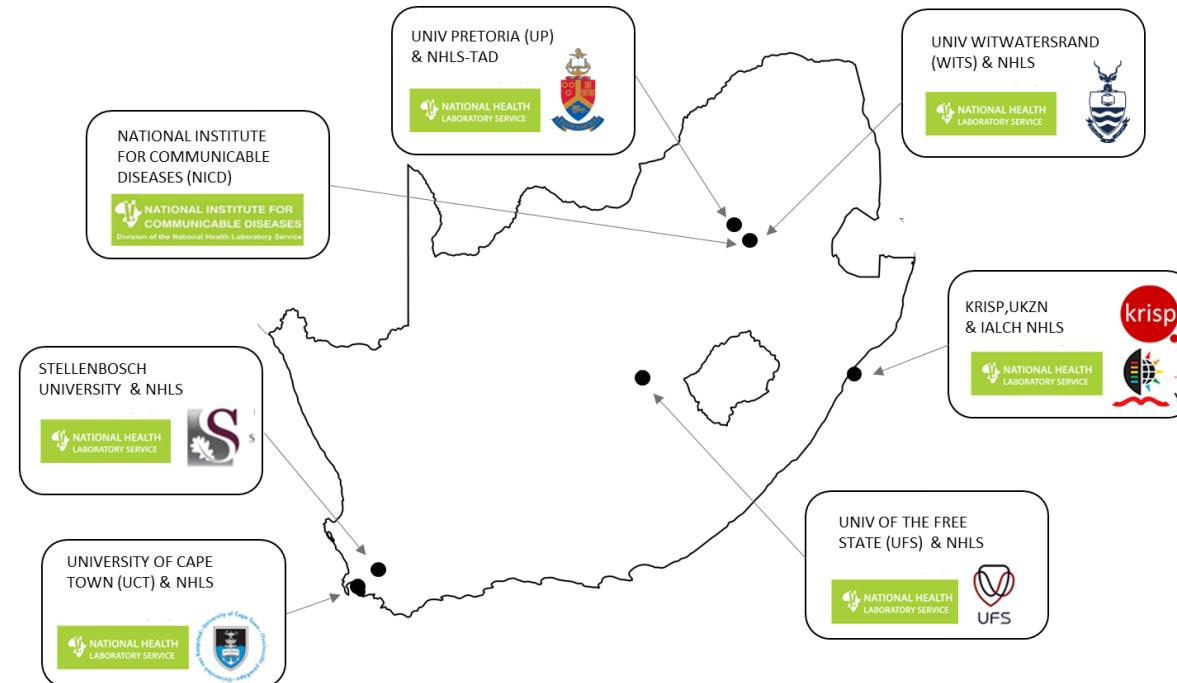


## SARS-CoV-2 Sequencing Update

### 6 May 2022

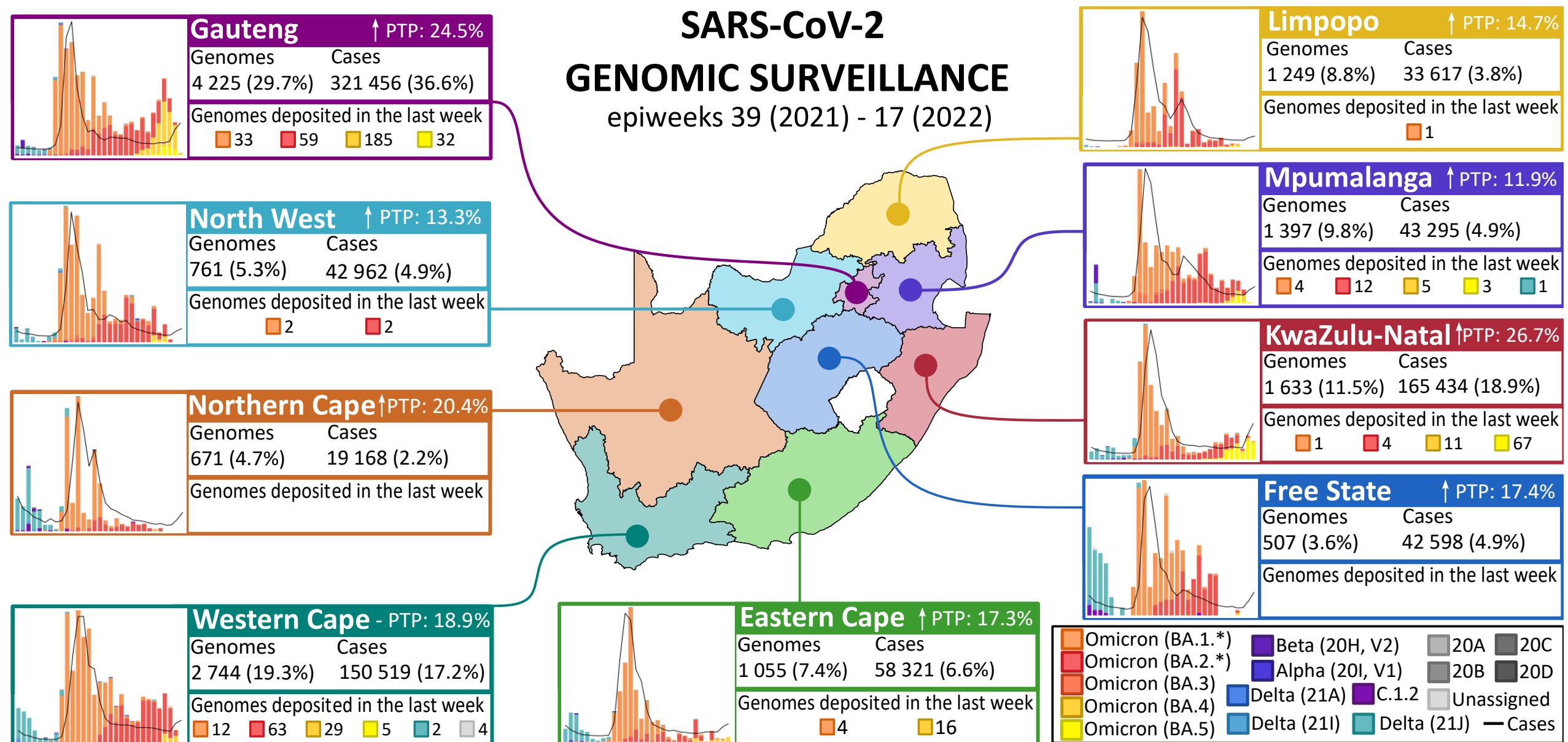


Supported by the DSI and the SA MRC

Msom N, Mlisana K, et al. Lancet Microbe 2020

# SARS-CoV-2 GENOMIC SURVEILLANCE

epiweeks 39 (2021) - 17 (2022)



557 genomes added since the previous report

Bar graphs represent genomes sequenced per epiweek, with lines representing cases by collection date (weeks 39 [2021] – 17 [2022])

Genomes and cases presented as provincial total (percentage of national total) for epiweeks 39 (2021) – 17 (2022)

PTP: percentage testing positive in week 17 (24 Apr 2022 – 30 May 2022); arrow indicates direction of change since previous week (17 Apr 2022 – 23 Apr 2022) if change was significant ( $P < 0.05$ )



The genomic data presented here are based on South African SARS-CoV-2 sequence data downloaded from GISAID ([www.gisaid.org](https://www.gisaid.org)) on 6 May 2022 at 14h23



Data license: <https://www.gisaid.org/registration/terms-of-use/>

Elbe, S., and Buckland-Merrett, G. (2017) Data, disease and diplomacy: GISAID's innovative contribution to global health. *Global Challenges*, 1:33-46. DOI: 10.1002/gch2.1018 PMCID: 31565258

Shu, Y., McCauley, J. (2017) GISAID: Global initiative on sharing all influenza data – from vision to reality. *EuroSurveillance*, 22(13) DOI: 10.2807/1560-7917.ES.2017.22.13.30494 PMCID: PMC5388101

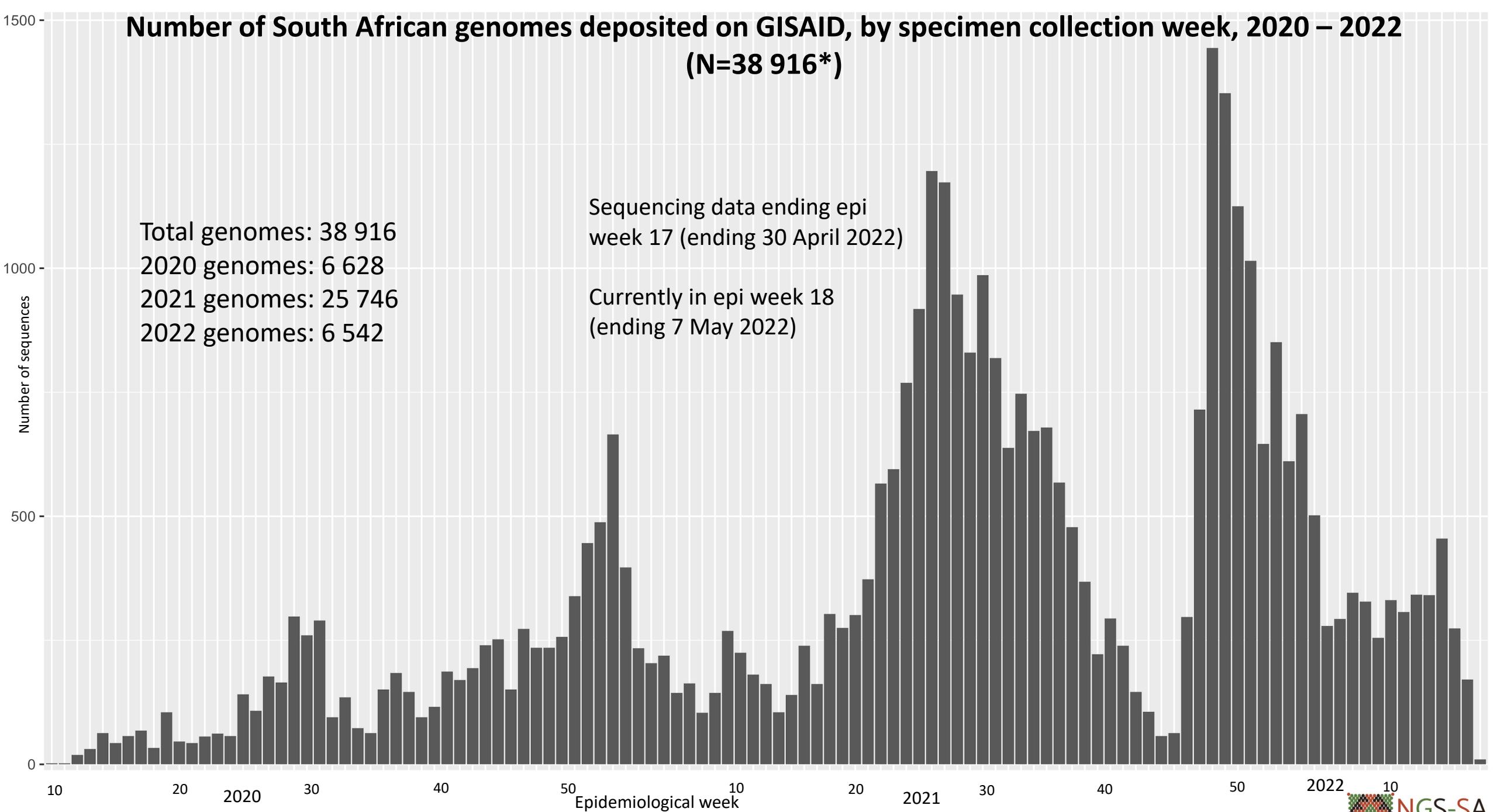
# Number of South African genomes deposited on GISAID, by specimen collection week, 2020 – 2022 (N=38 916\*)

Total genomes: 38 916  
2020 genomes: 6 628  
2021 genomes: 25 746  
2022 genomes: 6 542

Sequencing data ending epi week 17 (ending 30 April 2022)

Currently in epi week 18 (ending 7 May 2022)

