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

Foresight and Analysis of Infectious Disease Threats to Virginia's Public Health

October 12th, 2023

(data current to Sept 30th – Oct 10th) Biocomplexity Institute Technical report: TR BI-2023-247

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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 current and emerging Infectious Disease threats to the Commonwealth of Virginia using modeling and analytics

• Approach:

- Provide analyses and summaries of current infectious disease threats
- Survey existing forecasts and trends in these threats
- Analyze and summarize the current situation and trends of these threats in the broader context of the US and world
- Provide broad overview of other emerging threats





COVID-19 Activity levels continue to decline

- Declines in cases and hospitalizations have continued
- Other indicators continue to point towards continued declines or suggest no major change
- Wastewater based indicators similar mix of viral loads as in previous weeks

Genomic Surveillance maintains high diversity with no dominating variant

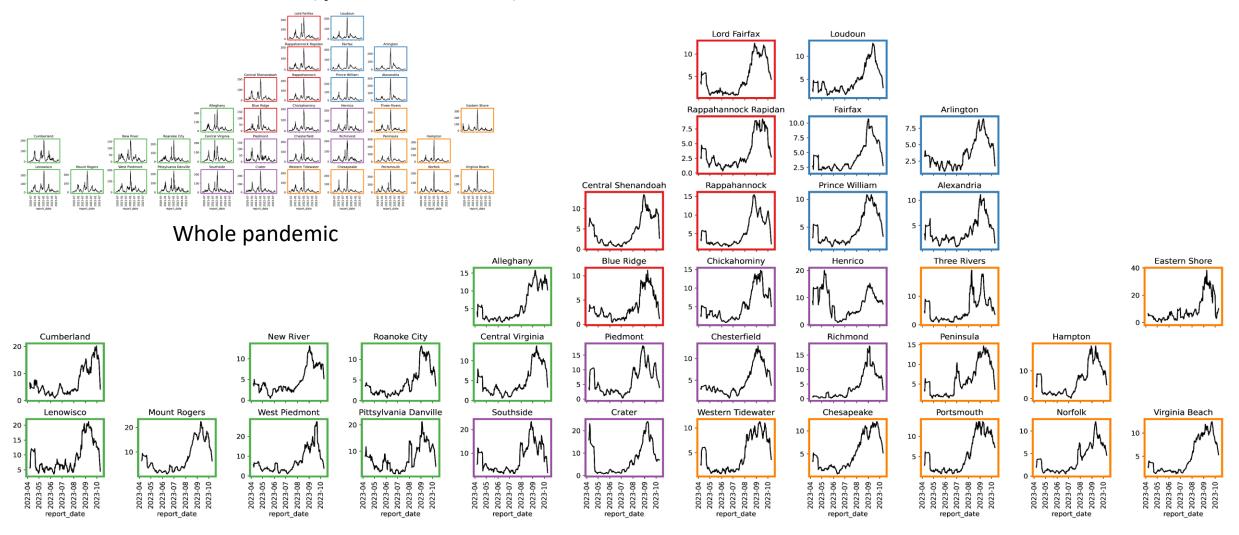
Together this suggests continued declines or easing into a plateau in near term



COVID-19 Surveillance



Case Rates (per 100k)





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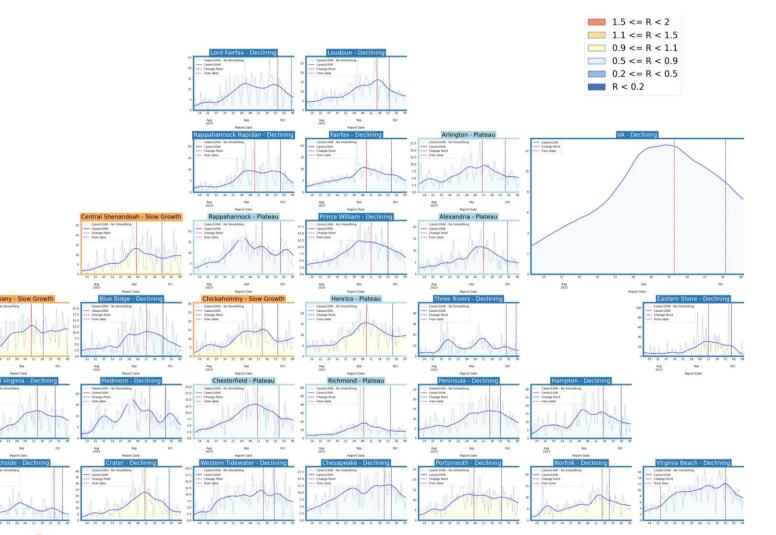
District Case Trajectories – last 10 weeks

nia Danville - Plate

<u>Rt estimates from EpiNow2</u>

Statuc	Number of Districts				
Status	Current Week	Last month			
Declining	24	(21)			
Plateau	8	(7)			
Slow Growth	3	(5)			
In Surge	0	(2)			

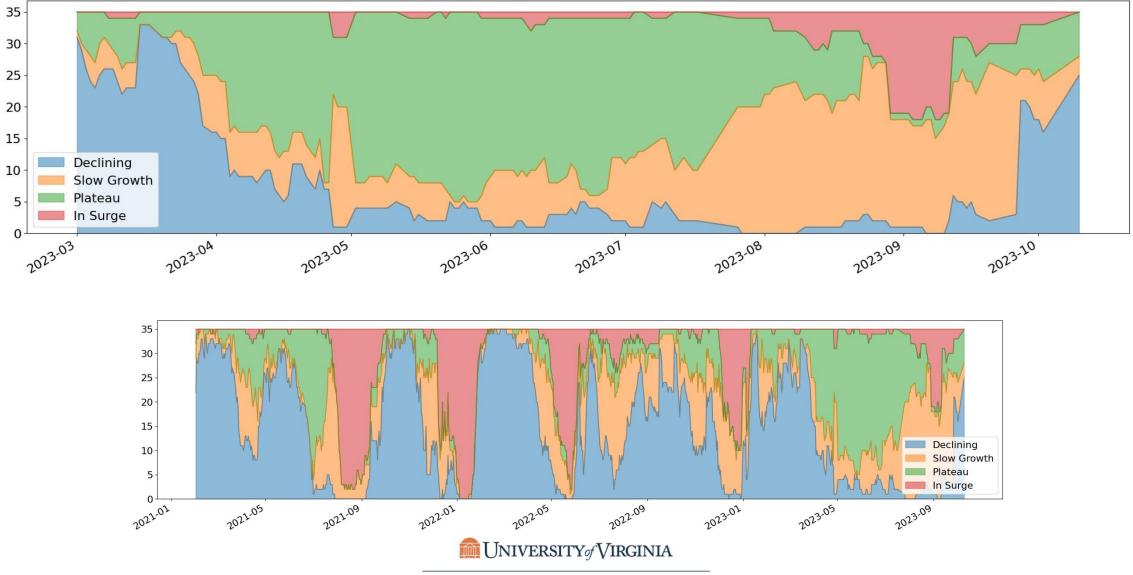
Curve shows smoothed case rate (per 100K) Trajectories of states in label & chart box Case Rate curve colored by Reproductive number



