### BV-BRC SARS-CoV-2 Emerging Variant Report December 6, 2022

Details of the emerging variants analysis can be found in "BV-BRC SARS-CoV-2 Emerging Variant Report – 20221206.xlsx" based on sequence data from GISAID.

Keep in mind that the information provided reflects sequence counts and sequence proportions and, as such, is impacted by sampling bias in the sequence databases and should not be interpreted as the prevalence of disease caused by these variants.

In addition, due to sequence anomalies (e.g., ambiguous nucleotides in many sequence records) and other issues, the absolute counts of Variants of Concern sequences are likely to be underestimates of the true sequence prevalence.

The key findings are summarized below.

## USA – VOC

## **OMICRON**

- Based on CDC Nowcast estimates (<u>https://covid.cdc.gov/covid-data-tracker/#variant-proportions</u>) for the week ending 12/03/22 in the US: virtually all sequences reported since July are **Omicron**.
  - BA.5 proportions are no longer dominating in the United States at 13.8% (down from 29.7% on 11/12/22)
  - **BA.4.6** proportions continue to decrease, now at 2.3%
  - BQ.1.1 and BQ.1 continue to grow drmatically. BQ.1 is currently estimated at 30.9%, up from last week. BQ.1.1 has overtaken BQ.1 and is currently estimated at 31.9%, up from last week.
  - *BF.7 has been added to the Nowcast tracker and is currently estimated at 6.3%, slightly down from last week*
  - **BA.5.2.6** is steady at 1.8%
  - XBB has been added to the tracker and has increased from last week to 5.5%
  - BN.1 has been added to the tracker and has increased from last week to 4.6%
  - $\circ$  BF.11 has been added to the tracker and has slightly decreased from last week to 0.9%
- Listed below are lineages with sequence prevalence >1.0% or a growth rate >3 fold and count >50 in November (ranked in order of sequence prevalence) in comprehensive data collected from GISAID. Lineages with fold growth >4 from October to November so far are highlighted in *bold italic*:
  - BQ.1.1 10%, 2.6 fold growth
  - $\circ$  BQ.1 8.8%, 2.3 fold growth
  - BA.5.2.1 7.1%, 0.48 fold growth
  - $\circ$  BA.5.2 6.1%, 0.56 fold growth
  - BA.4.6 3.9%, 0.51 fold growth
  - BA.2.75 3.3%, 1.3 fold growth
  - $\circ \quad BA.5.1-3.1\%,\, 0.54 \ fold \ growth$

- $\circ$  BF.7 3.1%, 0.93 fold growth
- $\circ$  BQ.1.12 1.7%, 4.0 fold growth
- *XBB* 1.6%, 5.0 fold growth
- $\circ$  BQ.1.3 1.5%, 2.7 fold growth
- BA.5.2.6 1.5%, 1.3 fold growth
- $\circ$  BQ.1.2 1.3%, 1.9 fold growth
- BQ.1.1.4 1.3%, 5.4 fold growth
- BQ.1.11 1.3%, 3.5 fold growth
- $\circ$  BF.10 1.3%, 0.58 fold growth
- $\circ \quad BQ.1.14-1.2\%,\, 2.5 \ fold \ growth$
- $\circ$  BQ.1.1.3 1.2%, 4.5 fold growth
- $\circ$  BQ.1.1.5 1.2%, 3.5 fold growth
- $\circ \quad BF.26-1.1\%,\, 0.75 \ fold \ growth$
- $\circ$  BA.5 1.1%, 0.59 fold growth
- BE.1.1 1.1%, 0.97 fold growth
- XBB.1 0.91%, 3.8 fold growth
- o BQ.1.13 0.77%, 3.7 fold growth
- o BQ.1.18 0.75%, 8.3 fold growth
- o BQ.1.1.18 0.74%, 3.9 fold growth
- o BQ.1.1.15 0.53%, 53 fold growth
- o CQ.2 0.5%, 10 fold growth
- o BQ.1.1.7 0.4%, 4.4 fold growth
- CR.1.1 0.34%, 11 fold growth
- BQ.1.8 0.33%, 3.3 fold growth
- CK.1 0.31%, 3.1 fold growth
- BQ.1.1.1 0.24%, 4 fold growth
- DELTA (B.1.617.2 and AY sub-lineages) (no significant change since previous report)
  - One Delta sequences in the US in November so far; three in October

