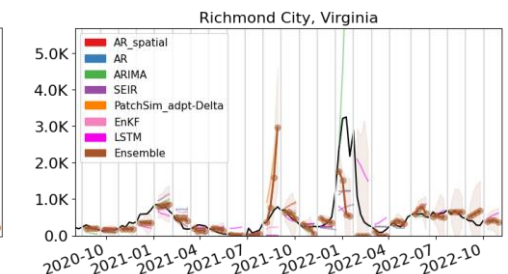
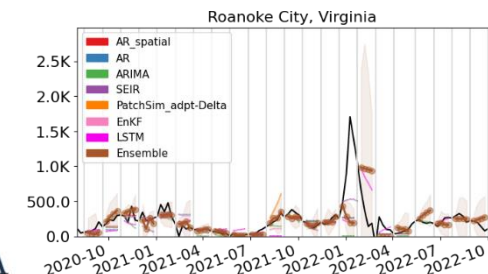
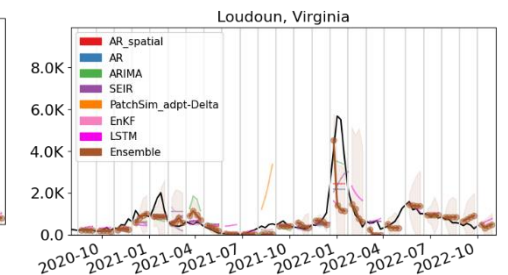
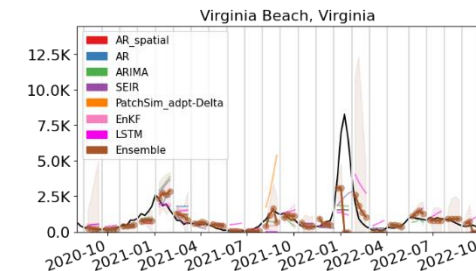
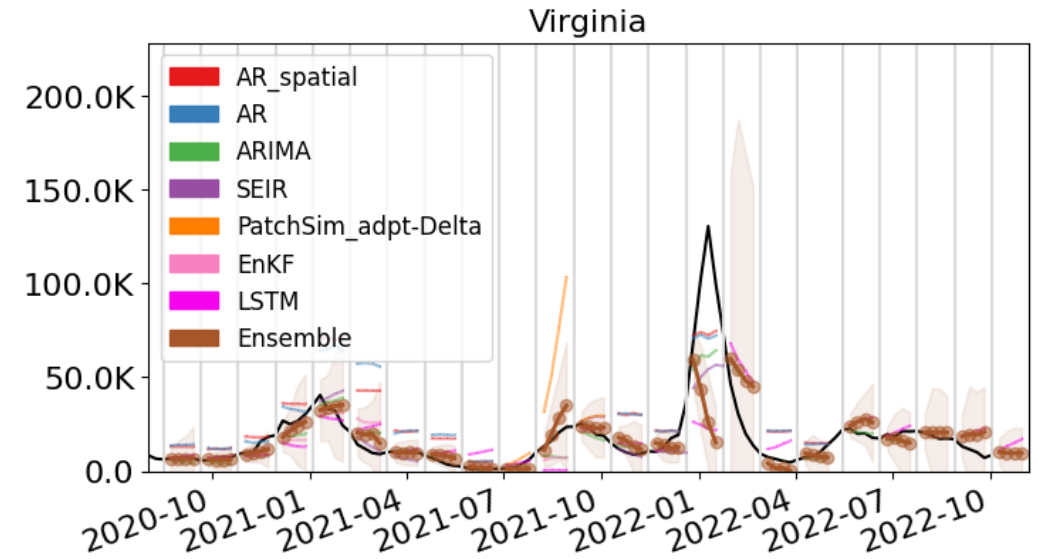
- Autoregressive (AR, ARIMA)
- Neural networks (LSTM)
- Kalman filtering (EnKF)

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.

