WEEKLY RESPIRATORY PATHOGENS SURVEILLANCE REPORT

SOUTH AFRICA WEEK 47 2021

COMMUNICABLE DISEASES

NATIONAL INSTITUTE FOR

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CUMULATIVE DATA FROM





HIGHLIGHTS: WEEK 47

-M

The 2021 influenza season has not yet started although sustained detections of influenza continue in all surveillance programmes. In week 47, transmission is below threshold and impact is low.

• 19 new cases of influenza from Western Cape (n=11), North West (n=3) and KwaZulu-Natal (n=5) surveillance sites were detected in week 47. To date, 346 influenza cases have been detected from Gauteng, Western Cape, North West, Eastern Cape, Mpumalanga and KwaZulu-Natal sentinel surveillance sites. From 1 January 2021 to date, influenza A(H1N1)pdm09 was the most commonly detected subtype in both influenza-like illness (ILI) surveillance (n=51/130, 39%) and pneumonia surveillance (n=55/168, 33%).

RSV activity remains below seasonal threshold in both ILI and pneumonia surveillance programmes. From 1 January 2021 to date, RSV subgroup A was the most commonly detected subgroup in both ILI surveillance (n=42/74, 57%) and pneumonia surveillance (n=216/411, 53%).

From 2 March 2020 to date, a total of 2 932 COVID-19 cases were detected from all surveillance programmes. A slight increase in number of cases in both pneumonia surveillance (from week 46) and ILI (in week 47) was noted. Of the 2 054 hospitalised COVID-19 cases reported with available data on outcome, 355 (17%) died.

From 1 January 2021 to date, of the 1211/1300 (93%) SARS-CoV-2 positive cases with variant type results, Delta (422/871, 48%) and Beta (157/319, 49%), were the most detected variants in pneumonia surveillance and in ILI, respectively. Delta variant predominated in both programmes (from week 22, week starting 31st May 2021 until week38). Beta variant predominated from week 47 of 2020 to week 21 of 2021.

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PROGRAMME DESCRIPTIONS

Programme	Influenza-like illness (ILI)	Viral Watch	National syndromic surveillance for pneumonia
Start year	2012	1984	2009
Provinces*	KZ NW WC** MP***	EC FS GP LP MP NC NW WC	GP KZ MP NW WC
Type of site	Primary health care clinics	General practitioners	Public hospitals
Case definition	ILI: An acute respiratory illness with a temperature (≥38°C) and cough, & onset ≤10 days Suspected pertussis Any person with an acute cough illness lasting ≥14 days (or cough illness of any duration for children <1 year), without a more likely diagnosis AND one or more of the following signs or symptoms: • paroxysms of coughing, • or inspiratory "whoop", • or post-tussive vomiting • or apnoea in children <1 year; OR Any person in whom a clinician suspects pertussis	ILI: An acute respiratory illness with a temperature (≥38°C) and cough, & onset ≤10 days	SRI: Acute (symptom onset≤10 days) or chronic (symptom onset >10) lower respiratory tract infection Suspected pertussis Any person with an acute cough illness lasting ≥14 days (or cough illness of any duration for children <1 year), without a more likely diagnosis AND one or more of the following signs or symptoms: paroxysms of coughing, or inspiratory "whoop", or post-tussive vomiting or apnoea in children <1 year; OR Any person in whom a clinician suspects pertussis.
	Suspected SARS-CoV-2 Any person presenting with an acute (≤14 days) respiratory tract infection or other clinical illness compatible with COVID-19 ^β	Suspected SARS-CoV-2 Any person presenting with an acute (<14 days) respiratory tract infection or other clinical illness compatible with COVID-19 ^g	Suspected SARS-CoV-2 Any person admitted with a physician diagnosis of suspected COVID-19 and not meeting SRI case definition.
Specimens collected	Oropharyngeal & nasopharyngeal swabs	Throat and/or nasal swabs or Nasopharyngeal swabs	Oropharyngeal & nasopharyngeal swabs
Main pathogens tested****	INF RSV BP SARS-CoV-2	INF RSV BP SARS-CoV-2	INF RSV BP SARS-CoV-2
Testing Methods	 INF and RSV Fast-Track Diagnostics multiplex real-time reverse transcription polymerase chain reaction (until 31 March 2021) B. pertussis Multiplex real-time PCR (Tatti et al., J Clin Microbiol 2011) and culture (if PCR cycle threshold ≤25) SARS-CoV-2 1 April 2020 – 31 March 2021: Roche E gene real-time PCR essay (Corman 	INF and RSV - Fast-Track Diagnostics multiplex real-time reverse transcription polymerase chain reaction (until 31 March 2021) B. pertussis Multiplex real-time PCR (Tatti et al., J Clin Microbiol 2011) and culture (if PCR cycle threshold ≤25) SARS-CoV-2 1 April 2020 – 31 March 2021: Roche E gene real-time PCR essay	INF and RSV - Fast Track Diagnostics multiplex real time reverse transcription polymerase chain reaction (until 31 March 2021) B. pertussis Multiplex real-time PCR (Tatti et al., J Clin Microbiol 2011) and culture (if PCR cycle threshold ≤25) SARS-CoV-2 1 April 2020 – 31 March 2021: Roche E gene real-time PCR essay (Corman et al., Euro Surv 2020)
	et al., Euro Surv 2020) 1 April 2021 to date: Allplex [™] SARS- CoV-2/FluA/FluB/RSV PCR kit • positivity assigned if PCR cycle threshold is <40 for ≥1 gene targets (N, S, OR RdRp)	Corman et al., Euro Surv 2020) 1 April 2021 to date: Allplex [™] SARS- CoV-2/FluA/FluB/RSV PCR kit • positivity assigned if PCR cycle threshold is <40 for ≥1 gene targets (N, S, OR RdRp)	1 April 2021 to date: Allplex™ SARS- CoV-2/FluA/FluB/RSV PCR kit • positivity assigned if PCR cycle threshold is <40 for ≥1 gene targets (N, S, OR RdRp)