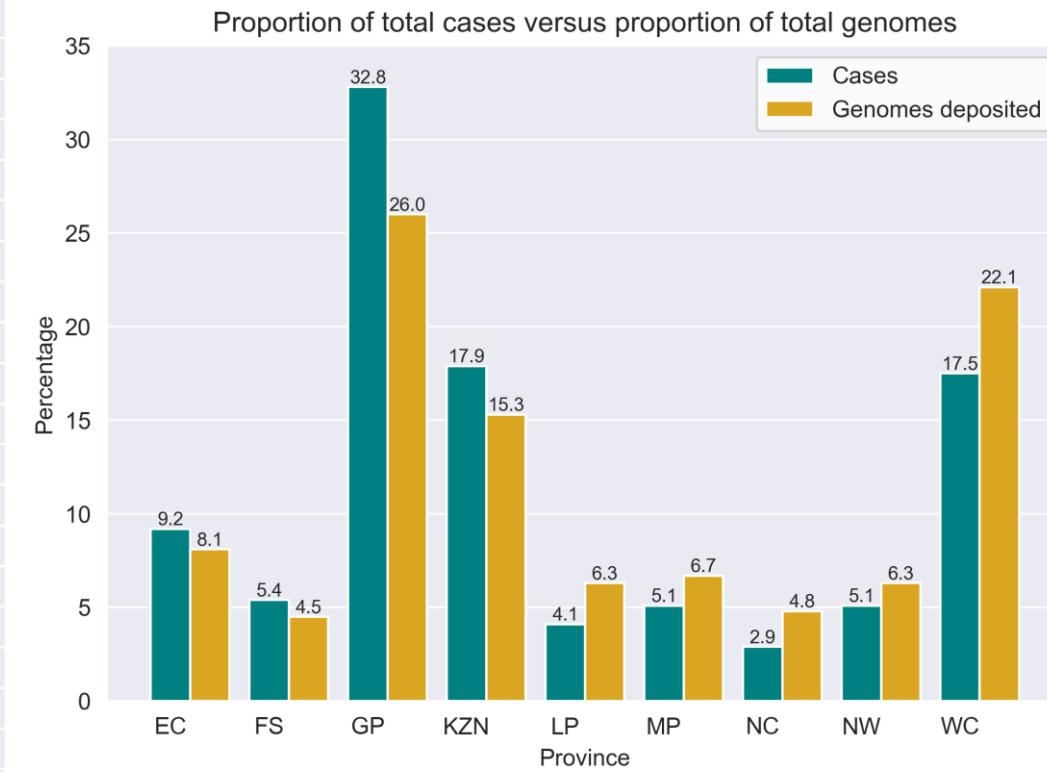
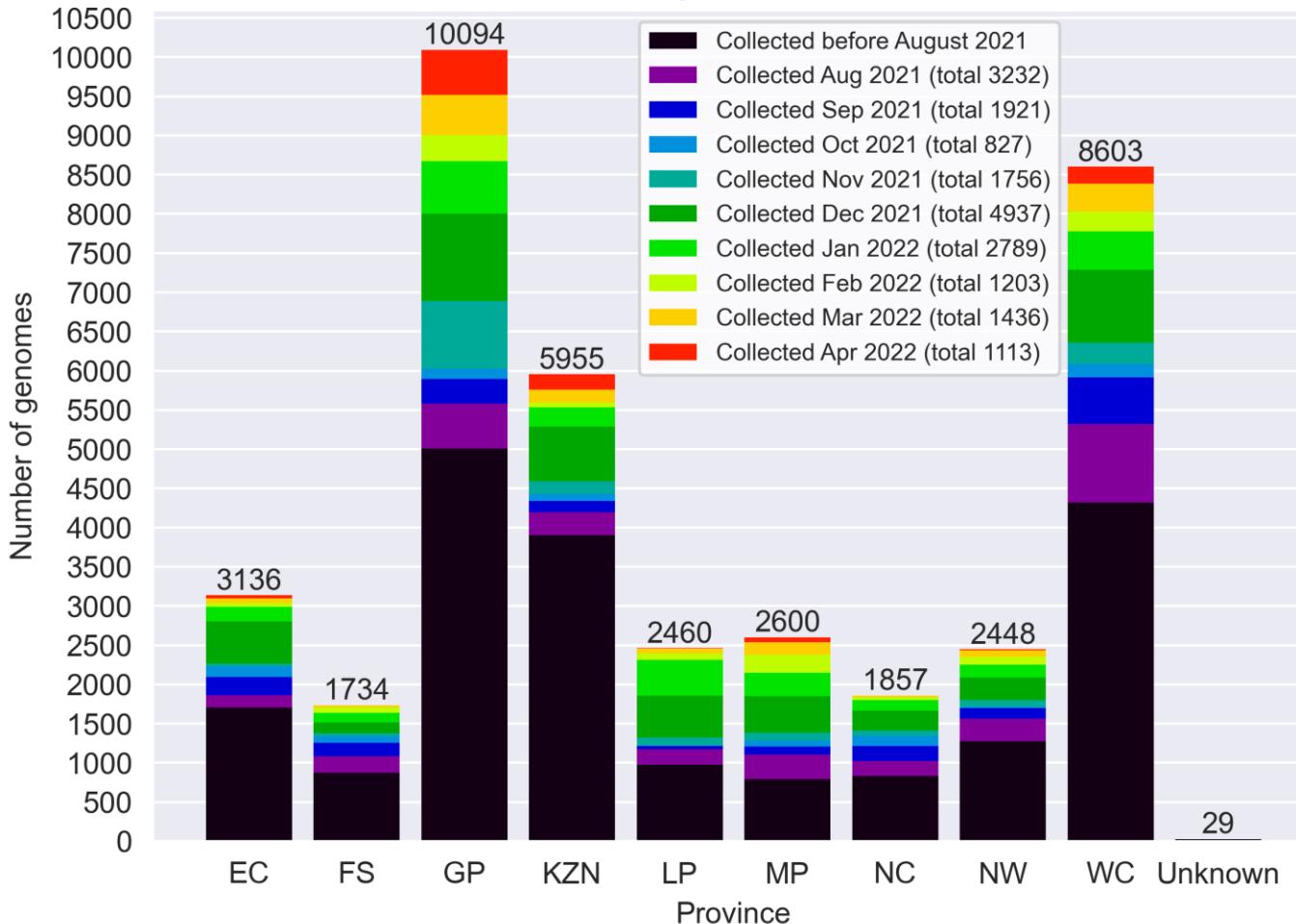
Number of sequences



\*This represents the cleaned, de-duplicated dataset of unique National and Pneumonia Surveillance sequences. This dataset will be used for all further figures.

# GISAID genomes vs total cases, 2020 – 2022 (N=38 916)

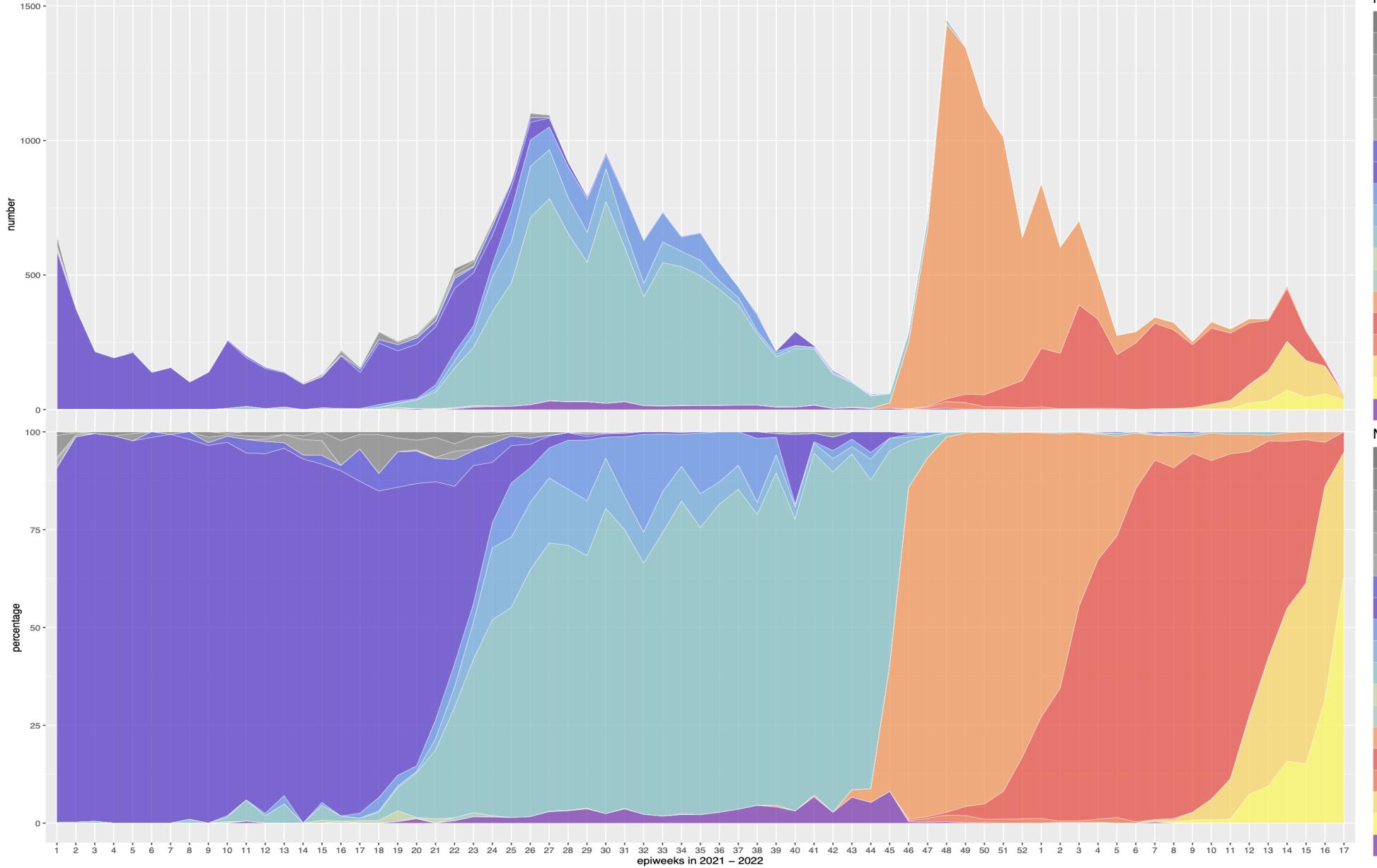
Provincial breakdown of genomes deposited into GISAID



All provinces, apart from GP, KZN, LP, MP, NC and WC, have comparable percentages of overall cases and overall sequenced genomes.

# Number and percentage of clades by epiweek in South Africa, 2021 – 2022 (32 203\*)

Distribution of Nextclade Clades in South Africa



Nextclade clades

- 19A
- 19B
- 20A
- 20B
- 20C
- 20D
- Alpha (20I, V1)
- Beta (20H, V2)
- Delta (21A)
- Delta (21I)
- Delta (21J)
- Eta (21D)
- Kappa (21B)
- Omicron (21K/BA.1.\*)
- Omicron (21L/BA.2.\*)
- Omicron (21M/BA.3)
- Omicron (22A/BA.4)
- Omicron (22B/BA.5)
- C.1.2 (20D)

Sequencing data  
ending epi week 17  
(ending 30 April  
2022)

Currently in epi  
week 18 (ending 7  
May 2022)

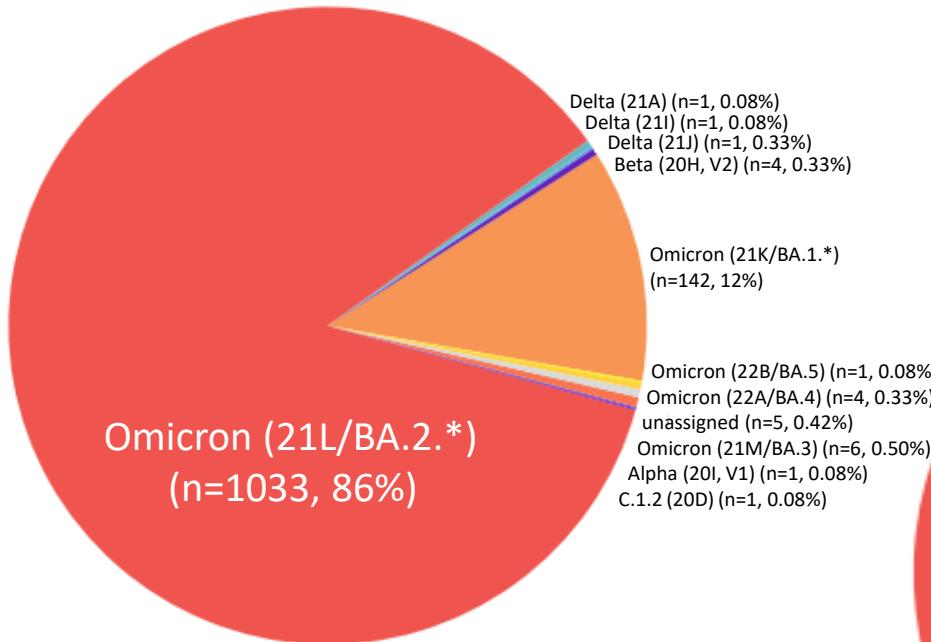
\*Excludes sequences  
missing collection dates,  
as well as those collected  
January 1<sup>st</sup> and 2<sup>nd</sup> 2021  
as they are part of  
epiweek 53 of 2020.