<u>USA – (other VOCs and VOIs)</u> (no significant change since previous report)

• None

### <u>USA – Recombinants</u>

- XBB 383 sequences in November so far; 200 sequences in October
- XBB.1 221 sequences in November so far; 153 sequences in October
- XBB.2 24 sequences in November so far; 15 sequences in October
- XBD 6 sequences in November so far; 8 sequences in October
- XBB.3 4 sequences in November so far; 19 sequences in October
- XAZ 4 sequences in November so far; 39 sequences in October
- XBE 3 sequences in November so far; 15 sequences in October
- XAV 1 sequences in November so far; 2 sequences in October
- XBB.4 1 sequences in November so far; 1 sequences in October
- XAS no sequences in November so far; 6 sequences in October
- XBB.1.1 no sequences in November so far; 6 sequences in October
- XBB.1.3 no sequences in November so far; 1 sequences in October

### World – VOC

### **OMICRON**

- Omicron remains dominant globally. Listed below are lineages showing the highest sequence prevalence (> 1.0%) or a fold growth > 3 fold and count >50 in November. Lineages with fold growth >4 from October to November are highlighted in *bold italic*:
  - BQ.1.1 9.4%, 2.2 fold growth
  - BA.5.2 7.5%, 0.53 fold growth
  - BA.2.75 6.2%, 1.9 fold growth
  - BA.5.2.1 6.0%, 0.51 fold growth
  - BQ.1 5.1%, 2.1 fold growth
  - $\circ$  BF.7 5.0%, 0.84 fold growth
  - BA.5.1 3.3%, 0.51 fold growth
  - $\circ$  XBB 2.8%, 5.3 fold growth
  - $\circ$  BF.5 2.5%, 0.66 fold growth
  - $\circ$  BA.4.6 2.2%, 0.60 fold growth
  - BA.5.2.6 1.4%, 1.0 fold growth
  - BQ.1.18 1.2%, 5.2 fold growth
  - BA.2 0.89%, 3.9 fold growth
  - BQ.1.12 0.57%, 3.6 fold growth
  - BQ.1.11 0.56%, 3.1 fold growth
  - BQ.1.1.15 0.5%, 3.3 fold growth
  - BQ.1.1.3 0.49%, 3.3 fold growth
  - BQ.1.1.10 0.45%, 3.8 fold growth
  - o BQ.1.13 0.36%, 3.3 fold growth
  - o CK.1 0.18%, 3 fold growth
  - o BN.1.3.1 0.16%, 5.3 fold growth
  - CQ.2 0.16%, 8 fold growth
  - o BQ.1.1.13 0.15%, 5 fold growth
  - BQ.1.10.1 0.11%, 3.7 fold growth
  - o BN.1.4 0.1%, 5 fold growth
  - o CR.1.1 0.1%, 10 fold growth
  - BN.1.2 0.09%, 3 fold growth
  - XBC.1 0.08%, 4 fold growth
- Substitutions in spike that we are monitoring (>3 fold growth from October to November with counts >50) include those listed below (but note that the numbers are still relatively small for some of these). Only one of substitutions (G252V) has a sequence prevalence >1% in November:
  - N148Q 0.24%, 12 fold growth
  - Y144L 0.24%, 12 fold growth
  - Y145P 0.24%, 12 fold growth
  - S673G 0.11%, 11 fold growth
  - N149Q 0.24%, 8 fold growth
  - o F486P 0.74%, 7.4 fold growth

- o H146K 0.26%, 6.5 fold growth
- o Q613H 0.19%, 4.8 fold growth
- V62I 0.14%, 4.7 fold growth
- o F486I 0.63%, 4.2 fold growth
- $\circ~$  M177T 0.12%, 4 fold growth
- K182I 0.08%, 4 fold growth
- K147Q 0.11%, 3.7 fold growth
- o T859S 0.11%, 3.7 fold growth
- $\circ~$  G485D 0.07%, 3.5 fold growth
- K147- 0.19%, 3.2 fold growth
- N148- 0.19%, 3.2 fold growth
- G252V 2.6%, 3.1 fold growth
- D253G 0.7%, 3.0 fold growth
- o L18F 0.06%, 3 fold growth
- o T19R 0.06%, 3 fold growth
- $\circ$  T208M 0.06%, 3 fold growth
- E156G 0.15%, 3 fold growth
- R158- 0.15%, 3 fold growth