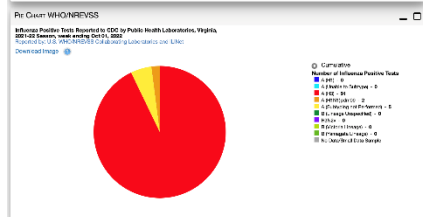
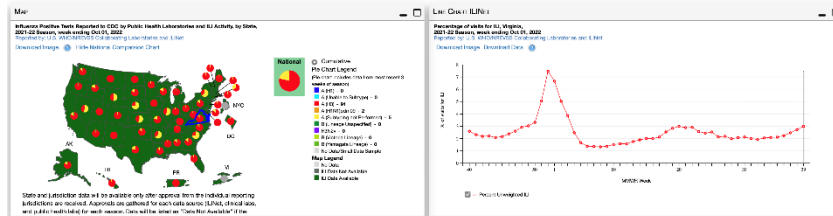
Also submitted to CDC Forecast Hub.



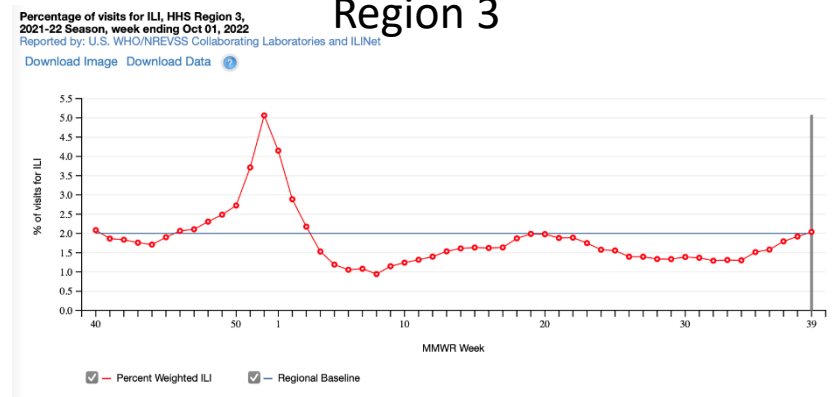
Current Influenza Hospitalization Forecast

Statistical models for submitting to CDC FluSight forecasting challenge

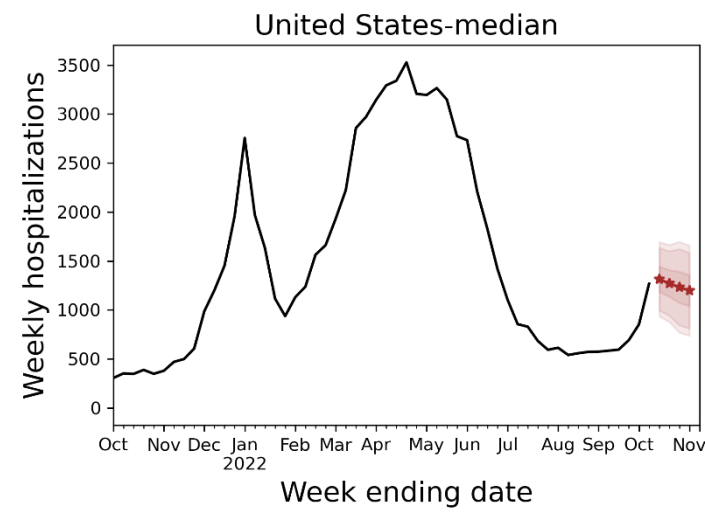
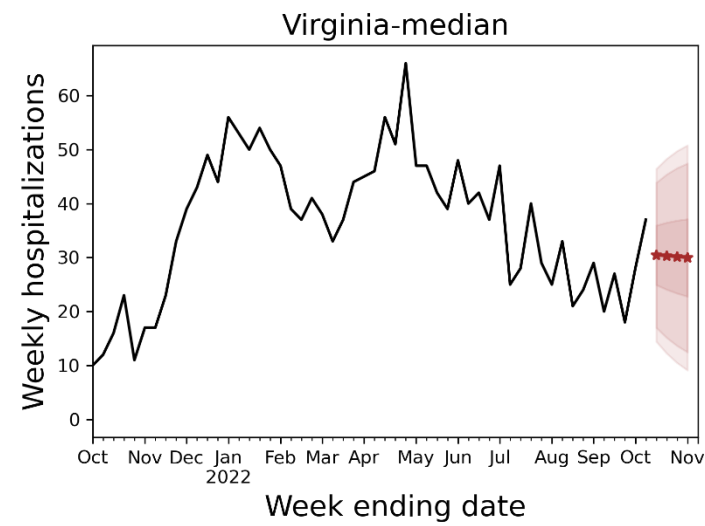
- Similar to COVID-19 case forecasts, uses a variety of statistical and ML approaches to forecast weekly hospital admissions for the next 4 weeks for all states in the US