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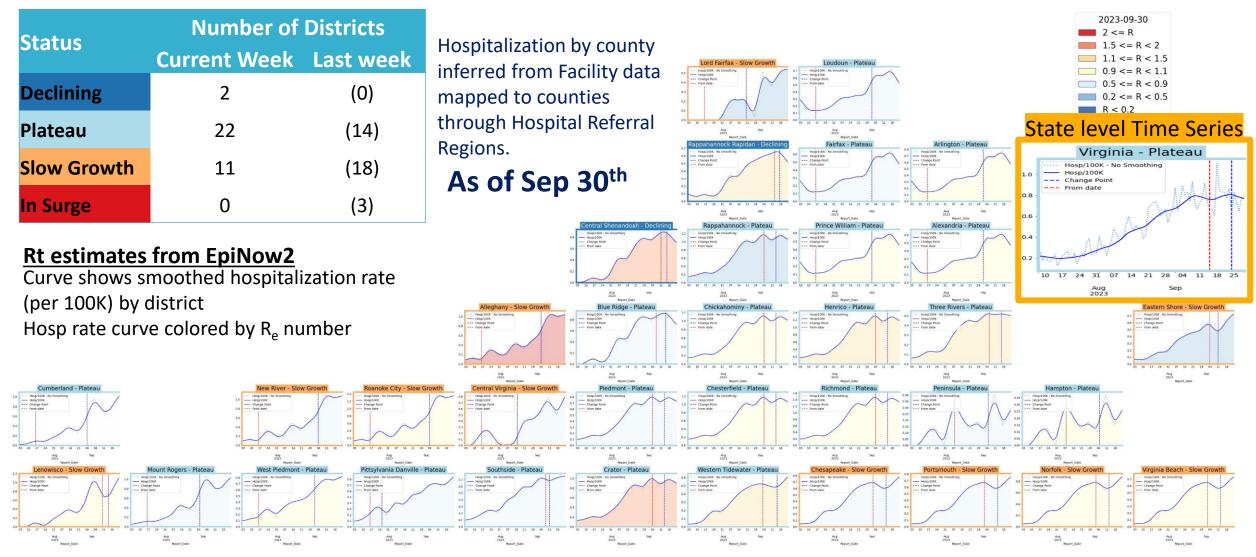
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District Case Trajectories – Recent 6 months



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District Hospital Trajectories – last 10 weeks



COVID-19 Growth Metrics



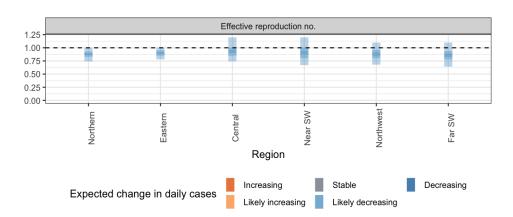
Estimating Daily Reproductive Number – VDH report dates – EpiNow2 estimation Re from VDH Cases (last 6 months)

Reproductive Estimate Summary as of October 10, 2023

Region	Reproductive number estimate	IQR	Trend forecast
State-wide cases	0.89	0.71 - 1.0	Likely decreasing
State-wide hosp	0.97	0.88 - 1.1	Likely decreasing
Central	0.93	0.74 - 1.2	Likely decreasing
Eastern	0.90	0.78 - 1.0	Likely decreasing
Far SW	0.86	0.64 - 1.1	Likely decreasing
Near SW	0.90	0.67 - 1.2	Likely decreasing
Northern	0.87	0.74 - 1.0	Likely decreasing
Northwest	0.88	0.68 - 1.1	Likely decreasing

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Estimate



Estimate based on partial data

Forecast

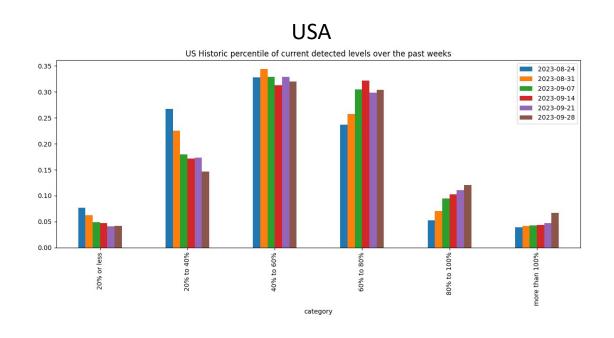
Methodology

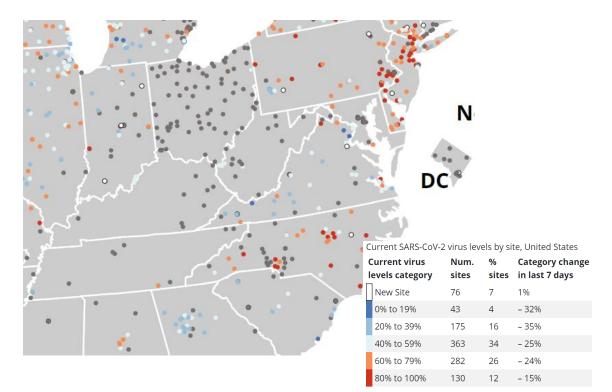
- Sam Abbott, Joel Hellewell, Katharine Sherratt, Katelyn Gostic, Joe Hickson, Hamada S. Badr, Michael DeWitt, Robin Thompson, EpiForecasts, Sebastian Funk (2020). **EpiNow2**: Estimate Real-Time Case Counts and Time-Varying Epidemiological Parameters. doi:10.5281/zenodo.3957489.
- Serial interval, generation time, and incubation period built into disease model via EpiNow2.
- Uses confirmation date but report date biases are better accounted for.
- Note: most recent data point for hospitalizations is 10 days prior to that of cases (HHS hospitalization through 9/30/23 vs. VDH case data through 10/10/23).

Wastewater Monitoring

Wastewater provides a coarse estimate of COVID-19 levels in communities

• Nationwide and in VA, sites have shifted from lower trend categories to higher trend categories





Total sites with current data: 1069

Total number of wastewater sampling sites: 1726

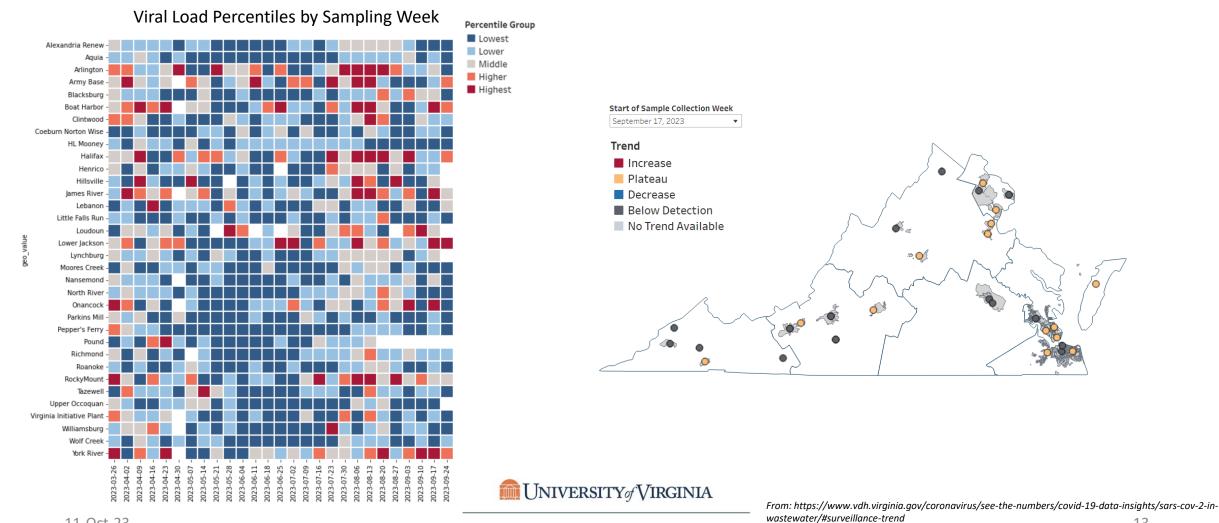
Data Source: CDC Data Tracker



Wastewater Monitoring – VA Sites

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

Some VA sites (esp. Eastern) are starting to shift to higher quintiles in wastewater percentile groups •



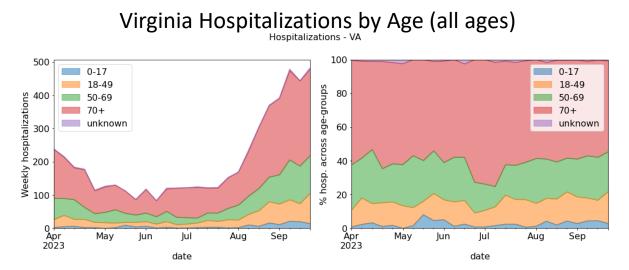
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Hospitalizations in VA by Age