Delta dominated in South Africa until October at >80%. Omicron has dominated from November onwards.

# Prevalence of Variants of Concern (VOC) and Variants of Interest (VOI) in

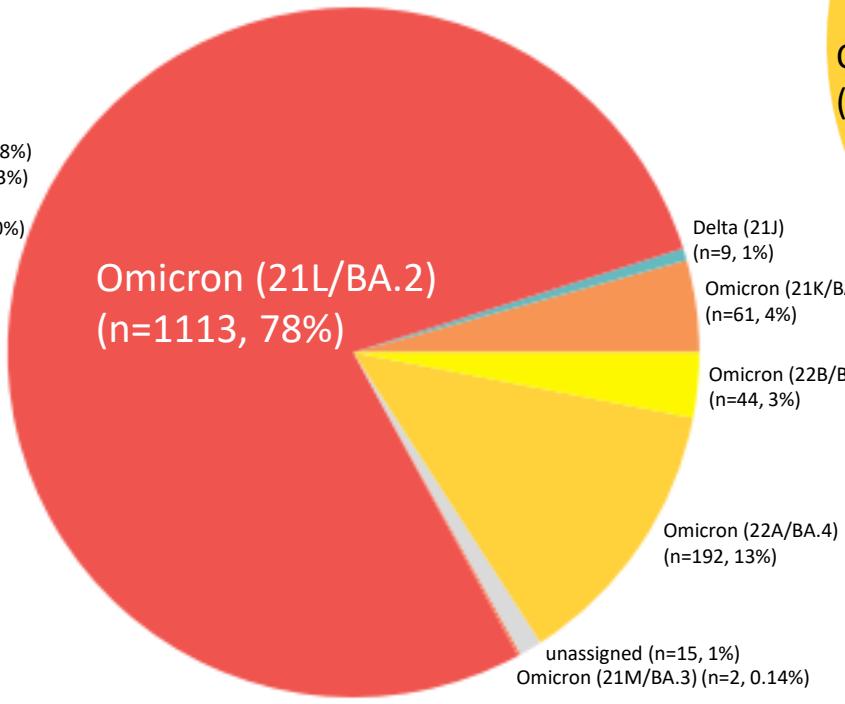
**Feb – Apr 2022**

**February (N=1203)**



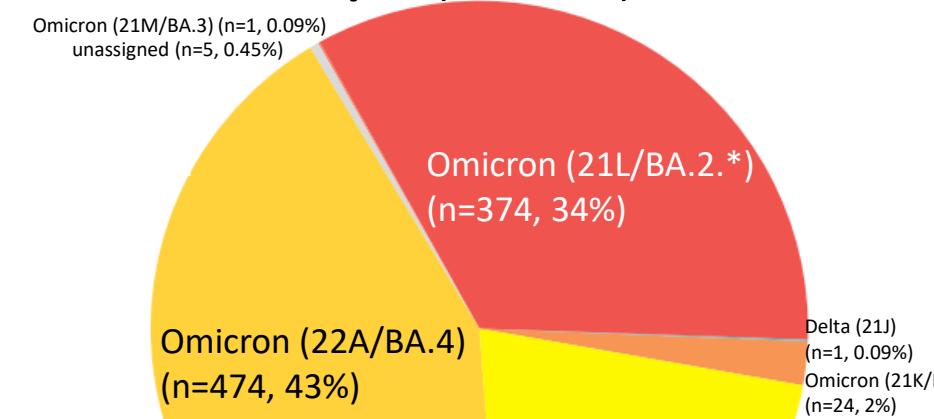
Total Omicron in Feb: 1186 (98.6%)

**March (N=1436)**



Total Omicron in Mar: 1412 (98.3%)

**April (N=1113)**



Total Omicron in Apr: 1106 (99.4%)

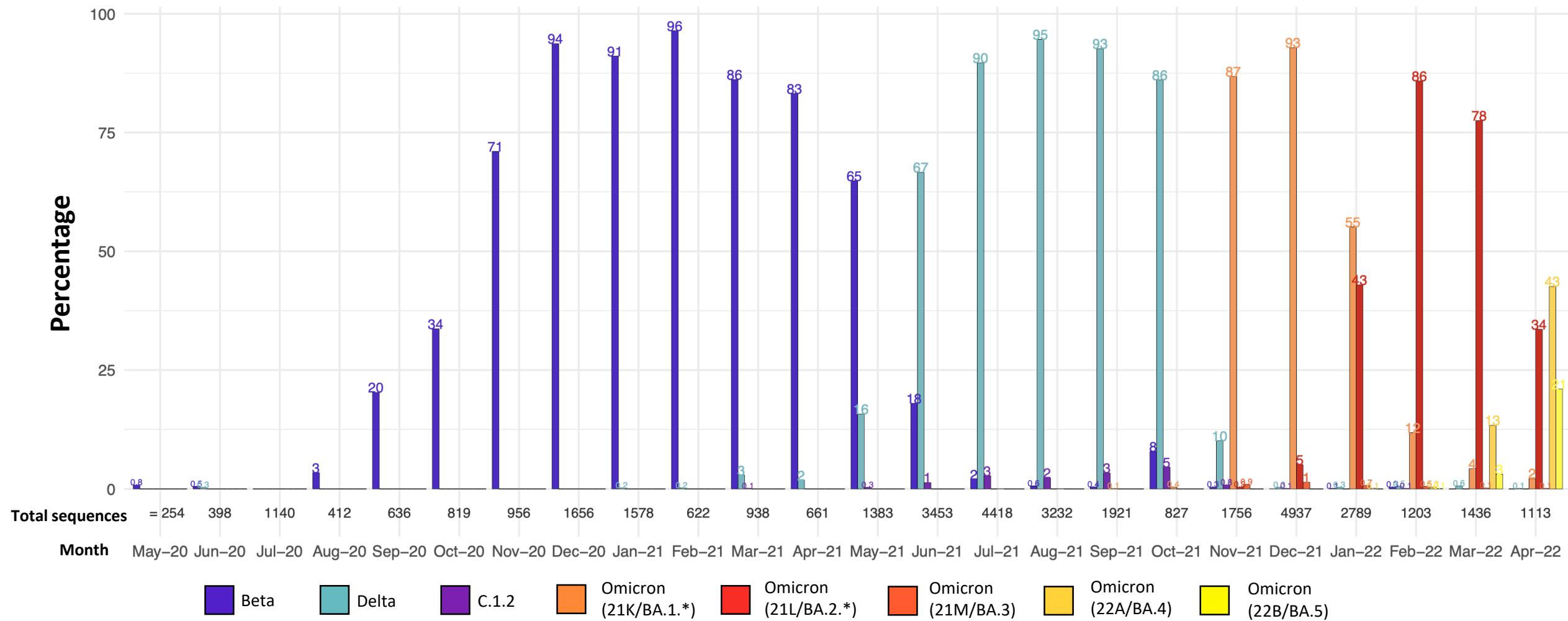
Legend:

- Omicron (21K/BA.1.\*)
- Omicron (21L/BA.2.\*)
- Omicron (21M/BA.3)
- Omicron (22A/BA.4)
- Omicron (22B/BA.5)
- Delta (21I)
- Delta (21J)
- C.1.2 (20D)
- Delta (21A)
- Beta (20H, V2)
- Alpha (20I, V1)
- Kappa (21B)
- Eta (21D)
- Other
- unassigned

Omicron dominated in February (98.6%, 1186/1203), March (98.3%, 1412/1436) and April (99.4%, 1106/1113). BA.2 was dominant in February and March. BA.4 and BA.5 together were dominant in April.

# Detection Rates: Beta, Delta, C.1.2 and Omicron

Detection rates of variants being monitored in South Africa

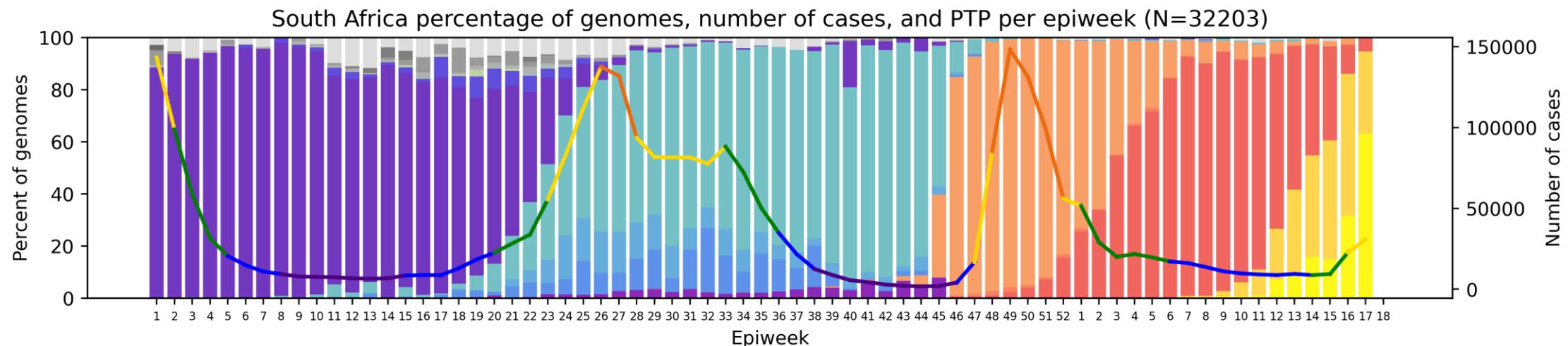
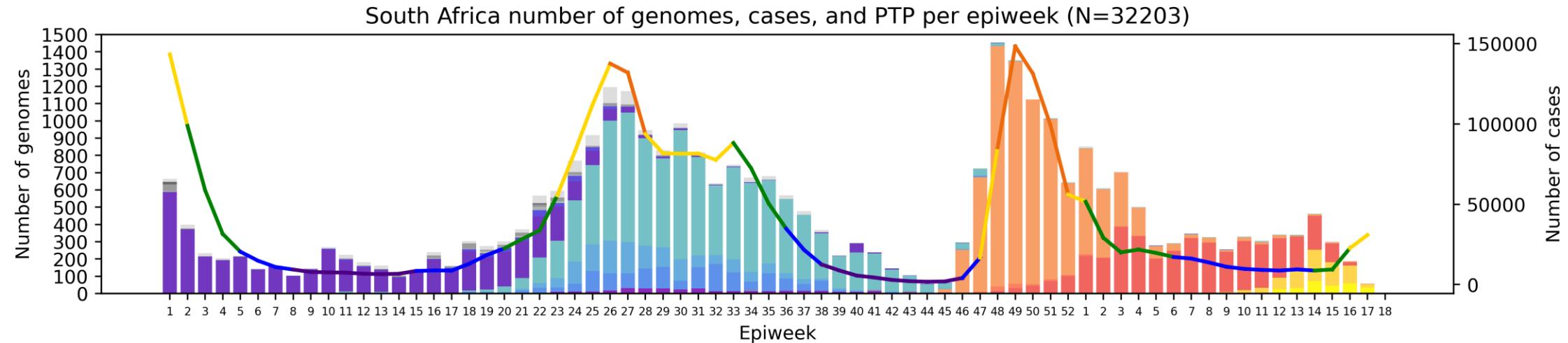


\*Bars represent percentage prevalence of variant for the month; total number sequences collected for the month are given below the bar

Omicron has been dominant since November (>85% in November, >98% in December – April).

BA.2 made up 43% of genomes in January, 86% in February, 78% in March and 34% in April. Newly designated lineages BA.4 and BA.5 together are dominant in April, at 43% and 21% respectively.