Note on convergent mutations: In addition to previously noted convergent mutations in the receptor binding domain (RBD) ( $L452^*$ :  $R346^*(T/I/S/K)$ ,  $K356^*(T/R)$ ,  $K444^*(M/T/R/N)$ ,  $V445^*(A/P)$ ,  $G446^*(D/S/V)$ ,  $N450^*$ ,  $N460^*(K/S)$ ,  $478^*$ ,  $485^*$ , & F486^\*), we have begin to notice convergent mutations in the N-terminal domain (NTD). More details are provided below:

## Positions between 144>152

144: in addition to Y144- (which has grown from  $\sim 2\%$  to  $\sim 19\%$  prevalence in the past 3 months) a bump in November of genomes with Y144L (only 207 genomes,  $\sim 0.2\%$ ) has also been noticed. Y144L has increased by <u>12 fold</u> in the past month.

145: Commonly seen mutations at this position include Y145P ( $\sim 0.2\%$ ), Y145H( $\sim 0.4\%$ ), & Y145-( $\sim 0.4\%$ ). Y145P has increased by <u>12 fold</u> in the past month.

146: Commonly seen mutations at this position include H146K (~0.3%), H146L (~0.2%), H146Q (~0.4%), & H146- (~0.1%). Frequency of multiple mutations at this position have increased in the past month: H146K (<u>6.5 fold</u> increase) H146L (~<u>3 fold</u> increase), H146Q (~<u>2.5 fold</u> increase).

147: Commonly seen mutations at this position include K147Q- (~0.1%), K147- (~0.2%), K147E

(~9% mainly from BA.2.75 lineages), K147I (~0.2%), K147N (~0.4%), & K147T (~0.1%).

Mutations at both K147Q & K147- have increase by more than <u>3 fold</u> in the past month.

148: Commonly seen mutations at this position include N148Q (~0.2%) & N148- (~0.2%). These have increased by <u>12 fold</u> and ~<u>3.2 fold</u> respectively in the past month.

149: One main mutation at this position: N149Q ( $\sim 0.2\%$ ), but has increased by <u>8 fold</u> in the past month.

**150**: One main mutation at this position **K150E** ( $\sim$ 0.4%), but has increased by <u>3 fold</u> in the past month.

Small sporadic increases in mutations at position 150, 151, 152 (BA.2.75 lineages).

## Positions between 156-158

There has been an increase in mutations at this position, including E156G, E156-, F157-, F157L, and R158-. Some of these are Delta like mutations and are attributable to recent increases in Delta/Omicron recombinant lineages such as **XBC** and **XAY** (and a few XBA). However there are also some BA.2.75

sublineages with E156-, F157- (mainly from Turkey), and of course most BA.2.75 already have **F157L**.

Positions between 208-215

Moderate increases in proportions of variants carrying mutations at these positions. These numbers are generally low, but appear to all occur in similar regions. This is a similar pattern to what was observed in Delta.

208: T208M (<0.1%, 3 fold increase in November)

209: **P209L** (0.5%, 2 fold increase in November)

210: I210V (7%, 1.9 fold increase in November, mainly due to BA.2.75 background)

211: N211- (0.15%, 2 fold increase in November, mainly occurs in BA.2.75 lineages)

212: L212I and L212S (both at ~0.15%, ~2 fold increase in November)

213: V213E and V213G (found in more than 95% genomes, both of these mutations are found in currently circulating XBB and BA.2/BA.4/5 lineages)

215: D215G (~0.1%, 2.6 fold increase) & D215H (~0.2% and 2 fold increase)

## Positions between 243-253

While mutations at these particular positions do not appear to be particularly convergent (different mutations at same site), the general area between these coordinates seems to be experiencing evolutionary pressure.