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Epidemic Threshold

Thresholds are calculated using the Moving Epidemic Method (MEM), a sequential analysis using the R Language, available from: http://CRAN.R-project.org/ web/package=mem) designed to calculate the duration, start and end of the annual influenza epidemic. MEM uses the 40th, 90th and 97.5th percentiles established from available years of historical data to calculate thresholds of activity. Thresholds of activity for influenza and RSV are defined as follows: Below seasonal threshold, Low activity, Moderate activity, High activity, Very high activity. For influenza, thresholds from outpatient influenza like illness (Viral Watch Programme) are used as an indicator of disease transmission in the community and thresholds from pneumonia surveillance are used as an indicator of impact of disease.

* EC: Eastern Cape; FS: Free State; GP: Gauteng; KZ: KwaZulu-Natal; LP: Limpopo; MP: Mpumalanga: NC: Northern Cape; NW: North West; WC: Western Cape **Started in 2019

***Started in November 2020

****INF: influenza virus; RSV: respiratory syncytial virus; BP: Bordetella pertussis; SARS-CoV-2: severe acute respiratory syndrome coronavirus 2

PSymptoms include ANY of the following respiratory symptoms: cough, sore throat, shortness of breath, anosmia (loss of sense of smell) or dysgeusia (alteration of the sense of taste), with or without other symptoms (which may include fever, weakness, myalgia, or diarrhoea). Testing for SARS-CoV-2 was initiated in all three surveillance programmes in week 10 of 2020 (week starting 2 March 2020).

COMMENTS

Influenza

The 2021 influenza season has not yet started although sustained detections of influenza continue in all surveillance programmes. Since the first influenza positive case of 2021 was detected in pneumonia surveillance in week 9 of 2021 (week ending on the 07 March 2021), sporadic cases have been reported from week 16 to date. Of the 346 influenza cases detected in surveillance sites in 2021, the majority (n=118, 34%) were influenza A(H1N1)pdm09. In week 47, transmission is below threshold and impact is low.

ILI programme: In 2021 to date, specimens from 1 771 patients meeting ILI case definition were received from 4 ILI sites. Influenza was detected in 130 (7%) patients, of which 51 (39%) were influenza A(H1N1)pdm09, 19 (15%) influenza A(H3N2), three (2%) influenza A(inconclusive), 15 (12%) influenza A(pending subtype results), 38 (29%) influenza B(Victoria) and four (3%) influenza B(lineage inconclusive).