# South Africa, 2021-2022, n = 32203\*



Clade key (bar graph)

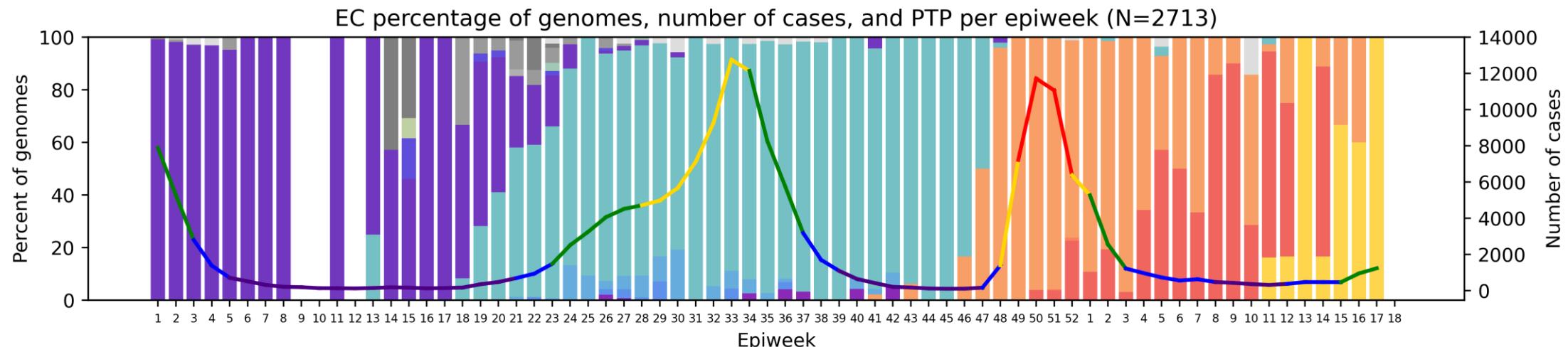
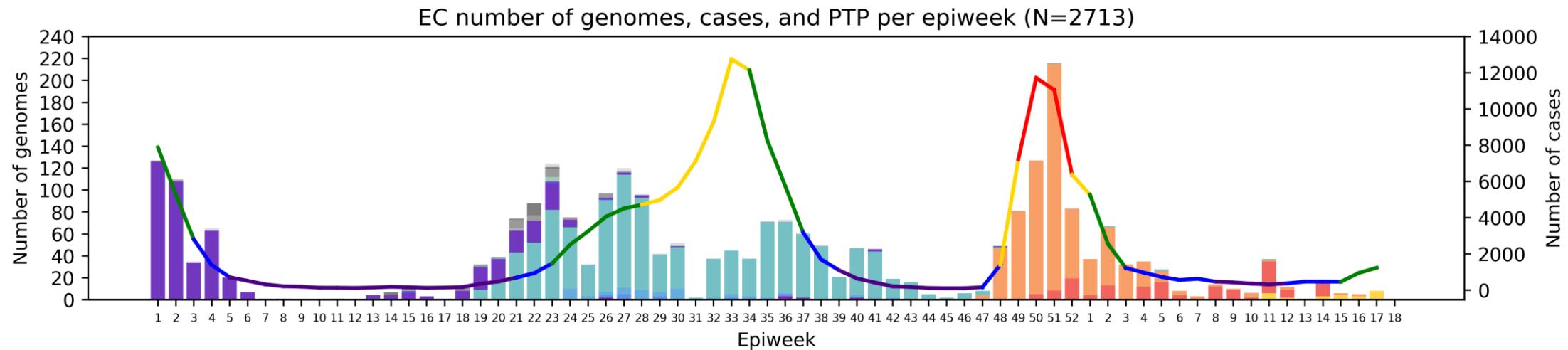
Omicron (21K/BA.1.*)	Omicron (21M/BA.3)	Omicron (22B/BA.5)	Beta (20H, V2)	Delta (21A)	Delta (21J)	Kappa (21B)	Eta (21D)	20D	20C	20B	20A	19B	19A	unassigned
Omicron (21L/BA.2.*)	Omicron (22A/BA.4)	C.1.2 (20D)	Alpha (20I, V1)	Delta (21I)	Delta (21)									

Weekly percentage testing positive key (line graph)

≤ 5	6 - 10	11 - 20	21 - 30	31 - 40	41 - 50	51 - 55
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\*Excludes sequences missing collection dates, as well as those collected January 1<sup>st</sup> and 2<sup>nd</sup> 2021 as they are part of epiweek 53 of 2020.

# Eastern Cape Province, 2021-2022, n = 2713



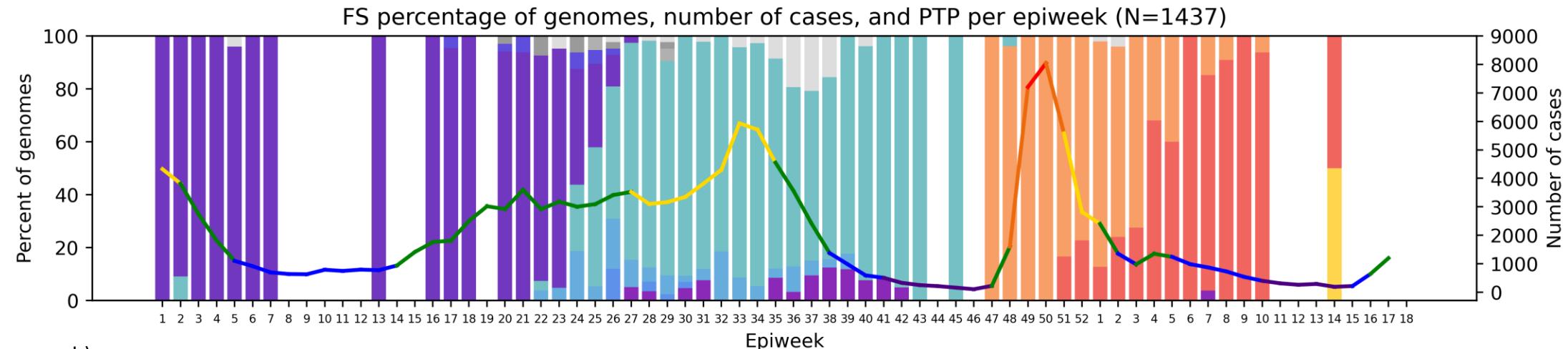
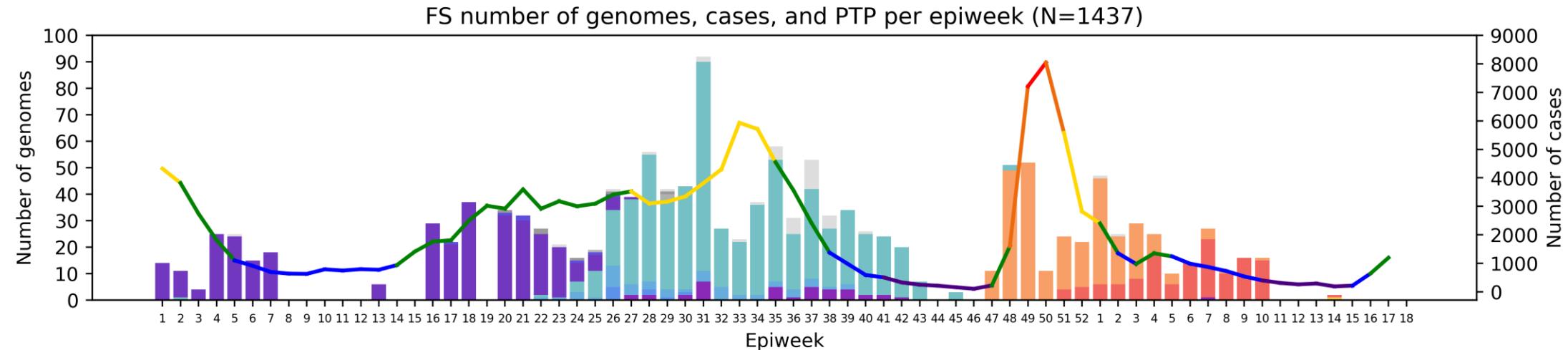
Clade key (bar graph)

Omicron (21K/BA.1.*)	Omicron (21M/BA.3)	Omicron (22B/BA.5)	Beta (20H, V2)	Delta (21A)	Delta (21J)	Kappa (21B)	Eta (21D)	20D	20C	20B	20A	19B	19A	unassigned
Omicron (21L/BA.2.*)		Omicron (22A/BA.4)	C.1.2 (20D)	Alpha (20I, V1)	Delta (21I)									

Weekly percentage testing positive key (line graph)

≤ 5	6 - 10	11 - 20	21 - 30	31 - 40	41 - 50	51 - 55
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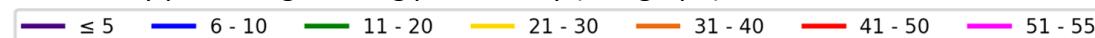
# Free State Province, 2021-2022, n = 1437



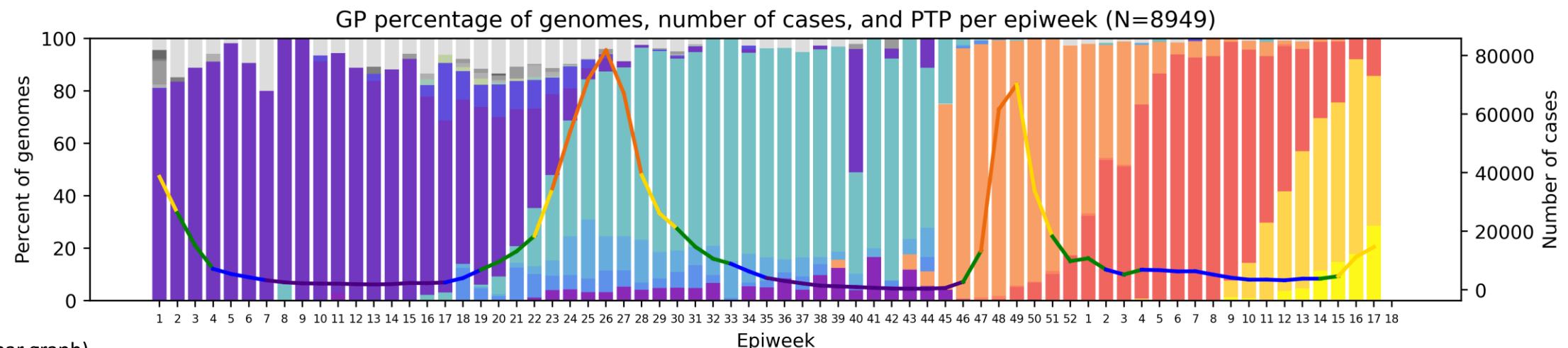
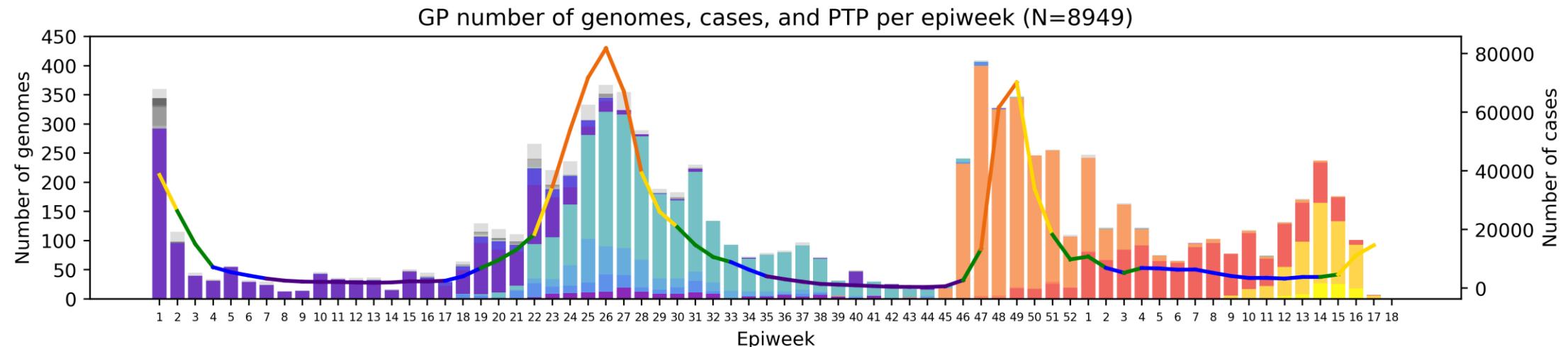
Clade key (bar graph)



Weekly percentage testing positive key (line graph)