Influenza A activity increasing in VA
Labs show high levels of H3 this season
(Influenza A H3N2 is more severe)



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



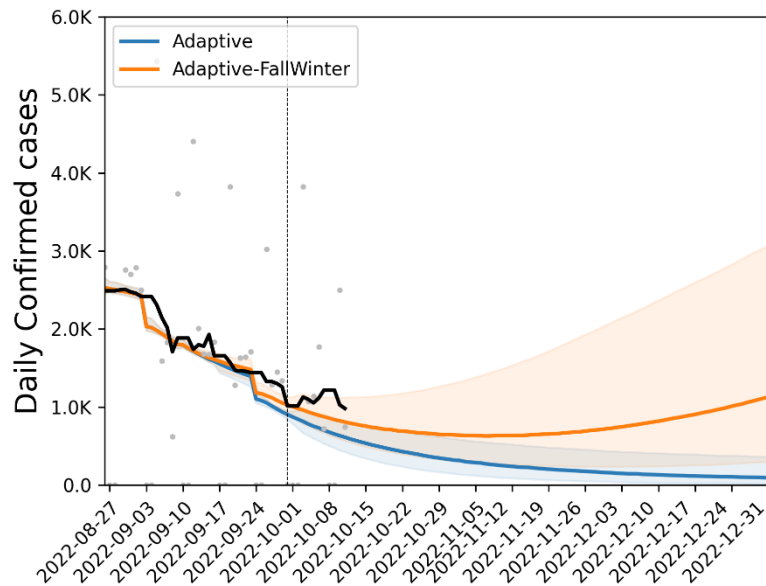
Initial forecasts have wide uncertainty due to noisiness in data due to low numbers of hospitalizations