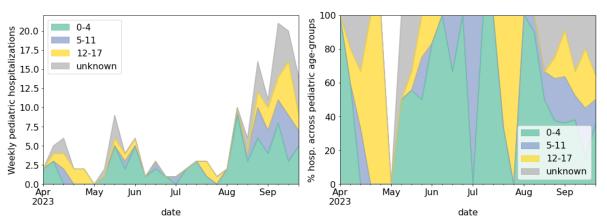
Age distribution in hospitals showing slight shift towards younger age groups

- Overall hospitalizations increasing across all age groups
- Increase in pediatric hospitalizations (0-4 age group), remain high, but have come down a little in last week





Pediatric hospitalizations - VA

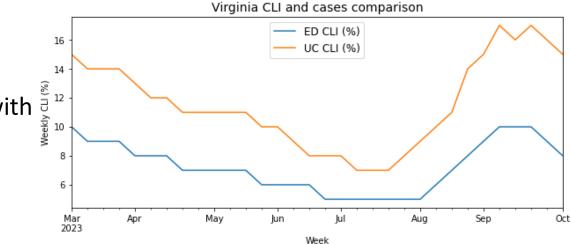


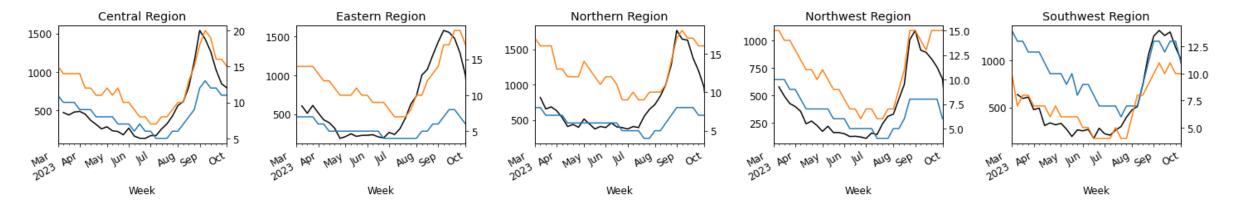
Note: These data are lagged and based on HHS hospital reporting

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 may be influenced by testing for other URIs
- CLI continues to decline
- Levels now back to those seen last in late winter

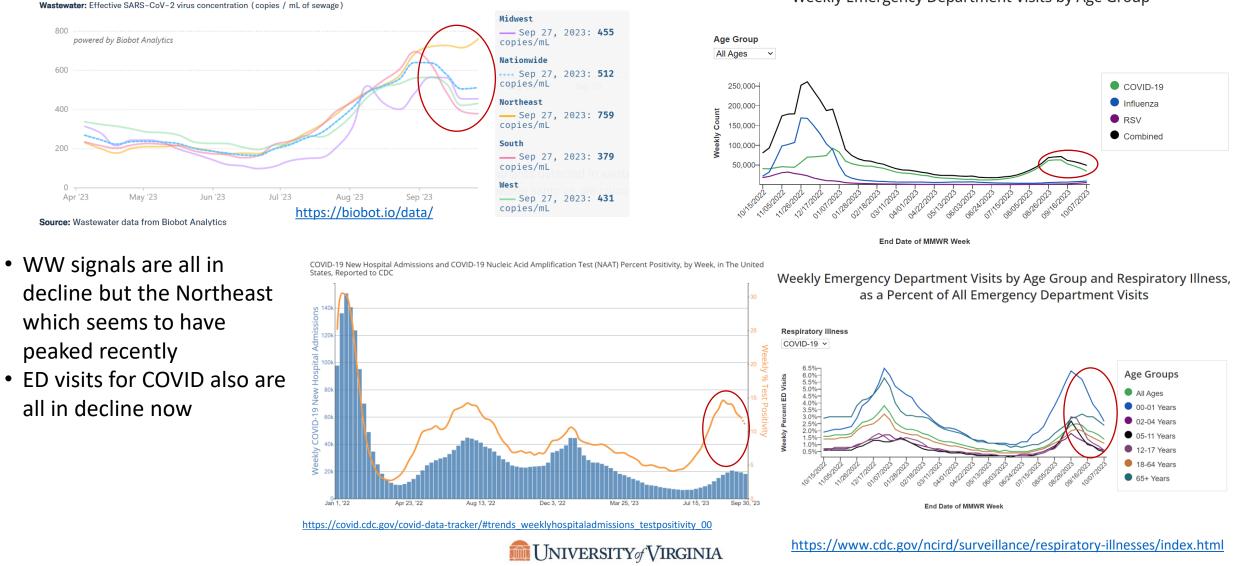






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Wastewater, ED visits, and Test positivity



Weekly Emergency Department Visits by Age Group

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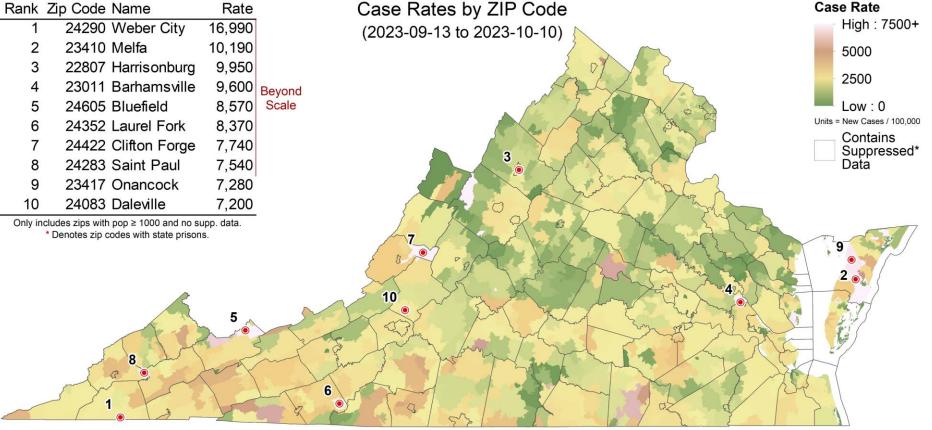
COVID-19 Spatial Epidemiology



ZIP Code level case rate since last meeting

New cases per 100k in the last four weeks

- Statewide COVID-19 case rates have declined since the last meeting.
- Divide rates by four to calculate average weekly incidence.
- Rates remain elevated in the Southwest and near Richmond. Areas north of Richmond have seen significant declines.
- There are no prisons in this top 10. But two ZIPs from the Eastern Shore appear this week.



Based on Spatial Empirical Bayes smoothed case rates, with an 8:1 ascertainment ratio, for five weeks ending 2023-10-10.

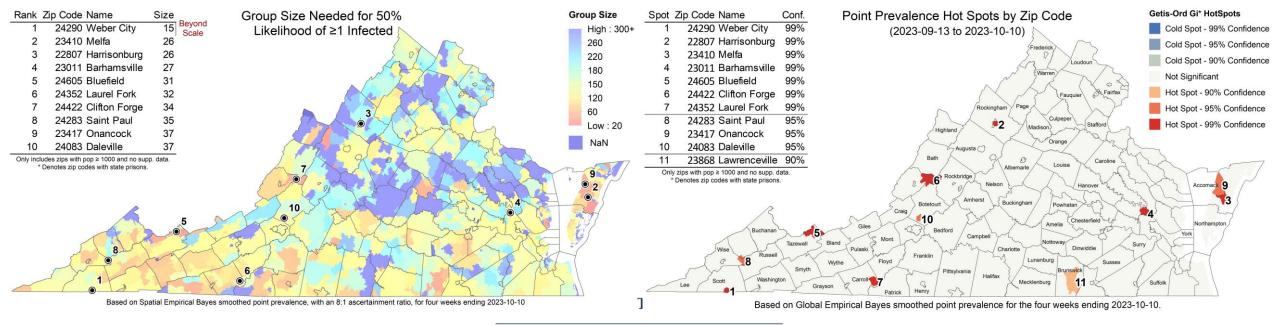
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Risk of Exposure and Hot Spots

Case rates since last meeting by zip code used to calculate risk of encountering someone infected in a gathering of randomly selected people and find spatial hot spots