Viral Watch programme: In 2021 to date, specimens were received from 215 patients from Viral Watch sites in 6 of the 8 provinces participating in surveillance. Influenza was detected in 26 (12%) patients, of which eight (31%) were influenza A(H1N1)pdm09, five (19%) influenza A(H3N2), two (8%) influenza A(pending results), seven (27%) influenza B(Victoria), three (12%) influenza B(lineage inconclusive) and one (4%) influenza B(lineage pending results). (Fig7, Table5)

Pneumonia surveillance: Since the beginning of 2021, specimens from 5 716 patients with severe respiratory illness (SRI) were received from the 6 sentinel sites. Influenza was detected in 168 (3%) patients, of which 55 (33%) were influenza A(H1N1)pdm09, 29 (17%) influenza A(H3N2), five (3%) influenza A(subtype inconclusive), 23 (14%) influenza A(pending subtype results), 48 (29%) influenza B(Victoria), seven (4%) were influenza B(lineage inconclusive) and one (1%) influenza B(lineage pending results). (Fig12, Table9)

In addition, influenza was detected in 22 (3%) of 808 specimens, of which four (18%) were influenza A(H1N1)pdm09, 17 (77%) influenza B(Victoria) and one (5%) was influenza B(inconclusive) from patients who met suspected SARS-CoV-2 case definition but did not meet pneumonia/ILI surveillance case definition.

Respiratory syncytial virus

In 2021 to date, RSV detection has been reported from all surveillance programmes, activity remains below seasonal threshold. Of the 512 RSV cases detected in 2021, the majority (n=268, 52%) were RSV subgroup A.

ILI programme: In 2021 to date, 1 771 specimens from patients meeting the ILI case definition were tested and RSV was detected in 74 (4%) patients. Of which, 42 (57%) were RSV subgroup A, 31 (42%) RSV subgroup B and one (1%) was RSV (subgroup inconclusive). (Fig4, Table2)

Viral Watch programme: In 2021 to date, 215 specimens from viral watch patients were tested and RSV was detected in specimens of six (3%) patients. Of which, three (50%) were RSV subgroup A, two (33%) RSV subgroup B and one (17%) was RSV (subgroup inconclusive). (Fig9, Table6)

Pneumonia surveillance: Since the beginning of 2021, 5 716 specimens were tested and RSV was detected in specimens of 411 (7%) patients. Of which, 216 (53%) were RSV subgroup A, 190 (46%) RSV subgroup B and five (1%) RSV (subgroup inconclusive). (Fig14, Table10)

In addition, RSV was detected in 21 of 808 (3%) specimens, of which seven (33%) were RSV subgroup A, 13 (62%) RSV subgroup B and one (5%) was RSV(subgroup inconclusive) from patients who met suspected SARS-CoV-2 case definition but did not meet pneumonia/ ILI surveillance case definition.

SARS-CoV-2 (Severe acute respiratory syndrome coronavirus 2)

ILI programme: From March 2020 to date, 3 085 patients were tested and SARS-CoV-2 was detected in 574 (19%) patients. From 1 January 2021 to date, of the 319/338 (94%) with data on variant type, majority (157/319, 49%) were Beta variant which predominated from week1 to week 24, followed by Delta (136/319, 43%) variant which predominated from week 25 to week 36. (Fig6, Table4)

Viral Watch programme: From March 2020 to date, 494 patients presenting with ILI were tested and SARS-CoV-2 was detected in 87 (18%) patients. From 1 January 2021, of the 21/87 (24%) with data on variant type, majority were (19/21, 90%) were Delta variant which dominated from week 25 to week 30. (Fig11, Table8)

Pneumonia surveillance: From March 2020 to date, 9 731 patients with severe respiratory illness (SRI) were tested and SARS-CoV-2 was detected in 2 078 (21%) patients. From 1 January 2021 to date, of the 871/941 (93%) with data on variant type, majority were (422/871, 49%) were Delta variant which dominated from week 22 to week 38 followed by Beta (403/871, 46%) variant which dominated from week 1 to 25. (Fig17, Table12)

In addition, SARS-CoV-2 was detected in 193 of 1 011 (19%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet the pneumonia/ILI surveillance case definitions.