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

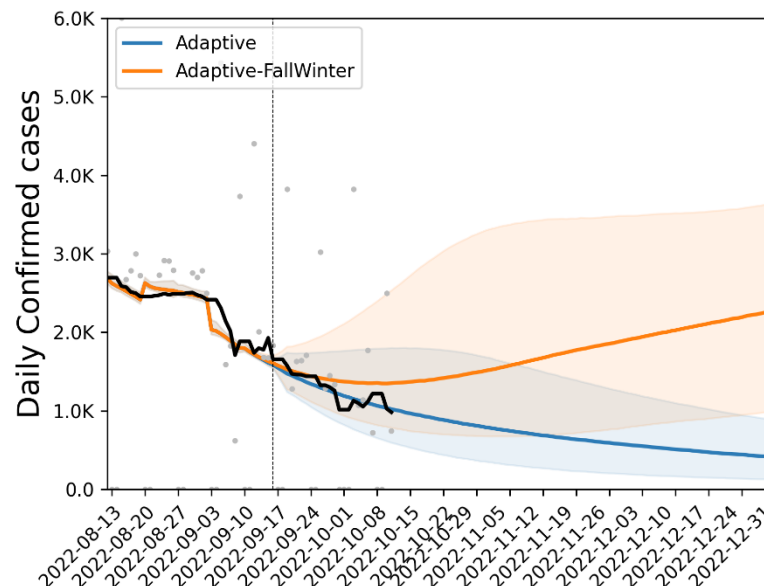
Projection from 1 weeks ago

Virginia Daily Confirmed - Comparison 2022-09-30



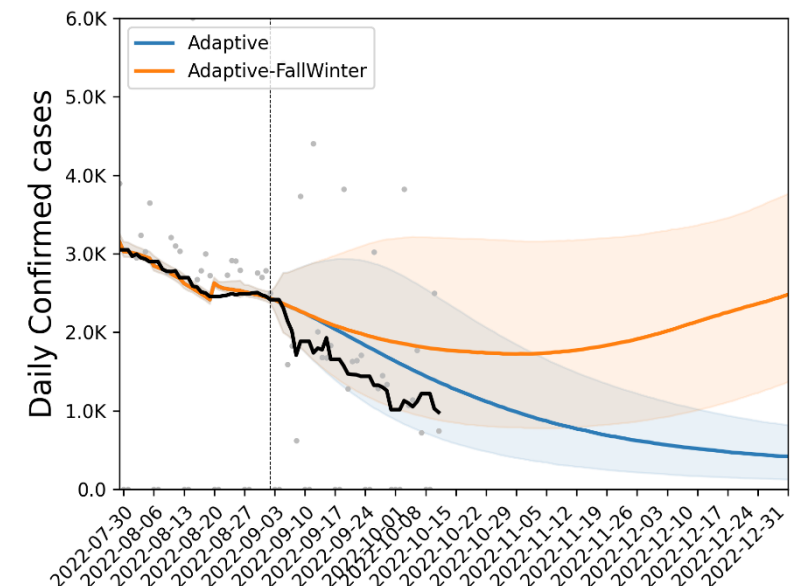
Projection from 3 weeks ago

Virginia Daily Confirmed - Comparison 2022-09-16



Projection from 5 weeks ago

Virginia Daily Confirmed - Comparison 2022-09-02

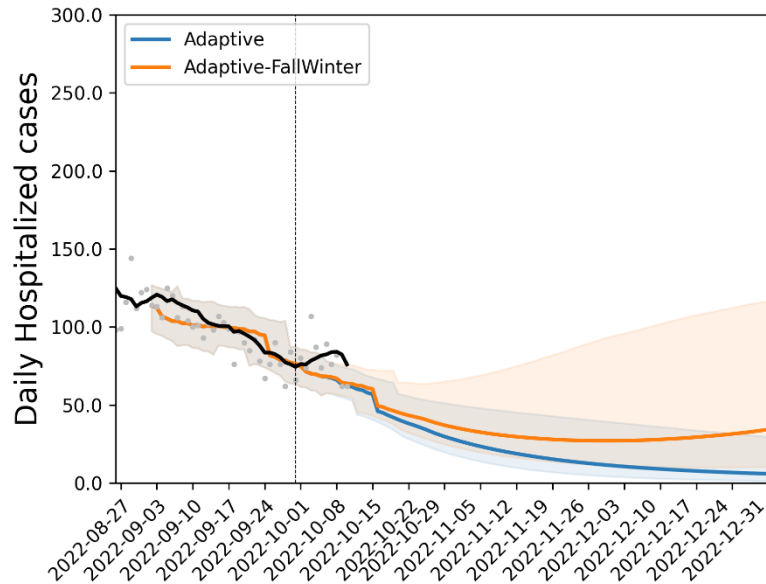


Previous projections comparison - Hospitalizations

- Previous projections have tracked observed hospitalizations well
- Projections track the decline well, however, recent rise in hospitalizations diverge a bit
- Hospitalization increase is not seen in cases as well