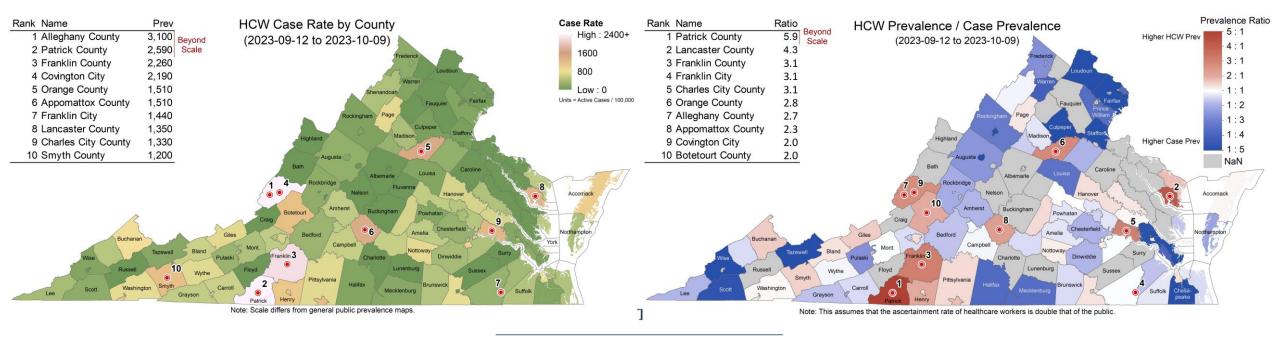
- **Group Size**: Assumes **8 undetected infections** per confirmed case (ascertainment rate from recent seroprevalence survey) and shows minimum size of a group with a 50% chance an individual is infected by zip code (e.g., in a group of 15 in Weber City, there is a 50% chance someone will be infected).
- **Spatial Clustering**: Getis-Ord Gi* based hot spots compare clusters of zip codes with four-week case rates higher than nearby zip codes to identify larger areas with statistically significant deviations.



COVID-19 among Healthcare Workers

COVID-19 case rates for the public and for healthcare workers (HCW) were compared to find regions where HCW suffered unusually high burdens of disease

- HCW Rate: Case rate among health care workers (HCW) over four weeks ending October 9
- HCW Ratio: Case rate among health care workers (HCW) over the same four weeks using patient facing health care workers as the numerator, and the population's case rate as the denominator.
- Unusual high rates and ratios are seen in the West Piedmont and Alleghany Health Districts.



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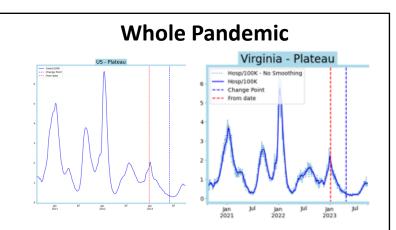
COVID-19 Broader Context



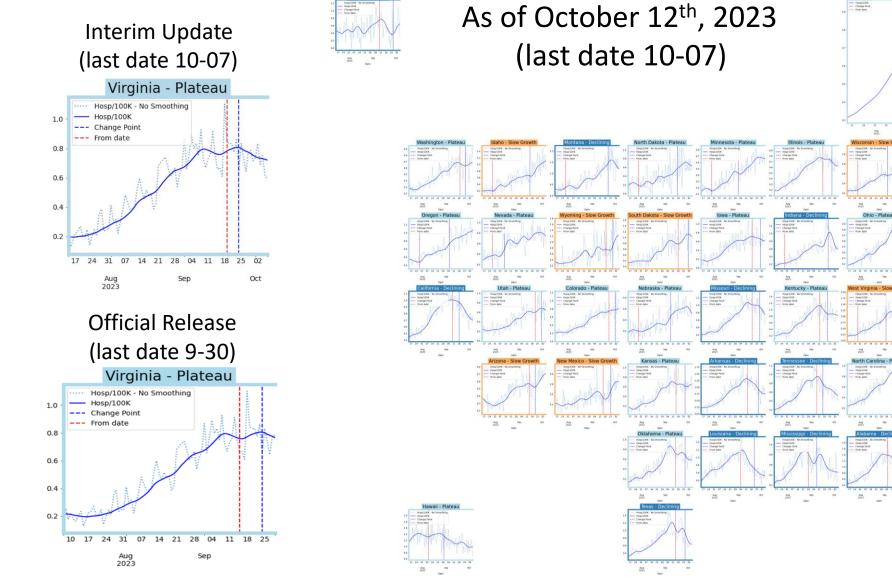
United States Hospitalizations



Number of States Status Current Last Week Month Declining 11 (4) Plateau 26 (23)**Slow Growth** (26) 16 (0) In Surge 0



United States Hospitalizations – Interim Update



11-Oct-23

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Aug Seg 1023

- Aug 2023

Virginia - Platea

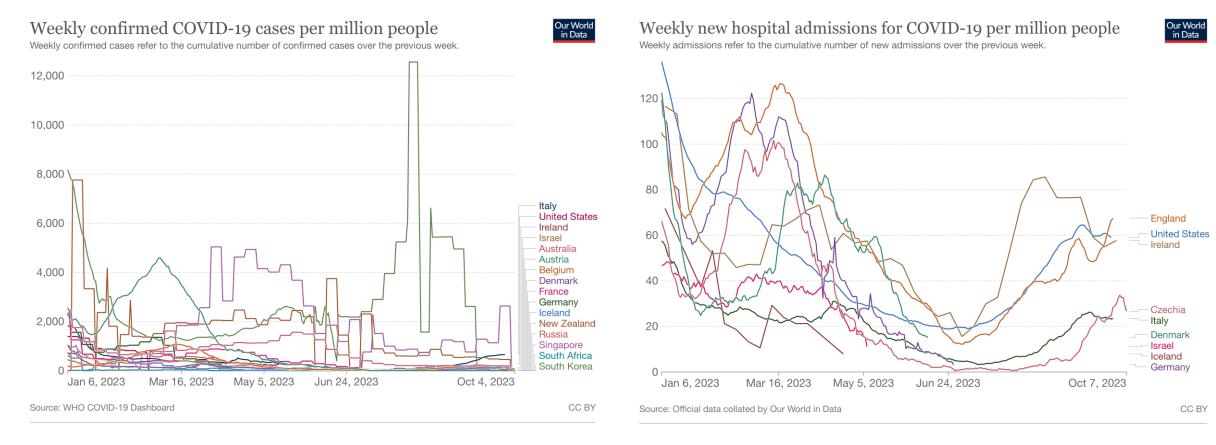
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Around the World – Various trajectories

Confirmed cases





11-Oct-23



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Hospitalizations

COVID-19 Genomic Update



SARS-CoV2 Variants of Concern

Emerging variants have potential to continue to alter the future trajectories of pandemic and have implications for future control

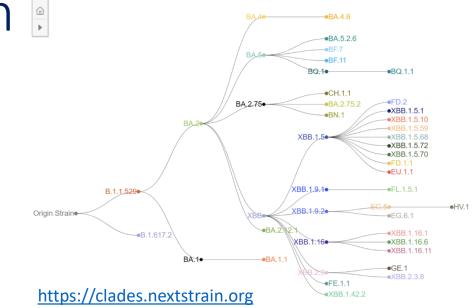
• Variants have been observed to: increase transmissibility, increase severity (more hospitalizations and/or deaths), and limit immunity provided by prior infection and vaccinations