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INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE PRIMARY HEALTH CARE CLINICS

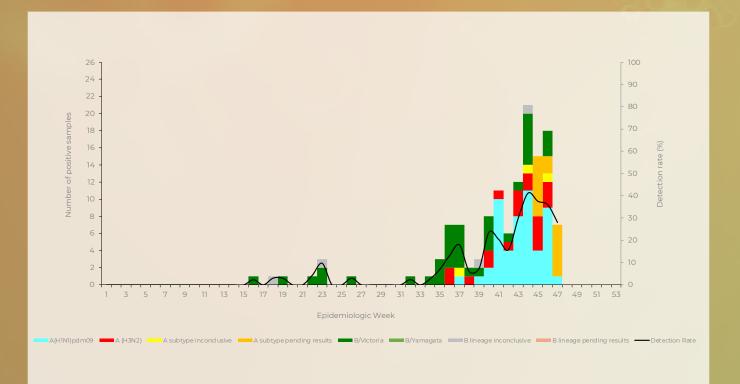


Figure 1. Number of influenza positive cases* by influenza subtype and lineage** and detection rate*** by week, Influenza-like illness (ILI) surveillance in primary health care clinics, 04/01/2021 – 28/11/2021

*Specimens from patients with influenza-like illnesses at 5 sentinel sites in 4 province

**Influenza was detected in 16 (3%) of 624 specimens, of which 2 (13%) were influenza A(H1N1)pdm09, 13 (81%) influenza B(Victoria) and one (6%) was influenza B(inconclusive) from patients who met suspected SARS-CoV-2 case definition but did not meet Influenza-like illness (ILI) case definition. These are not included in the epidemiological curve.

***Only reported for weeks with >10 specimens submit

Inconclusive: insufficient viral load in sample and unable to characterise further

Table 1. Number of laboratory confirmed influenza cases by subtype and lineage** and total number of samples tested byclinic and province, Influenza-like illness (ILI) surveillance in primary health care clinics, 04/01/2021 – 28/11/2021

Clinic (Province)	A(H1N1) pdm09	A(H3N2)	A subtype inconclusive	A subtype pending results ^ß	B/ Victoria	B/ Yamagata	B lineage inconclusive	B lineage pending results [®]	Total samples
Agincourt (MP)		2			0	0	0	0	243
Eastridge (WC)	2	6	0	2	6	0	0	О	234
Edendale Gateway (KZ)			0	5	15	Ο		0	250
Jouberton (NW)	42	5			13	0	0	Ο	777
Mitchell's Plain (WC)	5	5	1	4	4	0	1	0	267
Total:	51	19	3	15	38	0	4	0	1 771

KZ: KwaZulu-Natal; NW: North West; WC: Western Cape; MP: Mpumalanga

**Influenza was detected in 16 (3%) of 624 specimens, of which 2 (13%) were influenza A(H1N1)pdm09, 13 (81%) influenza B(Victoria) and one (6%) was influenza B(inconclusive) from patients who met suspected SARS-CoV-2 case definition but did not meet Influenza-like illness (ILI) case definition. These are not included in the table. *pinfluenza A subtype or B lineage results are pending

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INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE PRIMARY HEALTH CARE CLINICS

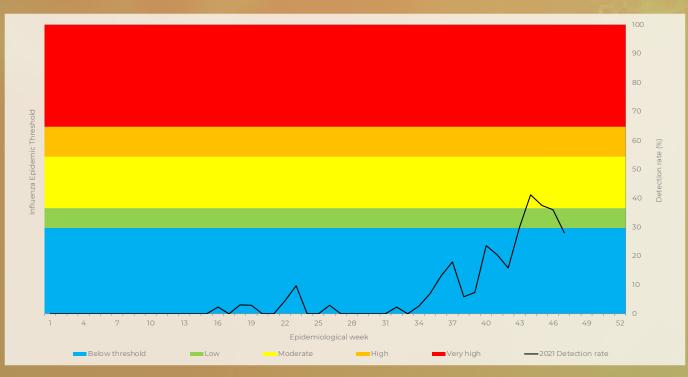


Figure 2. Influenza percentage detections and epidemic thresholds*, Influenza-like illness (ILI) surveillance in primary health care clinics, 04/01/2021 – 28/11/2021

*Thresholds based on 2012-2019 data

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INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE PRIMARY HEALTH CARE CLINICS