243 & 244: Deletions (A243-,L244-) occur simultaneously at both these positions in XBC\* lineages (~0.1%, 2.2 fold increase)

245: H245N, H245Y (0.9% and 0.1% respectively, no fold increase)

247: **S247N** (0.2%, 2.2 fold increase, BF\*)

248: **Y248-** (0.15%, 2 fold increase)

251: **P251H** (0.18%, 2.2 fold increase)

- 252: G252V (0.26%, 3 fold increase)
- 253: G253G (0.7%, 3 fold increase)

# **DELTA (B.1.617.2** and **AY** sub-lineages)

• 12 Delta genomes in November

# World (other VOIs)

• 2 Alpha genome in November so far

# <u> World – Recombinants</u>

- XBB 2394 sequences in November so far; 1371 sequences in October
- XBB.1 630 sequences in November so far; 1646 sequences in October
- XBB.2 81 sequences in November so far; 206 sequences in October
- XAZ 32 sequences in November so far; 245 sequences in October
- XBB.3 30 sequences in November so far; 244 sequences in October
- XBD 27 sequences in November so far; 82 sequences in October
- XBB.1.1 19 sequences in November so far; 143 sequences in October
- XBB.4 12 sequences in November so far; 22 sequences in October
- XBE 3 sequences in November so far; 52 sequences in October
- XBB.5 2 sequences in November so far; 11 sequences in October

- XBC 2 sequences in November so far; 15 sequences in October
- XAV 1 sequences in November so far; 3 sequences in October
- XBB.1.3 1 sequence in November so far; 7 sequences in October
- XAH no sequences in November so far; 1 sequence in October
- XAK no sequences in November so far; 1 sequence in October
- XAY no sequences in November so far; 10 sequences in October
- XBB.1.2 no sequences in November so far; 5 sequences in October
- XBB.3.1 no sequences in November so far; 20 sequences in October
- XE no sequences in November so far; 1 sequence in October
- XAN no sequences in November so far; 3 sequences in October
- XAS no sequences in November so far; 7 sequences in October

## Variants that have been mentioned in the media and/or social media:

- Boston-area COVID Wastewater Data Spikes 88%
- Covid hospitalizations rising post-Thanksgiving after an autumn lull
- <u>COVID-19 cases spike all across Florida after Thanksgiving holiday</u>
- <u>Covid evolution wipes out another antibody treatment, threatening the country's medicine</u> <u>cabinet</u>
- <u>SARS-CoV-2 BW.1</u>, a fast-growing Omicron variant from southeast Mexico bearing relevant escape mutations
- <u>Gradual emergence followed by exponential spread of the SARS-CoV-2 Omicron variant in</u> <u>Africa</u>
- <u>Convergent evolution of the SARS-CoV-2 Omicron subvariants leading to the emergence of BQ.1.1 variant</u>

## **Recombinant lineages:**

One new recombinant lineage has been added this week. A summary table of recombinant lineages can be found below with more details. We have added the number of recombinants genomes for each lineage. XA, XB, and XC have not been detected in the past 6 months. XD-XBJ are Omicron/Omicron or Omicron/Delta recombinants. With the exception of XBB, the majority of these recombinant lineages do not appear to have taken off and several have not been detected in recent weeks. The XBB lineage appears to have the largest number of genomes.

Name	number	Github#	Lineage 1	Lineage 2	Location first detected
XA	45	NA	B.1.1.7	B.1.177	UK
<u>XB</u>	3435	<u>#189</u>	B.1.634	B.1.631	N. America
<u>XC</u>	25	<u>#263</u>	Delta (AY.29)	B.1.1.7	Japan
<u>XD</u>	34	<u>#444</u>	Delta (AY.4)	BA.1	France