Weighted Estimates in HHS Region 3 for 2-Week Periods in 6/11/2023 – 9/30/2023

Nowcast Estimates in HHS Region 3 for 9/17/2023 – 9/30/2023

Description of the set of the set

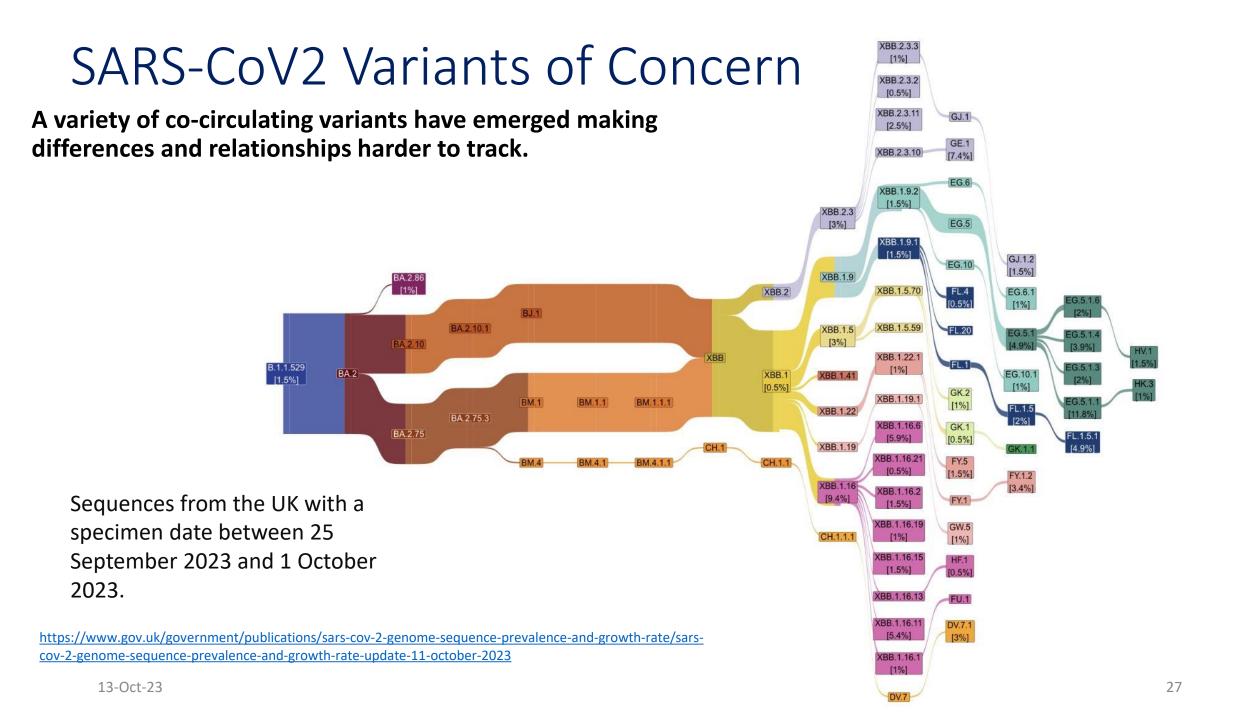
	ted Estimat icing results		proportions b	ased on rep	orted genon	nic	projected estimates of variant proportions		WHO label	Lineage #	%Total	95%PI	
									Omicron	EG.5	29.2%	26.0-32.6%	
										HV.1	15.4%	12.7-18.6%	
00% -										FL.1.5.1	14.3%	12.5-16.3%	
		EG.5	EG.5	EG.5	ŝ					XBB.1.16.6	9.2%	7.1-11.8%	
		B	ü	ů.	EG.5	9	LO,	10		XBB.2.3	5.2%	4.0-6.7%	<u> </u>
		-	_		ш	Ö	EG.5	EG.5		XBB.1.16	4.1%	3.4-4.9%	
30% -							u u	ш		XBB.1.5.70	2.8%	1.4-5.6%	
0.70										XBB.1.16.15	2.8%	1.7-4.4%	
										XBB.1.16.1	2.7%	2.1-3.4%	
		LO,								XBB.1.16.11	2.6%	2.0-3.4%	
	XBB.1.5	XBB.1.5	XBB.1.5							XBB	1.9%	1.3-2.8%	
0% -	2	8	E C							GE.1	1.6%	0.8-3.1%	
		×	ē							EG.6.1	1.1%	0.7-1.8%	
			~				5			XBB.1.9.1	1.0%	0.8-1.2%	
							HV.1	-		HE.1	1.0%	0.6-1.7%	
								HZ.		XBB.1.5	0.9%	0.7-1.1%	
0%-										GK.2	0.8%	0.4-1.8%	
			16							XBB.1.5.72	0.8%	0.5-1.3%	
										XBB.1.9.2	0.5%	0.4-0.8%	
			XBB.1.16							XBB.1.5.68	0.3%	0.2-0.6%	
0%-			×							XBB.1.5.10	0.3%	0.2-0.5%	
0 70										XBB.2.3.8	0.3%	0.2-0.5%	
										XBB.1.42.2	0.2%	0.1-0.5%	
										CH.1.1	0.2%	0.1-0.4%	
										BA.2	0.1%	0.0-0.5%	
0%										FD.1.1	0.1%	0.1-0.3%	
	53	23	53	53	53	53	53	53		FE.1.1	0.1%	0.0-0.2%	
	6/24/23	7/8/23	7122/23	8/5/23	8/19/23	9/2/23	9/16/23	9/30/23		XBB.1.5.59	0.1%	0.0-0.2%	
	79	2	22	æ	8/1	6	1/6	5/6		EU.1.1	0.0%	0.0-0.1%	
	-									XBB.1.5.1	0.0%	0.0-0.0%	
								Selected 2-Week		BQ.1	0.0%	0.0-0.0%	
								We		BA.5	0.0%	0.0-0.0%	
								Se 2-		FD.2	0.0%	0.0-0.0%	
			Co	lection date	two-week per	iod endina			Other	Other*	0.1%	0.1-0.2%	



Omicron Updates*

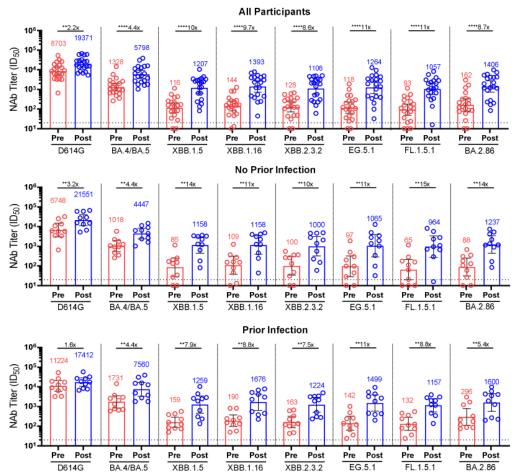
- HHS Region 3 estimates restored
- New lineage EG.5 (XBB.1.9*) up to 29.2 from 25.5%
- New lineage FL.1.5.1 (XBB.1.9*) up to 13.7 from 12%
- New lineage HV.1 (XBB.1.9*) up to 15.4%
- Most circulating variants are sublineages of XBB.1.9, XBB.1.16, XBB.1.5, and XBB.2.3





Updated Booster & Variants of Concern

Figure 2. Analysis of Neutralizing Antibody Titers Against Ancestral SARS-CoV-2 (D614G) and BA.4/BA.5, XBB.1.5, XBB.1.16, XBB.2.3.2, EG.5.1, FL.1.5.1 and BA.2.86 Variants in a Randomly-selected Subset of Participants Who Received Monovalent mRNA-1273.815