# Gauteng Province, 2021-2022, n = 8949



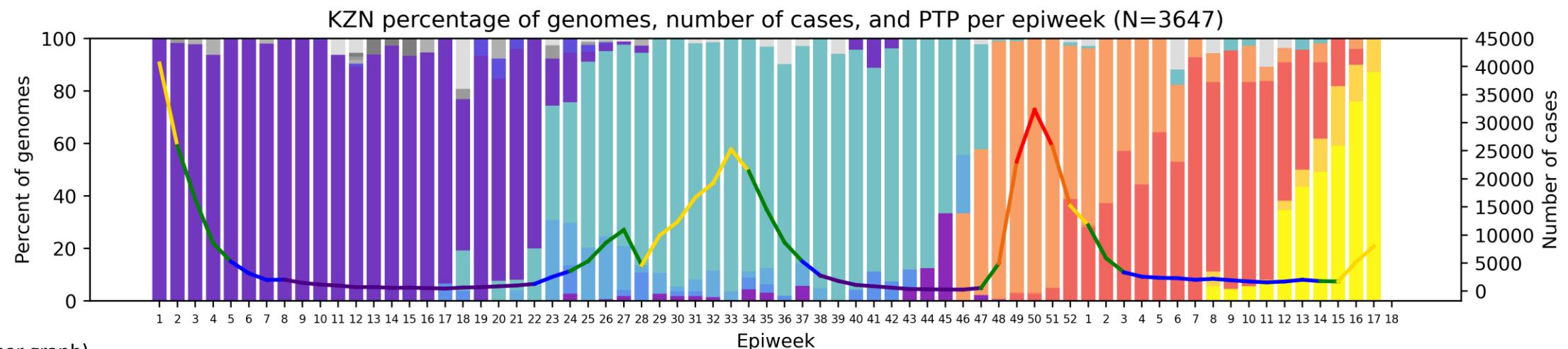
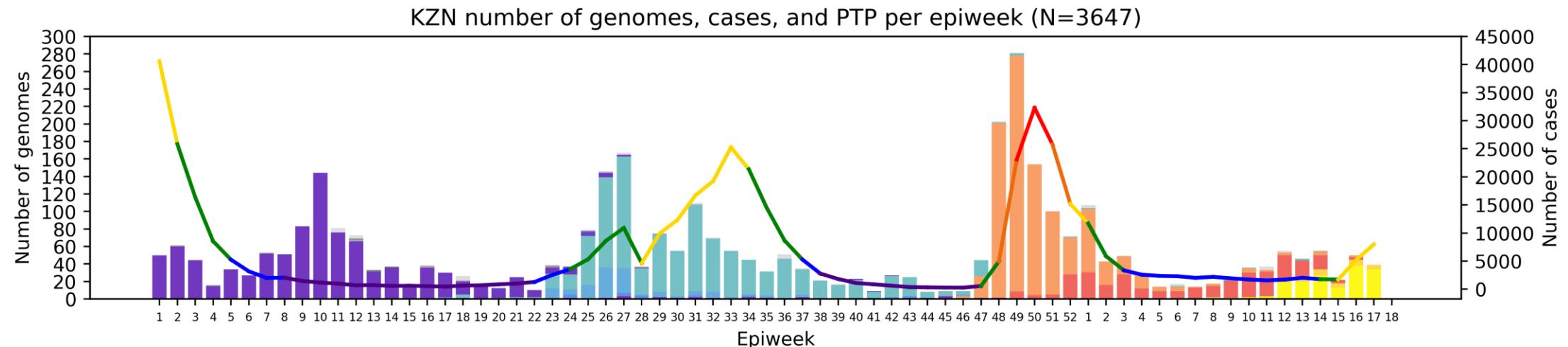
Clade key (bar graph)

Omicron (21K/BA.1.*)	Omicron (21M/BA.3)	Omicron (22B/BA.5)	Beta (20H, V2)	Delta (21A)	Delta (21J)	Kappa (21B)	Eta (21D)	20D	20C	20B	20A	19B	19A	unassigned
Omicron (21L/BA.2.*)	Omicron (22A/BA.4)	C.1.2 (20D)	Alpha (20I, V1)	Delta (21I)	Delta (21)									

Weekly percentage testing positive key (line graph)

≤ 5	6 - 10	11 - 20	21 - 30	31 - 40	41 - 50	51 - 55
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# KwaZulu-Natal Province, 2021-2022, n = 3647



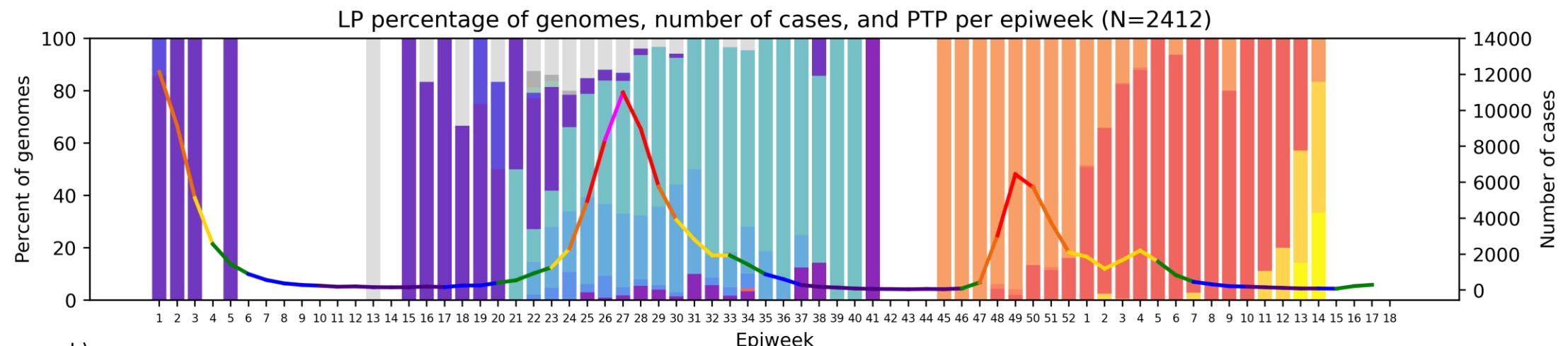
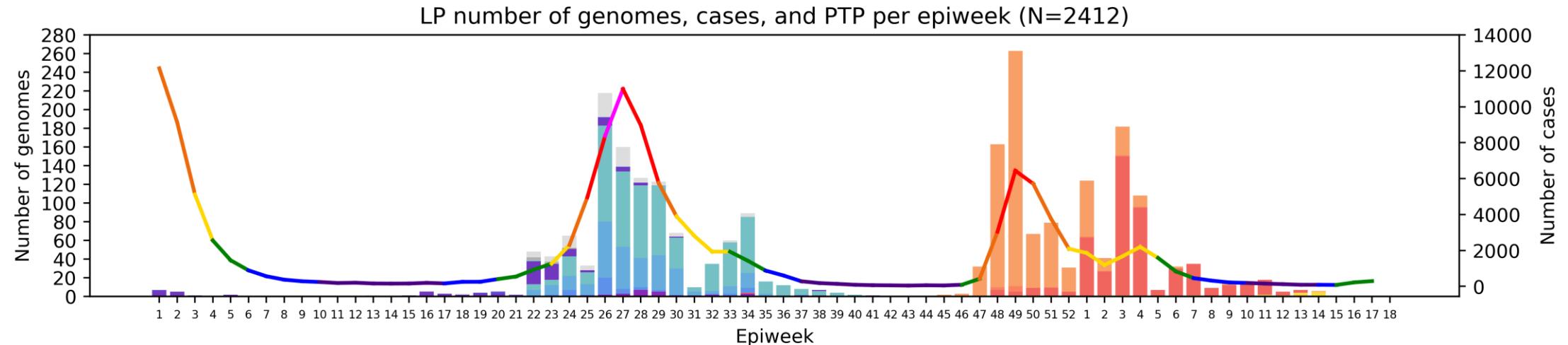
Clade key (bar graph)

Omicron (21K/BA.1.*)	Omicron (21M/BA.3)	Omicron (22B/BA.5)	Beta (20H, V2)	Delta (21A)	Delta (21J)	Kappa (21B)	Eta (21D)	20D	20C	20B	20A	19B	19A	unassigned
Omicron (21L/BA.2.*)	Omicron (22A/BA.4)	C.1.2 (20D)	Alpha (20I, V1)	Delta (21I)	—	—	—	20D	20C	20B	20A	19B	19A	—

Weekly percentage testing positive key (line graph)

≤ 5	6 - 10	11 - 20	21 - 30	31 - 40	41 - 50	51 - 55
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# Limpopo Province, 2021-2022, n = 2412



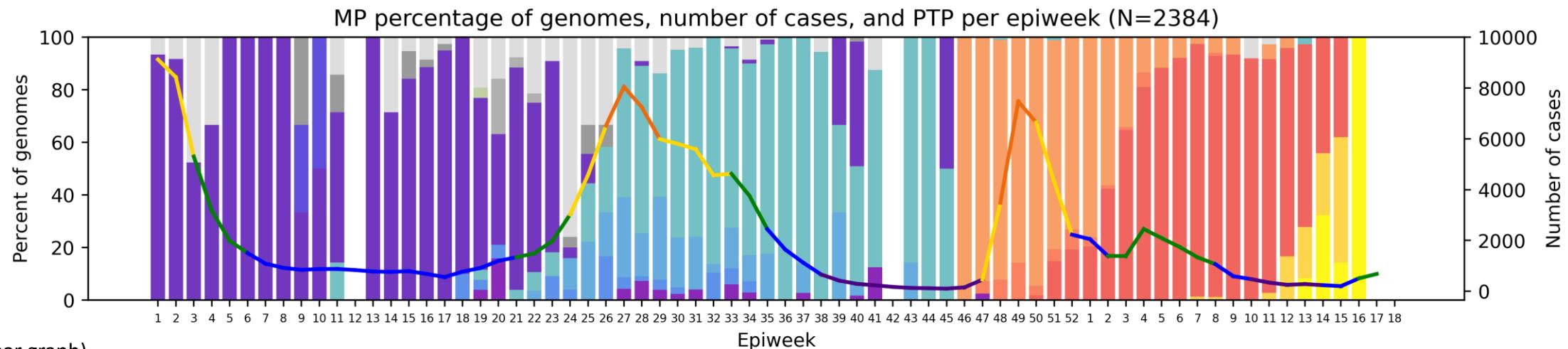
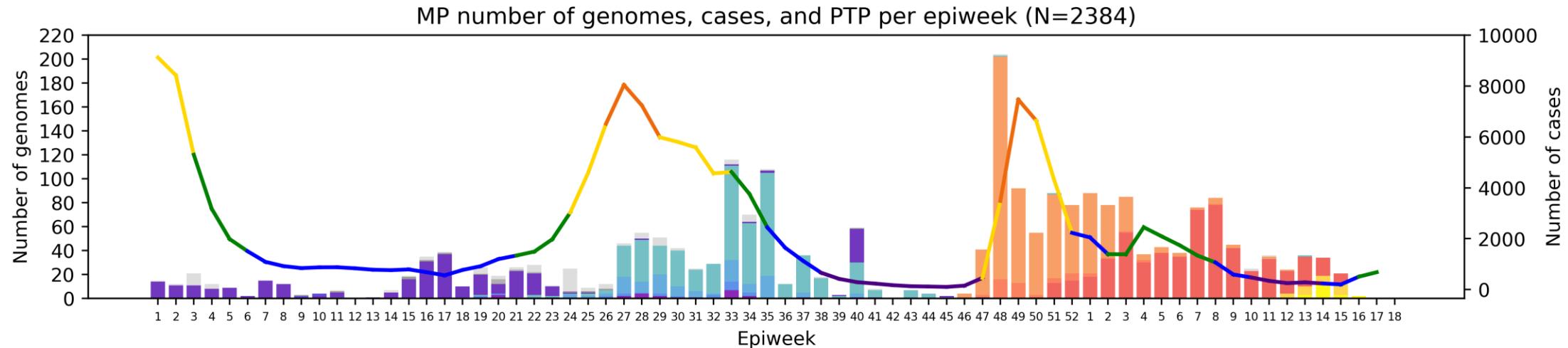
Clade key (bar graph)

Omicron (21K/BA.1.*)	Omicron (21M/BA.3)	Omicron (22B/BA.5)	Beta (20H, V2)	Delta (21A)	Delta (21J)	Kappa (21B)	Eta (21D)	20D	20C	20B	20A	19B	19A	unassigned
Omicron (21L/BA.2.*)	Omicron (22A/BA.4)	C.1.2 (20D)	Alpha (20I, V1)	Delta (21I)	—	—	—	—	—	—	—	—	—	—

Weekly percentage testing positive key (line graph)

≤ 5	6 - 10	11 - 20	21 - 30	31 - 40	41 - 50	51 - 55
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# Mpumalanga Province, 2021-2022, n = 2384