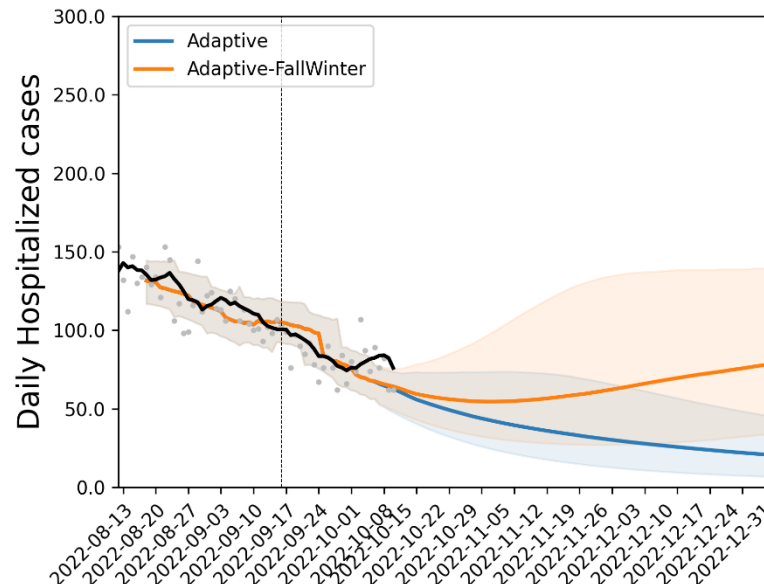
Projection from 1 weeks ago

Virginia Daily Hospitalized - Comparison 2022-09-30



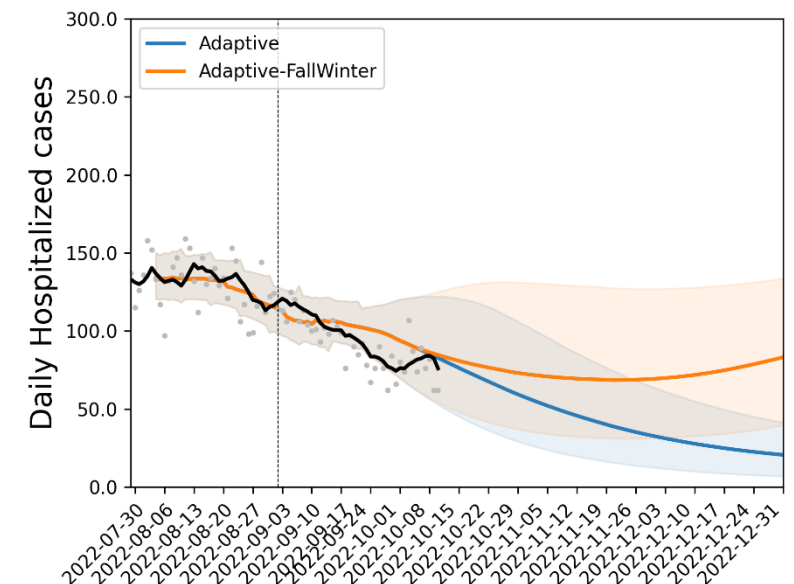
Projection from 3 weeks ago

Virginia Daily Hospitalized - Comparison 2022-09-16



Projection from 5 weeks ago