https://www.medrxiv.org/content/10.1101/2023.08.22.23293434v2

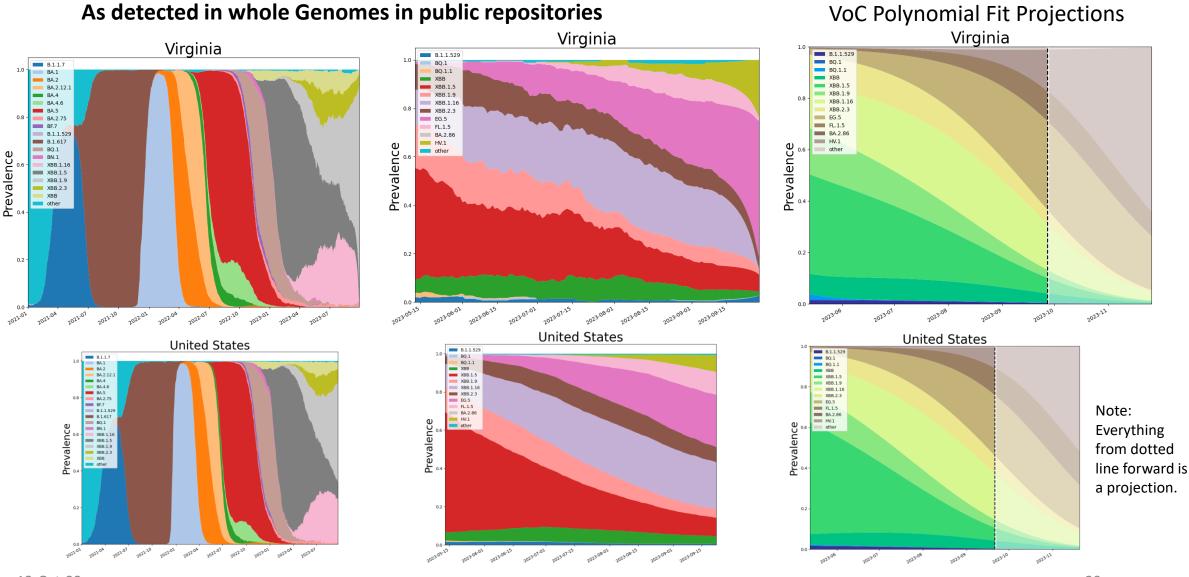
Phase 2/3 participants in updated booster trial show significantly increased neutralization levels for XBB.2.3.2, EG.5.1, FL.1.5.1 and BA.2.86.

The occurrence of solicited adverse reactions and unsolicited adverse events were overall similar to those previously reported for the original mRNA-1273 50-µg and omicron BA.4/BA.5-containing bivalent mRNA-1273 vaccines.

SARS-CoV2 Omicron Sub-Variants

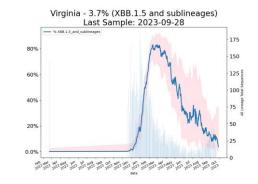
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covSPECTRUM



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SARS-CoV2 Omicron Sub-Variants

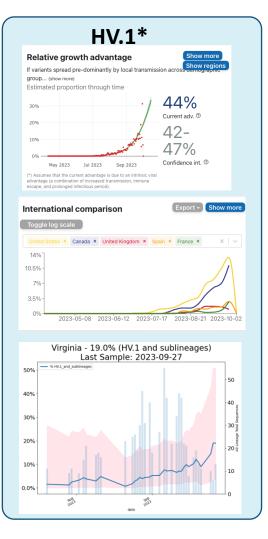


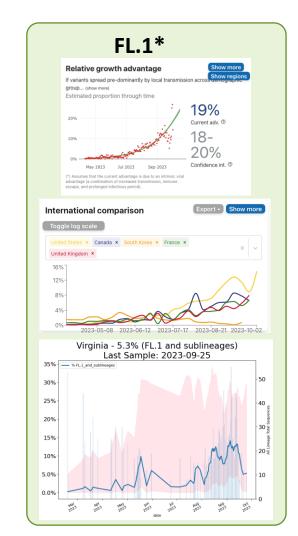
EG.5* Relative growth advantage If variants spread pre-dominantly by local transmissio group... (show more) Estimated proportion through time 60% 22% Current adv. @ 22-23% Confidence int. @ May 2023 Jul 2023 Sen 2023 (*) Assumes that the current advantage is due to an intrinsic viral advantage (a combination of increased transmission, immune escape, and prolonged infectious period). Export - Show more International comparison Toggle log scale United States × China × Canada × Japan × South Korea × 100% 75%-50%-25% 2023-05-08 2023-06-12 2023-07-17 2023-08-21 2023-10-02 Virginia - 66.7% (EG.5 and sublineages) Last Sample: 2023-09-27 - % EG.5 and sublinea 80% 60% 40% 20%-0.0% N013

<u>COV-spectrum</u> "Editor's choice" Variants to watch

Known variants Which variant would you like to explore? Editor's choice V 98.2% DV.7.1* XBB* 0.6% 0.3% HV.1* 11.3% BA.2.86* 2.9% FL.1.5.1* 11.1% HK 3* 10.9% EG.5.1* 39.9% XBB.1.16.6* XBB.2.3* 7.5% XBB.1.5* 8.2% S:F456L 70.6% S:F456L + S:L455F 10.9% 89.3% ORF9b:i5T 82.4% ORF8:G8* **covSPECTRUM**

Enabled by data from **GISAID**





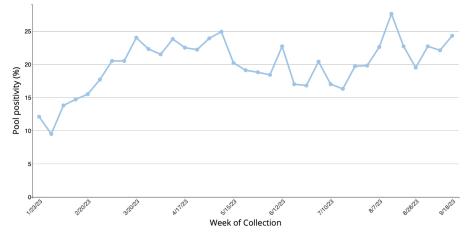
Global SARS-CoV2 Variant Status

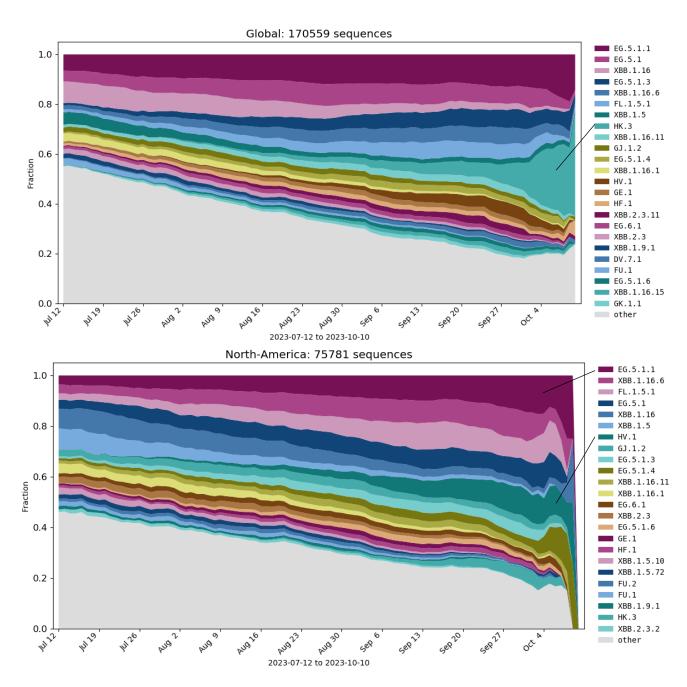
Traveller Surveillance

B.1.1.529 BA.2 BA.2.75 BN.1 CH.1.1 BA.5 BA.2.86 XBB XBB.1.5.10 BE 7 BQ.1.1 XBB.1.5 XBB.1.5.1 XBB.1.5.70 EU.1.1 FD.2 FL.1.5.1 XBB.1.5.59 XBB.1.5.72 XBB.1.9.1 XBB.1.16.1 XBB.1.16.11 XBB.1.9.2 EG.5 EG.6.1 XBB.1.16 XBB.1.16.6 Other FE.1.1 XBB.2.3 GE.1 Viral lineages among positive pools (%) 8 8 8 8 8 AIT723 518123 Week of Collection

Variants Detected, by Collection Week

Positivity Rate for Pooled Samples, by Collection Week





https://cov.lanl.gov/components/sequence/COV/sparks.comp https://covid.cdc.gov/covid-data-tracker/#traveler-genomic-surveillance