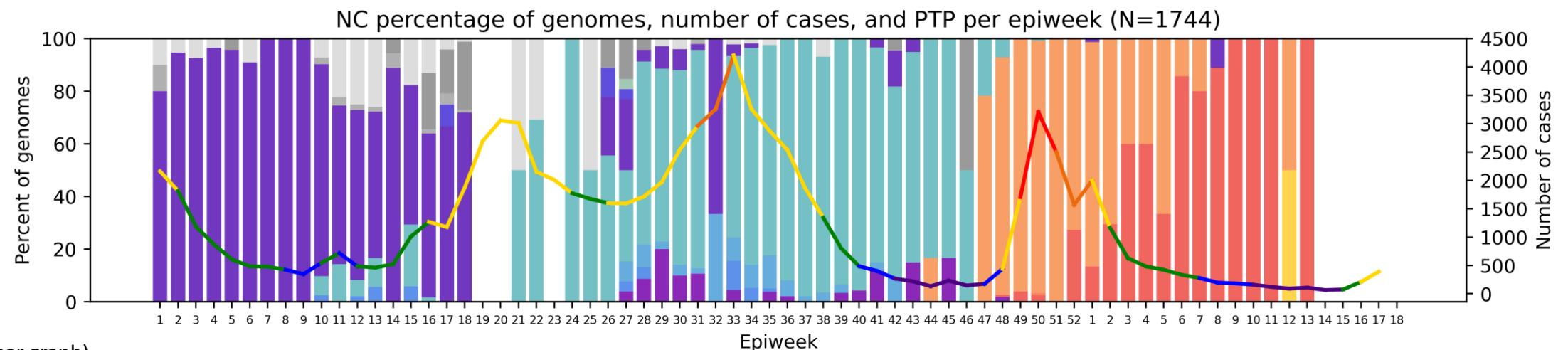
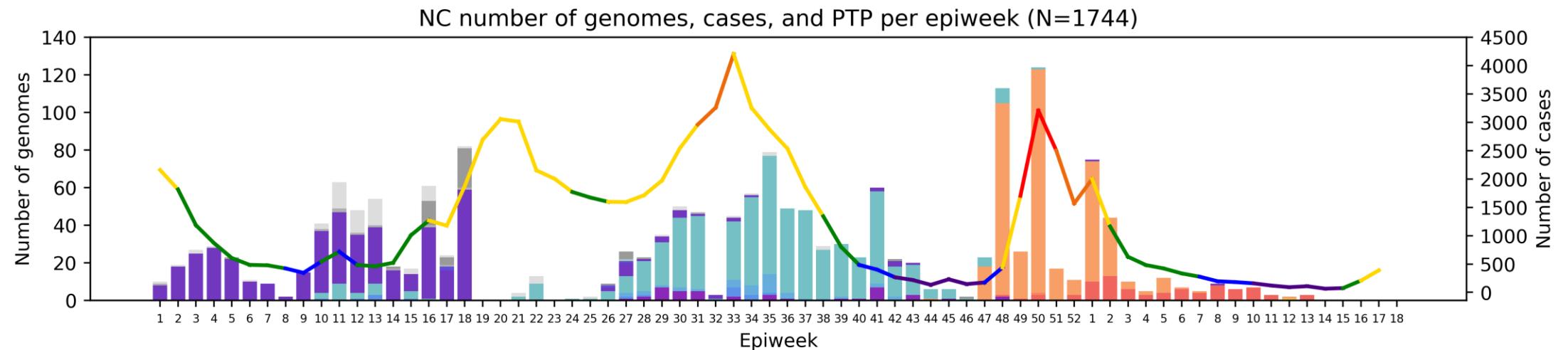
Clade key (bar graph)

Omicron (21K/BA.1.*)	Omicron (21M/BA.3)	Omicron (22B/BA.5)	Beta (20H, V2)	Delta (21A)	Delta (21J)	Kappa (21B)	Eta (21D)	20D	20C	20B	20A	19B	19A	unassigned
Omicron (21L/BA.2.*)	Omicron (22A/BA.4)	C.1.2 (20D)	Alpha (20I, V1)	Delta (21I)	Delta (21)			20D	20C	20B	20A	19B	19A	

Weekly percentage testing positive key (line graph)

≤ 5	6 - 10	11 - 20	21 - 30	31 - 40	41 - 50	51 - 55
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# Northern Cape Province, 2021-2022, n = 1744



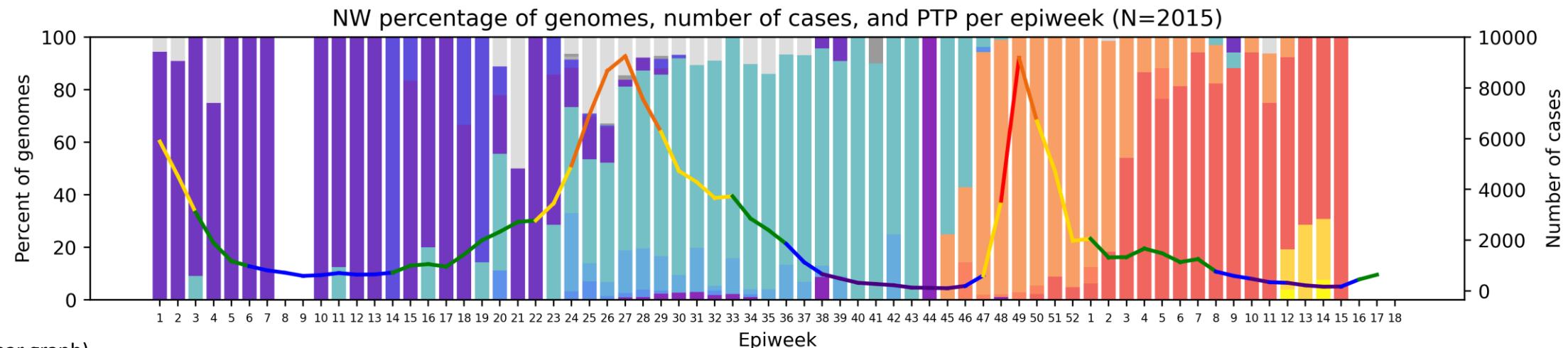
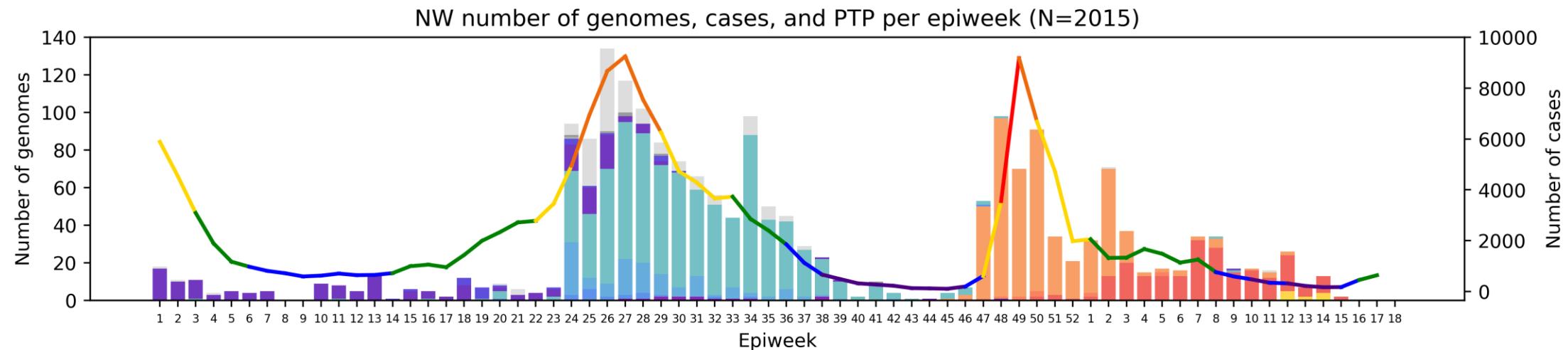
Clade key (bar graph)

Omicron (21K/BA.1.*)	Omicron (21M/BA.3)	Omicron (22B/BA.5)	Beta (20H, V2)	Delta (21A)	Delta (21J)	Kappa (21B)	Eta (21D)	20D	20C	20B	20A	19B	19A	unassigned
Omicron (21L/BA.2.*)	Omicron (22A/BA.4)	C.1.2 (20D)	Alpha (20I, V1)	Delta (21I)	Delta (21)			20D	20C	20B	20A	19B	19A	

Weekly percentage testing positive key (line graph)

≤ 5	6 - 10	11 - 20	21 - 30	31 - 40	41 - 50	51 - 55
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# North West Province, 2021-2022, n = 2015



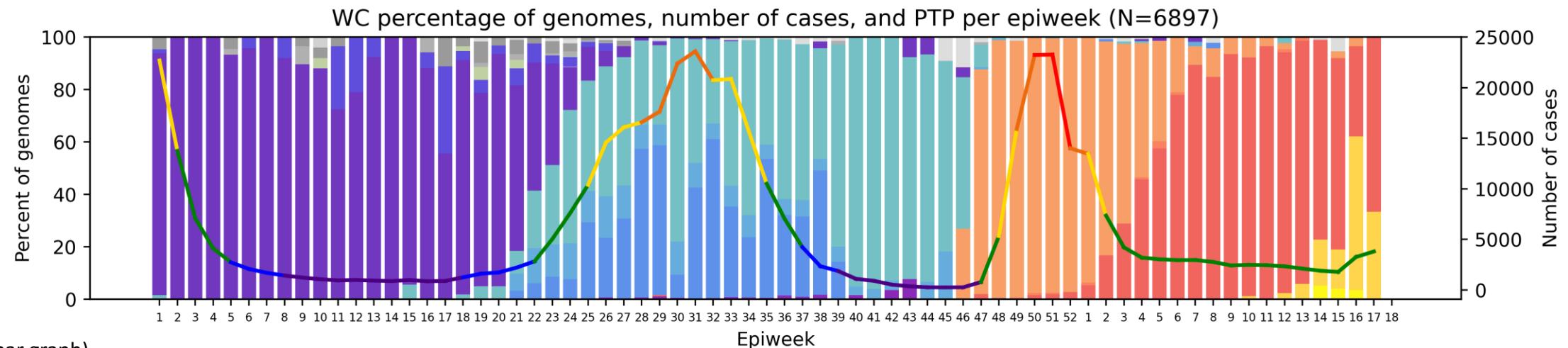
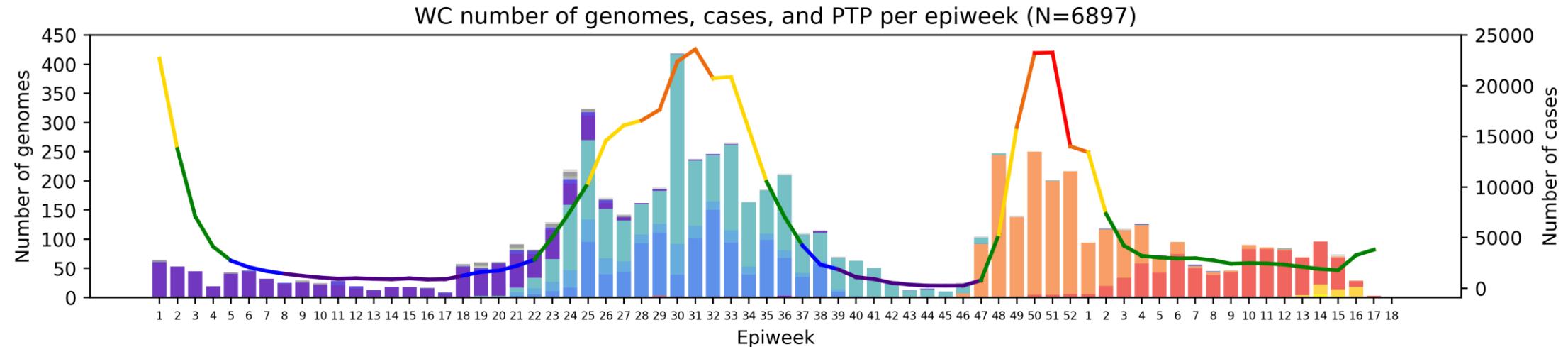
Clade key (bar graph)

Omicron (21K/BA.1.*)	Omicron (21M/BA.3)	Omicron (22B/BA.5)	Beta (20H, V2)	Delta (21A)	Delta (21J)	Kappa (21B)	Eta (21D)	20D	20C	20B	20A	19B	19A	unassigned
Omicron (21L/BA.2.*)	Omicron (22A/BA.4)	C.1.2 (20D)	Alpha (20I, V1)	Delta (21I)	—	—	—	—	—	—	—	—	—	—

Weekly percentage testing positive key (line graph)

≤ 5	6 - 10	11 - 20	21 - 30	31 - 40	41 - 50	51 - 55
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# Western Cape Province, 2021-2022, n = 6897



Clade key (bar graph)

Omicron (21K/BA.1.*)	Omicron (21L/BA.2.*)	Omicron (22B/BA.5)	Beta (20H, V2)	Delta (21A)	Delta (21J)	Kappa (21B)	Eta (21D)	20D	20C	20B	20A	19B	19A	unassigned
Omicron (21L/BA.2.*)	Omicron (22A/BA.4)	C.1.2 (20D)	Alpha (20I, V1)	Delta (21)	Delta (21)									