Virginia Daily Hospitalized - Comparison 2022-09-02



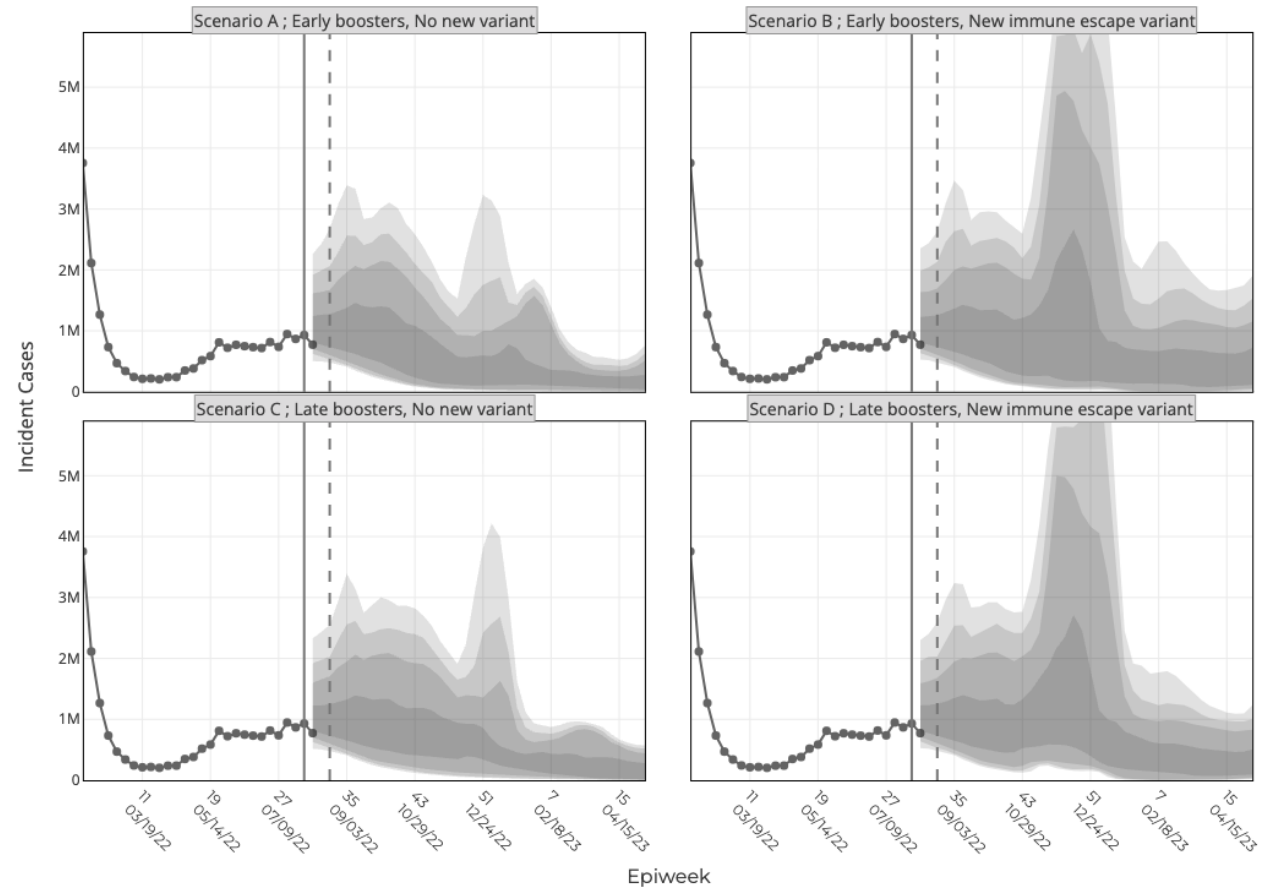
Scenario Modeling Hub – COVID-19 (Round 15)

Collaboration of multiple academic teams to provide national and state-by-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)