Influenza Update

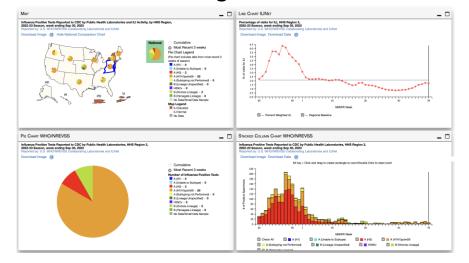


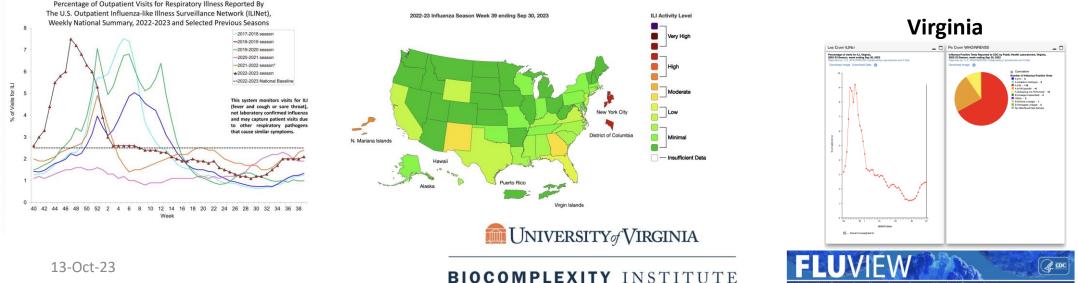
Current Influenza Situation – ILI Activity

Influenza Activity is below threshold

- Virginia is in "Low" level as some states and regions renew upward growth after a couple week pause.
- National ILI activity remains below threshold but has had ٠ some slight growth after 4 weeks of plateau
- Only Region 2 (NY, NJ, PR, VI) are over the seasonal threshold for ILI activity

Region 3





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National Modeling Hub Updates

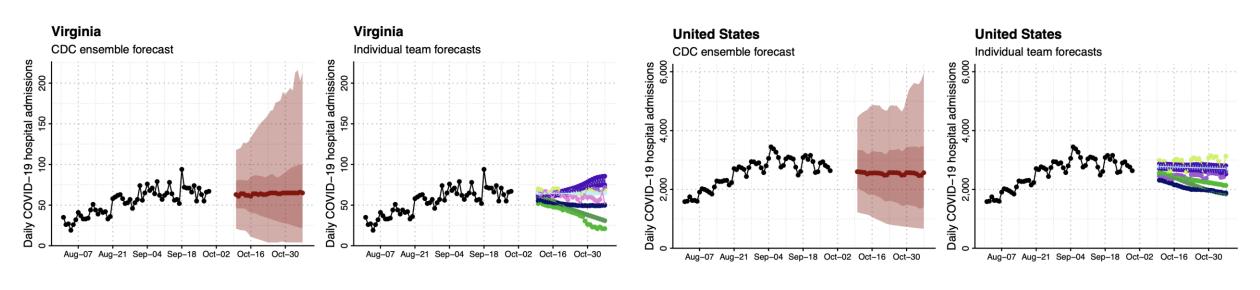


Current COVID-19 Hospitalization Forecast

Statistical models for submitting to CDC COVID Forecasting Hub

 Uses a variety of statistical and ML approaches to forecast weekly hospital admissions for the next 4 weeks for all states in the US

Hospital Admissions for COVID-19 and Forecast for next 4 weeks (CDC COVID Ensemble)



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Preliminary Influenza Hospitalization Forecast

Statistical models for submitting to CDC Influenza Forecasting Hub

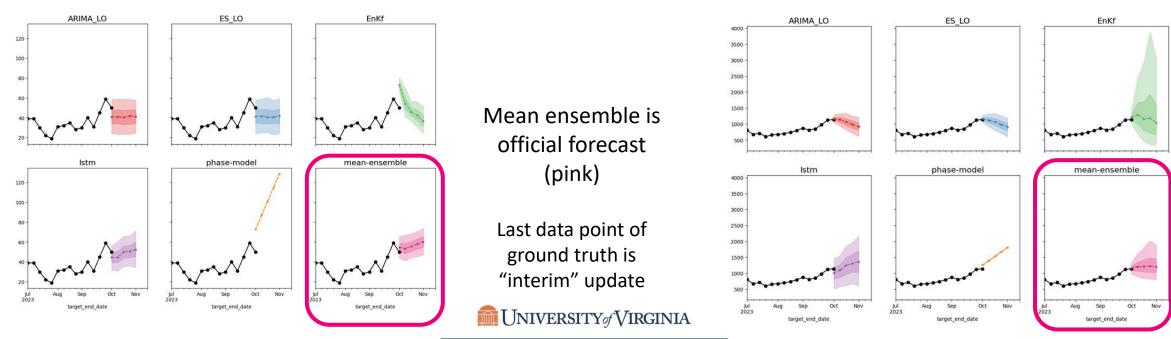
 Uses a variety of statistical and ML approaches to forecast weekly hospital admissions for the next 4 weeks for all states in the US

UVA models only

Hospital Admissions for Influenza and Forecast for next 4 weeks

United States

US || ref_date = 2023-10-14 || avl_date = 2023-10-07



Virginia

Virginia || ref date = 2023-10-14 || avl date = 2023-10-07

Scenario Modeling Hub – Influenza (Round 4)

Collaboration of multiple academic teams to provide national and state-by-state level projections for 6 aligned scenarios

- Round Designed to explore a season dominated by H3 vs. H1 with different levels of seasonal flu vaccination coverage
- Based on data till September 2nd, 2023

Scenario Dimensions:

Influenza type A/H3 vs. A/H1:

- H3 higher hospitalization rates with vax efficacy weaker in older groups
- H1 lower hospitalization rates and efficacy even across age groups

Vaccination levels (compared to 2021-22 season):

Low (20% less) vs. Business as Usual (same) vs. Higher (20% more)

https://fluscenariomodelinghub.org/viz.html

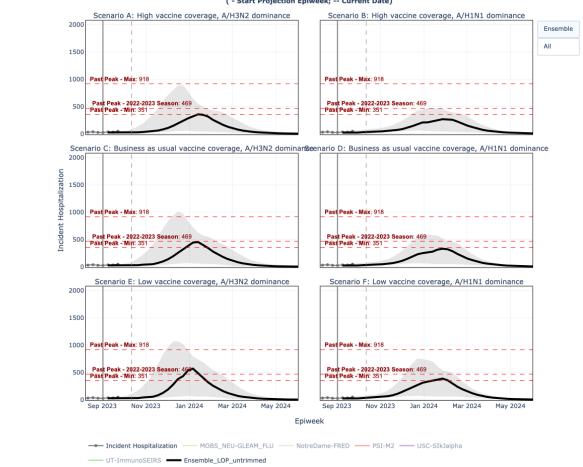
	Season dominated by influenza A/H3N2, indexed on age distribution of 2017-18 season. VE = 40% against medically attended illnesses and hospitalizations, VE drops in older age groups	Season dominated by influenza A/H1N1, indexed on age distribution of 2019-20 season. VE = 40% against medically attended illnesses and hospitalizations, similar VE across all age groups
 Higher than Usual Vaccine Coverage Vaccine coverage is 20% higher than in the 2021-22 flu season in all age groups and jurisdictions. (20% is a relative change, ie a 50% coverage for age group <i>a</i> and jurisdiction <i>j</i> in 2021-22 translates to a 50%*1.20=60% coverage for 2023-24). Overall, the US coverage is about 60% in this scenario. 	Scenario A	Scenario B
 Business as Usual Vaccine Coverage Vaccine coverage is the same as in the 2021-22 flu season in all age groups and jurisdictions. Overall, the US coverage is about 50% in this scenario. 	Scenario C	Scenario D
 Low Vaccine Coverage Vaccine coverage is 20% lower than in the 2021-22 flu season in all age groups and jurisdictions. Overall, the US coverage is about 40% in this scenario. 	Scenario E	Scenario F

Scenario Modeling Hub – Influenza (Round 4)

• Severity of season has a strong influence.