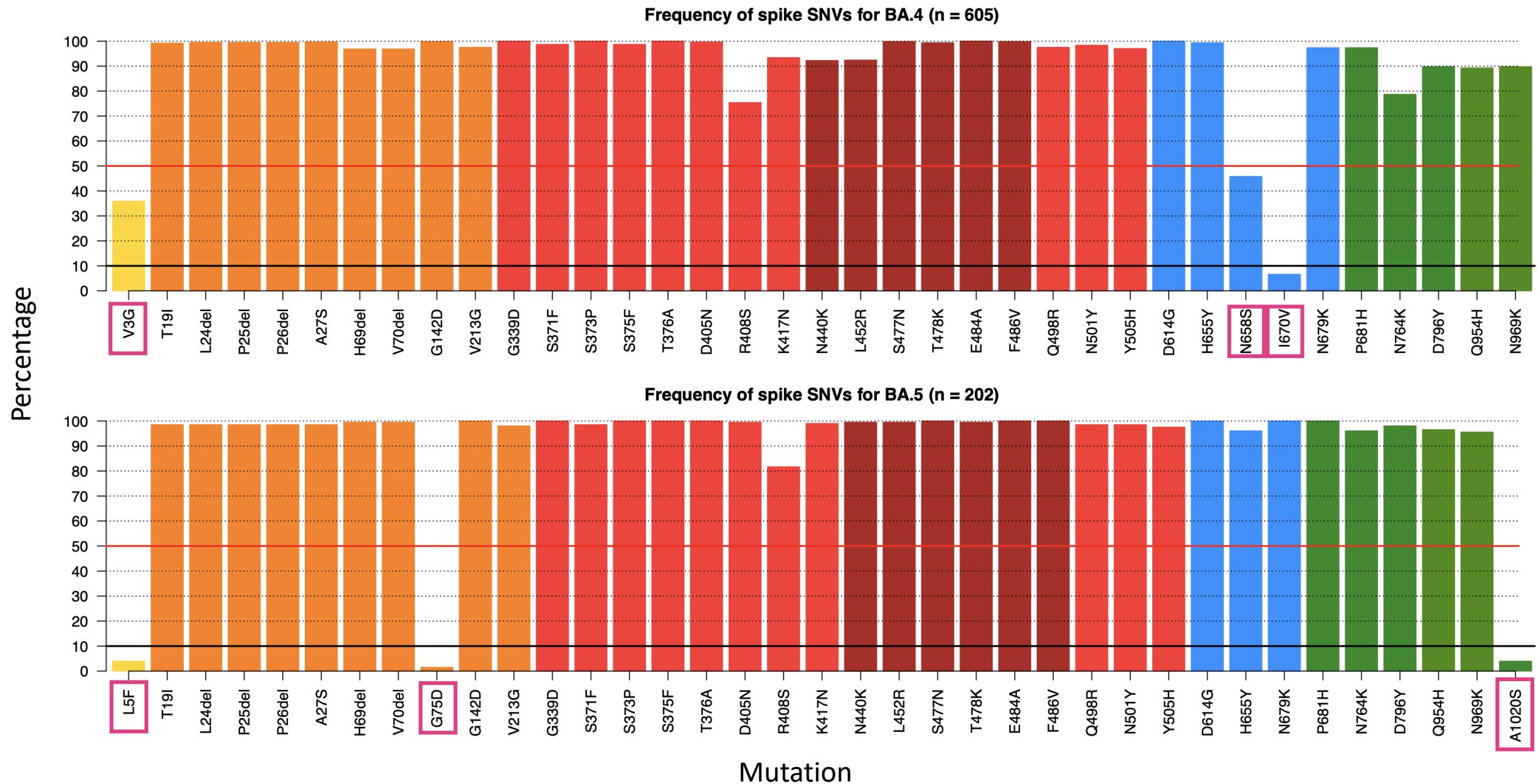
Weekly percentage testing positive key (line graph)

≤ 5	6 - 10	11 - 20	21 - 30	31 - 40	41 - 50	51 - 55
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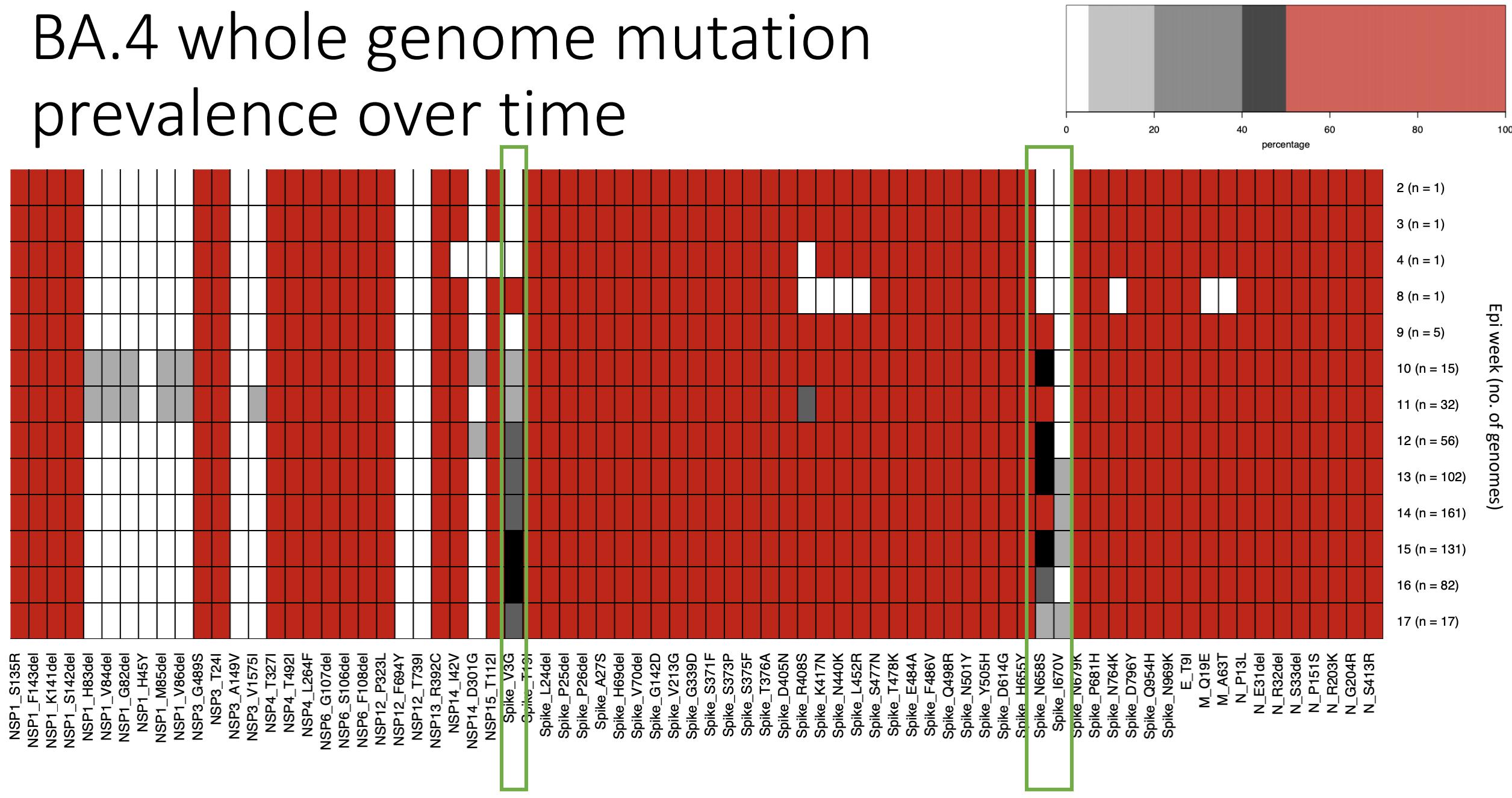
# Summary

- **Variant of Concern Omicron in South Africa**
  - Dominates 2022 sequencing data at >98% of genomes.
  - While BA.1 (and sub-lineages) was the predominant lineage in January (55%), BA.2 dominated in February (86%) and March (78%).
  - Omicron lineages BA.4 and BA.5 increased in prevalence in March (16%), and together are dominant in April (64%).
  - BA.3 continues to be detected at low levels.
  - NGS-SA teams are monitoring sequencing data for recombinants.
- Low frequency of previously circulating variants such as Delta still detected in recent data.

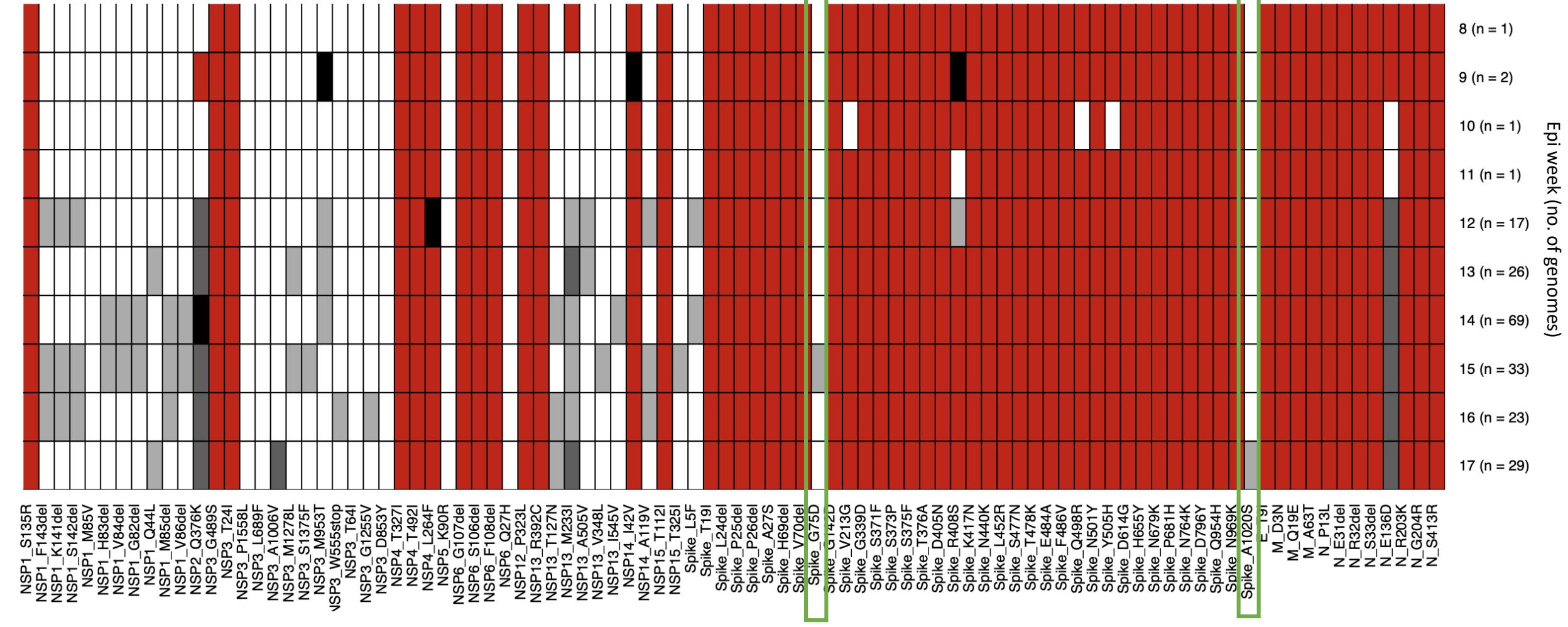
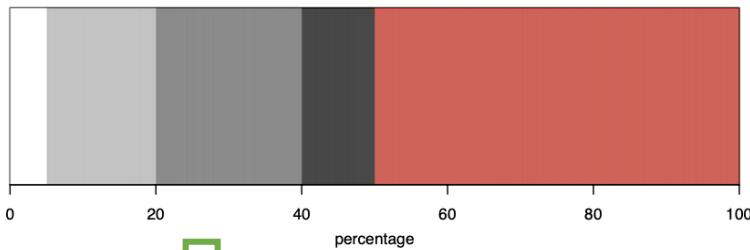
# BA.4 and BA.5 spike mutations