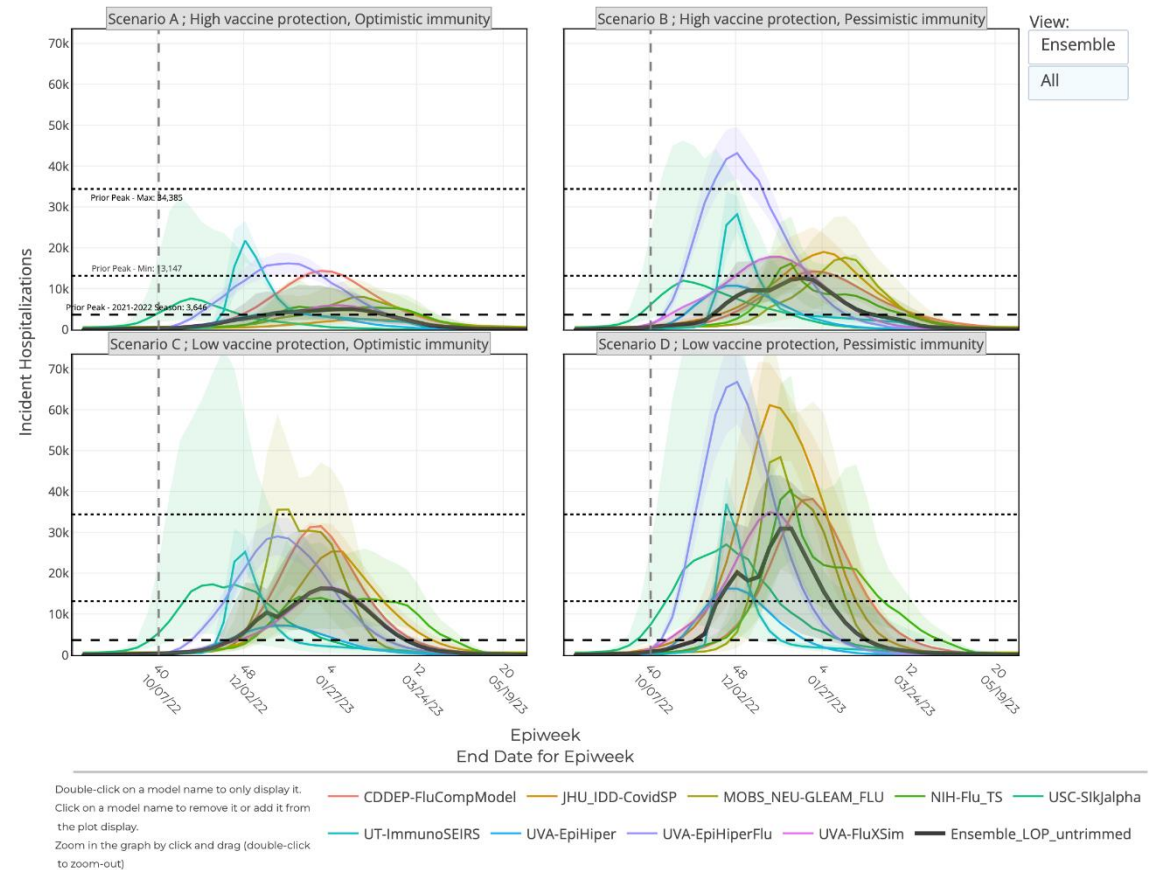
Scenario Modeling Hub – Influenza (Round 1)

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

- Round 1 results recently published
 - Impact of missed flu seasons on pre-season 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>

Projected Incident Hospitalizations by Epidemiological Week and by Scenario for Round 1 - US
(- Projection Epiweek; -- Current Week)



Key Takeaways

Projecting future cases precisely is impossible and unnecessary.

Even without perfect projections, we can confidently draw conclusions:

- **Case rates have remained flat as have hospitalizations**
- VA weekly case rate flat at 206 per 100K from 203 per 100K
 - US weekly case rate is down to 174 per 100K from 189 per 100K
 - VA hospital occupancy (rolling 7 day mean of 791 slightly down from 798 a week ago) currently on month plateau
- Sub-variant prevalence evolution as expected
- Projections from last week remain largely on target, though cases are diverging

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

Additional Analyses

References

Venkatramanan, S., et al. "Optimizing spatial allocation of seasonal influenza vaccine under temporal constraints." *PLoS Computational Biology* 15.9 (2019): e1007111.

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

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

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

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

Questions?

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