<u>XE</u>	2836	<u>#454</u>	BA.1	BA.2	UK
XF	34	<u>#445</u>	Delta	BA.1	UK
XG	479	<u>#447</u>	BA.1	BA.2	Denmark
<u>XH</u>	174	<u>#448</u>	BA.1	BA.2	Denmark
<u>XJ</u>	253	<u>#449</u>	BA.1	BA.2	Finland
XK	25	<u>#460</u>	BA.1	BA.2	Belgium
<u>XL</u>	120	<u>#464</u>	BA.1	BA.2	UK
<u>XM</u>	526	<u>#472</u>	BA.1.1	BA.2	Multiple EU
XN	288	<u>#480</u>	BA.1	BA.2	UK
<u>XP</u>	57	<u>#481</u>	BA.1.1	BA.2	UK
XQ	145	<u>#468</u>	BA.1.1	BA.2	UK
XR	183	<u>#469</u>	BA.1.1	BA.2	UK
<u>XS</u>	60	<u>#471</u>	Delta	BA.1.1	USA
<u>XT</u>	14	<u>#478</u>	BA.2	BA.1*	S. Africa
<u>XU</u>	16	<u>#522</u>	BA.1*	BA.2	India
XV	42	<u>#463</u>	BA.1*	BA.2*	Denmark
XW	195	<u>#591</u>	BA.1*	BA.2*	JP, DE, SI, CA, UK, US
<u>XY</u>	126	<u>#606</u>	BA.1*	BA.2*	FR, IL, UK, US
XZ	216	<u>#636</u>	BA.2*	BA.1*	Multiple
XAA	104	<u>#664</u>	BA.1*	BA.2*	US, IL
XAB	114	# <u>665</u>	BA.1*	BA.2*	IT, FR, DE, CH, DK
XAC	207	# <u>590</u>	BA.1*	BA.2*	IL, DE, CA, IR, NL, JP, UK, US
XAD	99	<u>#607</u>	BA.2*	BA.1*	CZ,DE,UK
XAE	133	<u>#637</u>	BA.2*	BA.1*	CA,US,NL,CH
XAF	300	<u>#676</u>	BA.1*	BA.2*	CR
XAG	412	<u>#709</u>	BA.1.1	BA.2.23	BR
XAH	169	<u>#755</u>	BA.2*	BA.1*	SI
XAJ	106	<u>#826</u>	BA.2.12.1	BA.4	UK
XAK	149	<u>#823</u>	BA.1*	BA.2*	DE
XAL	122	<u>#757</u>	BA.1.1	BA.2*	DE
XAM	546	<u>#759</u>	BA.1.1	BA.2.9	Panama
XAN	213	<u>#771</u>	BA.2*	BA.5.1	Multiple

XAP	359	<u>#789</u>	BA.2*	BA.1*	US
XAQ	81	<u>#798</u>	BA.1*	BA.2*	Canada
XAR	21	#860	BA.1*	BA.2*	Reunion/France
XAS	192	<u>#882</u>	BA.5*	BA.2*	N. America
XAT	32	<u>#885</u>	BA.2.3.13	BA.1*	Japan
XAU	44	<u>#894</u>	BA.1.1*	BA.2.9*	Multiple
XAV	82	<u>#911</u>	BA.2*	BA.5*	France
XAW	45	<u>#895</u>	BA.2*	AY.122	Russia
XAY*	71	<u>#844</u>	BA.2	AY.45	S. Africa
XAZ	2348	<u>#797</u>	BA.2.5, BA.5	BA.2.5	Multiple
XBA	4603	<u>#844</u>	AY.45	BA.2	S. Africa
XBB*	9868	# <u>1058</u>	BJ.1	BM.1.1.1	Singapore/US
XBC*	297	#1100	Delta	BA.2	Philippines
XBD	388	<u>#1137</u>	BA.2.75.2	BF.5	Multiple
XBE*	5397	<u>#1246</u>	BA.5.2	BE.4	USA
XBF	618	#1259	BA.5.2.3	CJ.1	Australia
XBG	50	<u>#896</u>	BA.2.76	BA.5.2	UK
XBH	~100	<u>#1229</u>	BA.2.3.17	BA.2.75.2	Multiple
XBJ	~25	#1268	BA.2.3.20	BA.5.2	Multiple

# Newly designated Pango lineages:

New lineages designated this week:

- BA.5.2.45: Sublineage of BA.5.2 (ORF1b:T1050N) with S:K444Q & ORF7a:A55V (92 seq)
- CH.3: BM.4.1.1 sublineage with S:F486P recently emerged in Indonesia and already exported to New Zealand, Denmark and Australia 14 sequences
- CV.2: New BM.1.1.3 sublineage with S:F486P + ORF1a:G514E , ORF1b:Y1147C circulating in Chile and Usa (28 sequences)
- BQ.1.26.1: Potential BQ.1.26 sublineage with ORF1a:S248N (136 good seqs as of 2022-12-03; Denmark, USA, Germany)