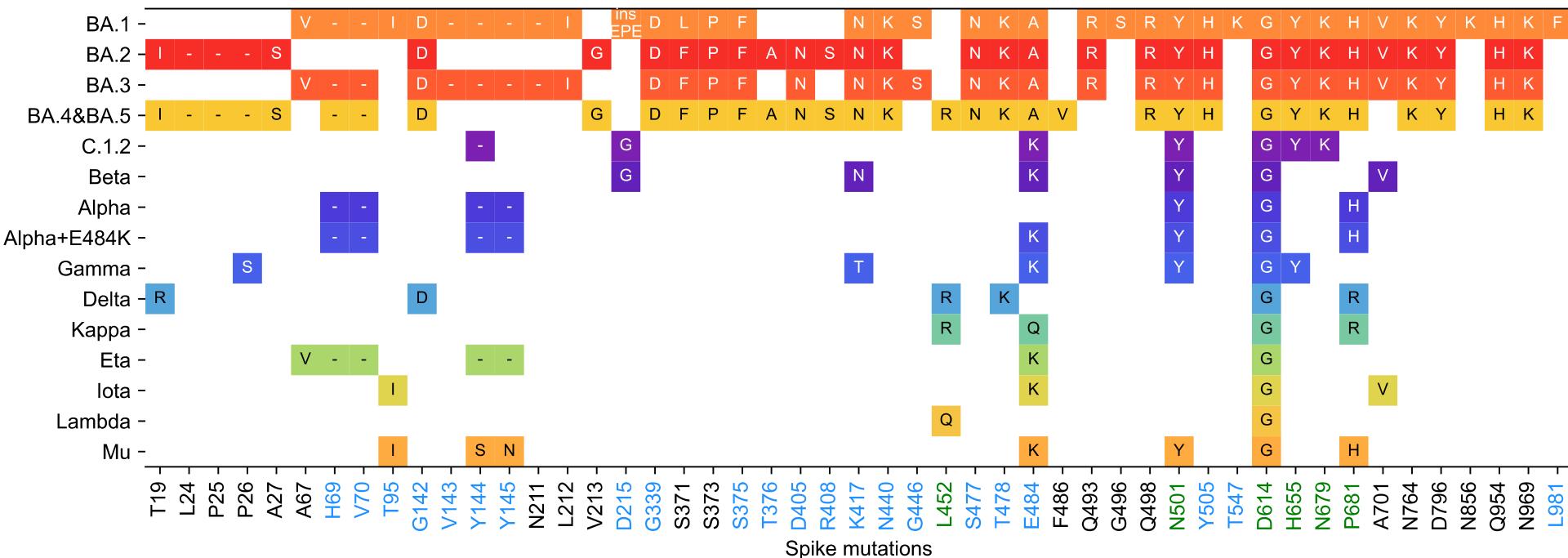
# BA.4 whole genome mutation prevalence over time



# BA.5 whole genome mutation prevalence over time



# Omicron spike mutations compared to other VOC/VOIs

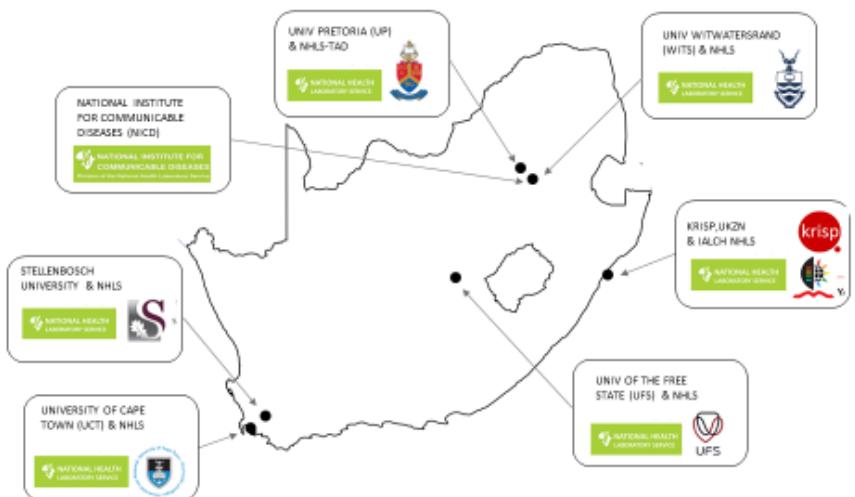


Only lineage-defining mutations are pictured here. Low prevalence mutations can be seen on the following slide.

## Mutation impact key

<span style="background-color: black; border: 1px solid black; display: inline-block; width: 15px; height: 15px;"></span>	Unknown or unconfirmed impact
<span style="background-color: blue; border: 1px solid black; display: inline-block; width: 15px; height: 15px;"></span>	Known/predicted immune escape
<span style="background-color: green; border: 1px solid black; display: inline-block; width: 15px; height: 15px;"></span>	Enhanced infectivity

- Multiple changes within the two immunogenic regions in S1 (NTD and RBD)
  - including a three amino acid insertion
- Accumulation of mutations surrounding the furin cleavage site
  - Including combination of N679K and P681H
- Effect of most spike S2 subunit changes have not been defined, but may be linked to immune escape



Supported by the DSI and the SA MRC



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UNIVERSITY



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ED C T P  
This project (RIA2020EF-3030) is part of the EDCTP2 programme supported by the European Union"



## University of Stellenbosch & NHLS Tygerberg Virology



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Wolfgang Preiser  
Gert van Zyl  
Tongai Maponga  
Bronwyn Kleinhans  
Shannon Wilson  
Karabo Phadu  
Tania Stander  
Kamela Mahlakwane  
Mathilda Claassen  
Diagnostic laboratory staff

## University of Cape Town, NHLS & Western Cape Government



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Nei-yuan Hsiao  
Diana Hardie  
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Ziyaad Valley-Omar

### WCG-UCT

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Hannah Hussey  
Andrew Boule  
Masudah Paleker  
Theuns Jacobs  
Erna Morden

### NHLS Greenpoint

Annabel Enoch



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Innocent Mudau  
Nokuzola Mbhele  
Fezokuhle Khumalo  
Thabang Serake  
Bruna Galvão  
Arghavan Alisoltani (U. California)  
Robert Wilkinson  
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Wendy Burgers  
Ntobeko Ntusi  
Rageema Joseph  
Sean Wasserman  
Linda Boloko



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Amy Strydom (Postdoc)  
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**NHLS Tshwane**  
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**HYRAX**  
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CAPE TOWN HVTN  
IMMUNOLOGY LABORATORY

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**AMPATH**  
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**PathCare**  
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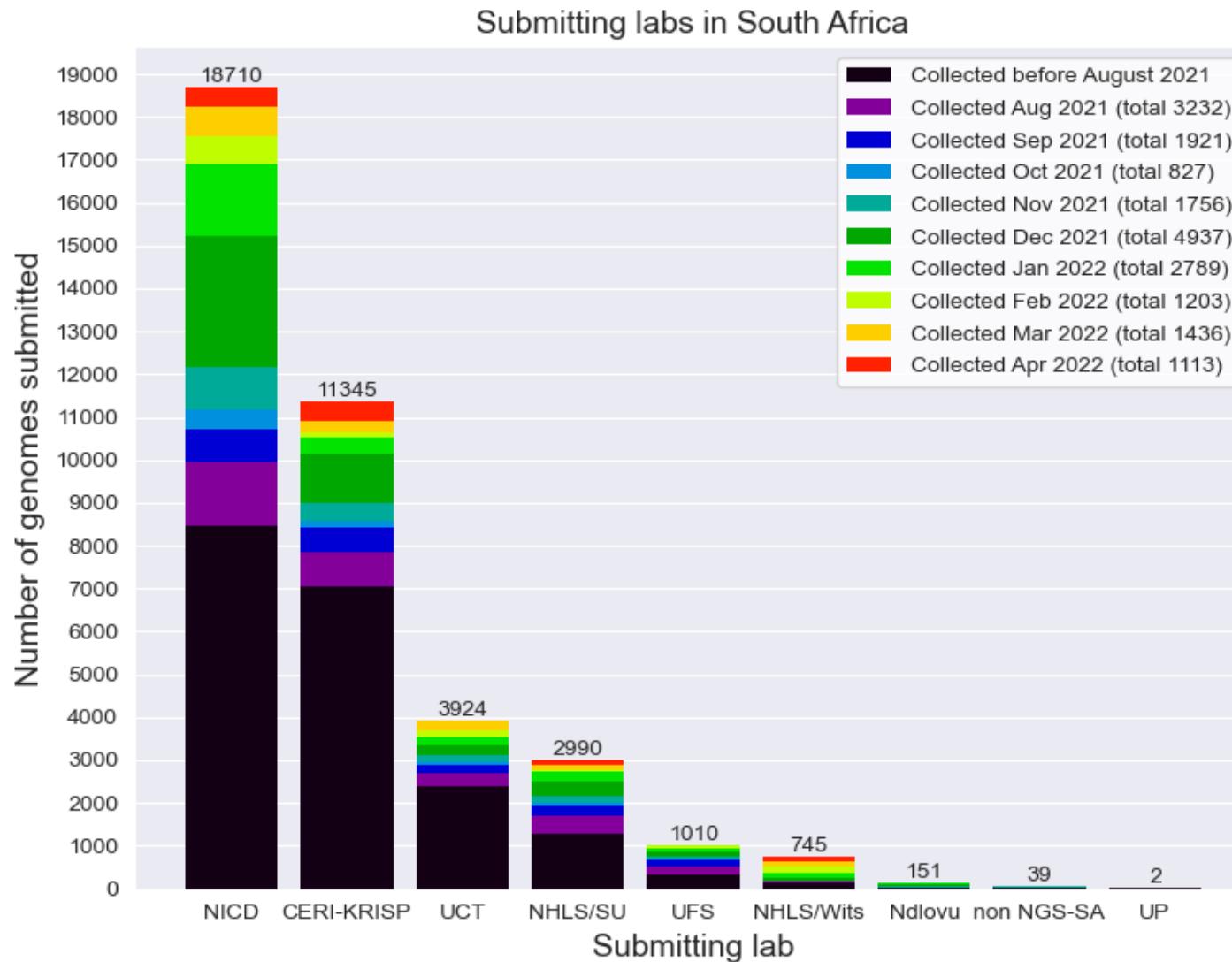
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# South African genomes submitted per submitting lab, 2020 - 2022 (N=38 916)



## NGS-SA Labs

**CERI:** Centre for Epidemic Response and Innovation

**KRISP:** KZN Research Innovation and Sequencing Platform

**NDLOVU:** Ndlovu Research Laboratories

**NICD:** National Institute for Communicable Diseases

**NHLS:** National Health Laboratory Service

**SU:** Stellenbosch University

**UCT:** University of Cape Town

**UFS:** University of the Free State

**UP:** University of Pretoria

Multiple labs from NGS-SA and collaborating public and private laboratories are contributing to sequencing, both as originating and as submitting (pictured here) laboratories.

# Currently circulating Variants of Concern (VOC)

WHO label	Pango lineage•	GISAID clade	Nextstrain clade	Additional amino acid changes monitored°	Earliest documented samples	Date of designation
Delta	B.1.617.2	G/478K.V1	21A, 21I, 21J	+S:K417N +S:K484K	India, Oct-2020	VOI: 4-Apr-2021 VOC: 11-May-2021
Omicron*	B.1.1.529	GR/484A	21K	+S:R346K	Multiple countries, Nov-2021	VUM: 24-Nov-2021 VOC: 26-Nov-2021

<https://www.who.int/en/activities/tracking-SARS-CoV-2-variants/> accessed 18 March 2022

•Includes all descendant lineages. See the cov-lineages.org and the Pango network websites for further details.

° Only found in a subset of sequences

# Previously circulating Variants of Concern

WHO label	Pango lineage•	GISAID clade	Nextstrain clade	Earliest documented samples	Date of designation
Alpha	B.1.1.7	GRY	20I (V1)	United Kingdom, Sep-2020	VOC: 18-Dec-2020 Previous VOC: 09-Mar-2022
Beta	B.1.351	GH/501Y.V2	20H (V2)	South Africa, May-2020	VOC: 18-Dec-2020 Previous VOC: 09-Mar-2022
Gamma	P.1	GR/501Y.V3	20J (V3)	Brazil, Nov-2020	VOC: 11-Jan-2021 Previous VOC: 09-Mar-2022

<https://www.who.int/en/activities/tracking-SARS-CoV-2-variants/> accessed 18 March 2022

- Includes all descendant lineages. See the cov-lineages.org and the Pango network websites for further details.

# Submission of routine specimens for sequencing

- representative of multiple geographic regions (provinces/districts/health facilities) from individuals of
  - all ages
  - over as many time periods during the SARS-CoV-2 epidemic in South Africa
- requested that testing laboratories in both the private and public sectors, submit respiratory samples to their closest NGS-SA sequencing laboratory on a routine basis (ideally every week) as follows, depending on the capacity of the testing laboratory:
  - All positives samples should be sent every week (NGS-SA laboratory will perform random sampling as described below) **OR**
  - A weekly selection of approximately 10%-20% of randomly selected positive samples should be sent every week. Number of selected samples will depend on the size of laboratory and how many other laboratories are drained by the submitting laboratory.

# Submission of special interest specimens for sequencing

In addition to routine samples mentioned above, please send specimens separately to above and clearly marked if:

- Suspected vaccine breakthrough ( $\geq 14$  days after vaccine), especially if hospitalised and clinically severe
- Suspected re-infection ( $\geq 90$  days after previous episode), especially if hospitalised and clinically severe
- Prolonged shedding with high SARS-CoV-2 viral loads (i.e. Ct values less than 30 for more than 1 month post-primary diagnosis) in immunocompromised individuals
- Possible animal-to-human transmission
- Suspected cases of importation from another country, especially countries known to harbour SARS-CoV-2 variants of concern or countries with little available information
- Clusters of “unusual” cases (e.g., in terms of disease presentation, patient groups affected, etc.)