Virginia Hospitalizations

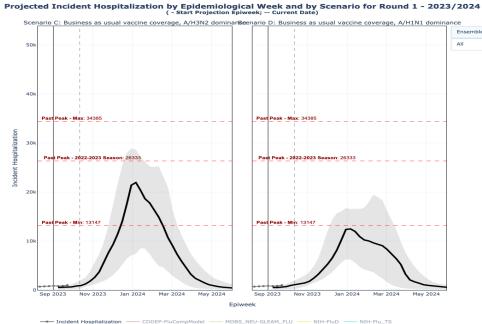
Projected Incident Hospitalization by Epidemiological Week and by Scenario for Round 1 - 2023/2024 (- Start Projection Epiweek; -- Current Date)



H3N2 dominated season (more severe)

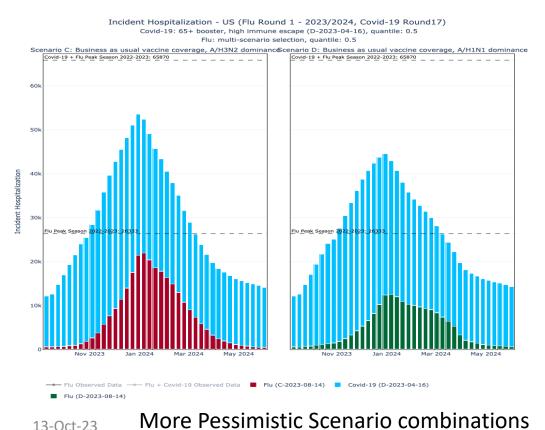
H1N1 dominated season (less severe)

United States Hospitalizations

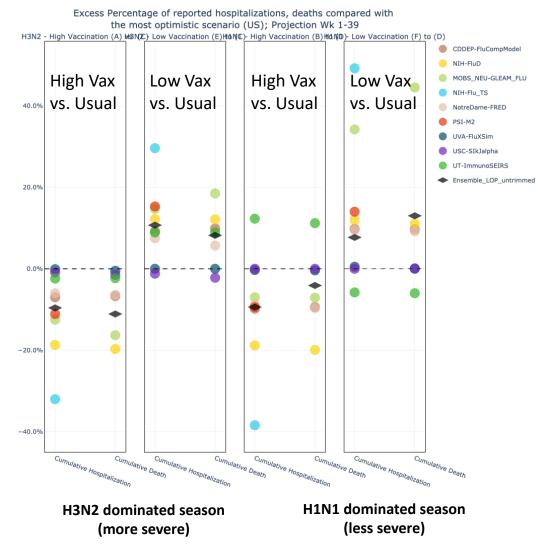


Scenario Modeling Hub – Influenza (Round 4)

- Higher coverage can reduce hospitalizations significantly, more so in an H3N2 season
- Projected COVID and Influenza hospitalizations could exceed 50k with serious influenza season aligned with high immune escape COVID



Virginia Hospitalizations



Scenario Modeling Hub – COVID-19 (Round 17)

Collaboration of multiple academic teams to provide national and state-by-state level projections for 6 aligned scenarios

- Preliminary Results
- Round Designed to explore different seasonal vaccination levels and the impact of Immune Escape

Scenario Dimensions:

Immune Escape (IE):

Slower IE (20%/yr) vs. Faster IE (50%/yr)

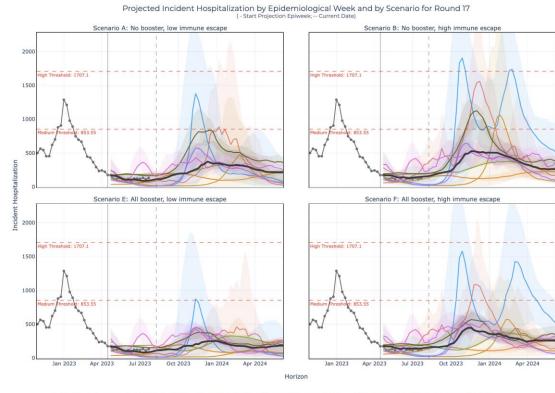
Vaccination levels:

None vs. Vulnerable and 65 + vs. Broader population of eligible https://covid19scenariomodelinghub.org/viz.html

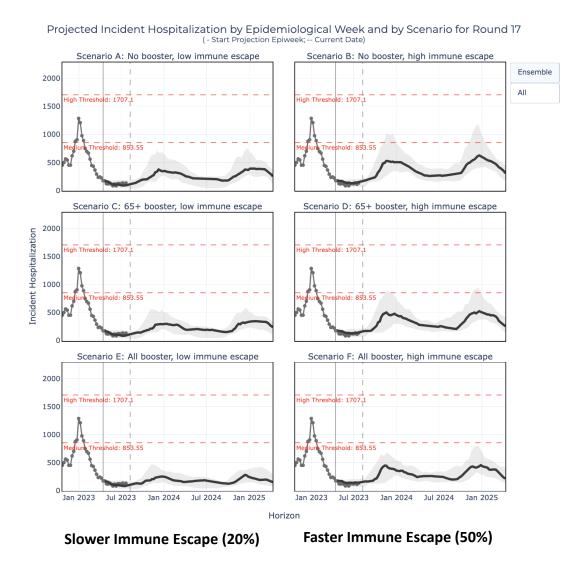
	 Low immune escape Immune escape occurs at a constant rate of 20% per year 	 High immune escape Immune escape occurs at a constant rate of 50% per year
 No vaccine recommendation Uptake negligible or continues at very slow levels based on existing 2022 booster trends 	Scenario A	Scenario B
 Reformulated annual vaccination recommended for 65+ and immunocompromised Reformulated vaccine has 65% VE against variants circulating on June 15 Vaccine becomes available September 1 Uptake in 65+ same as first booster dose recommended in September 2021 Uptake in individuals under 65 negligible or continues to trickle based on 2022 booster trends 	Scenario C	Scenario D
 Reformulated annual vaccination recommended for all currently eligible groups Reformulated vaccine has 65% VE against variants circulating on June 15 Vaccine becomes available September 1 65+ uptake same as first booster dose recommended in September 2021 Coverage in individuals under 65+ saturates at levels of the 2021 booster (approximately 34% nationally) 	Scenario E	Scenario F

SMH – COVID-19 (Round 17) – Virginia Results

- To date, immune escape evolution has been slow. Booster campaign size remains unknown.
- Significant variation in Fall-Winter 2023 outlook across models

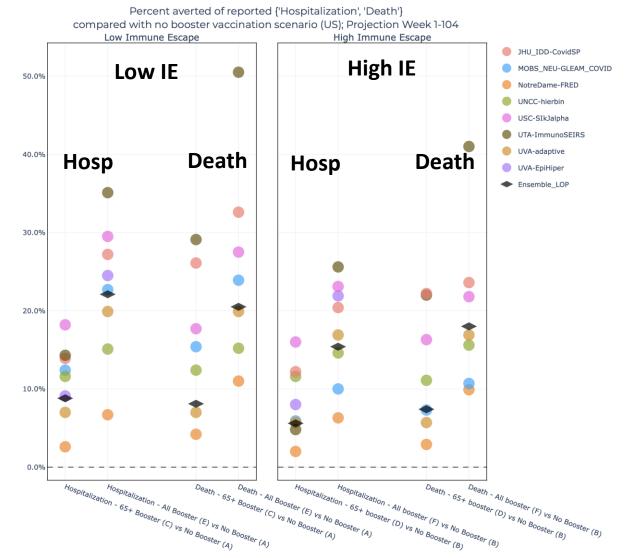


Incident Hospitalization JHU_IDD-CovidSP MOBS_NEU-GLEAM_COVID NotreDame-FRED UNCC-hierbin UVA-adaptive UVA-EpiHiper
 UTA-ImmunoSEIRS USC-SIkJalpha Ensemble_LOP



SMH – COVID-19 (Round 17) – Results – Booster Impact

- Models estimate potential reduction in hospitalizations ranging from 2% - 18% for a 65+ only campaign to 5% - 35% for a whole population campaign
- Reductions in deaths are similar with ensemble estimates of 8% reduction for 65+ campaign and 22% reduction for whole population campaign
- For high immune escape scenarios, the reductions are smaller and more pronounced for deaths than hospitalizations





COVID-19 Activity levels continue to decline

- Declines in cases and hospitalizations have continued
- Other indicators continue to point towards continued declines or suggest no major change
- Wastewater based indicators similar mix of viral loads as in previous weeks

Genomic Surveillance maintains high diversity with no dominating variant

Together this suggests continued declines or easing into a plateau in near term



Questions?

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