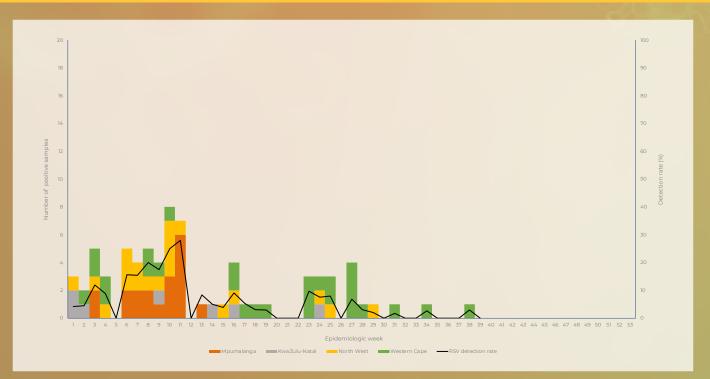


Figure 3. Number of patients testing positive for respiratory syncytial virus* by province and detection rate** by week, Influenzalike illness (ILI) surveillance in primary health care clinics, 04/01/2021 – 28/11/2021

**RSV was detected from 15 of 624 (2%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet influenza-like illness (ILI) case definition. These are not included in the epidemiological curve.

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INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE PRIMARY HEALTH CARE CLINICS

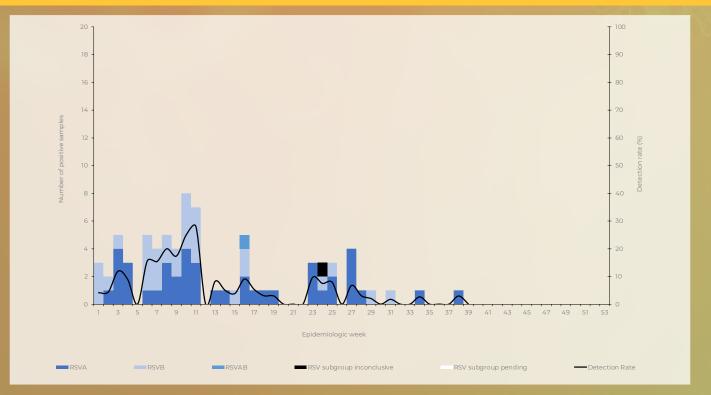


Figure 4. Number of patients testing positive for respiratory syncytial virus** by subgroup and detection rate by week,

Table 2. Number of patients testing positive for respiratory syncytial virus (RSV) by subgroups** identified and total number of samples tested by clinic and province, Influenza-like illness (ILI) surveillance in primary health care clinics, 04/01/2021 - 28/11/2021

Clinic (Province)	RSVA	RSVB	RSVAB	RSV subgroup inconclusive	RSV subgroup pending*	Total samples
Agincourt (MP)	12	7	0	0	Ο	243
Eastridge (WC)	24		0	Ο	Ο	234
Edendale Gateway (KZ)		6	0	Ο	0	250
Jouberton (NW)	2	17	0		Ο	777
Mitchell's Plain (WC)	3	0	0	О	0	267
Total	42	31	0	1	0	1 771

Inconclusive: insufficient viral load in sample and unable to characterise further RSV AB: Both RSV A and B subgroup identified *RSV results for subgroups are pending

**RSV was detected from 15 of 624 (2%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet influenza-like illness (ILI) case definition. These are not included in the table.

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INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE PRIMARY HEALTH CARE CLINICS

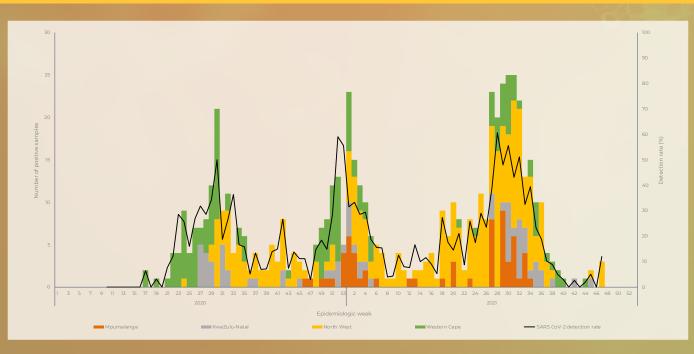


Figure 5. Number of patients testing positive for SARS-CoV-2* by province and detection rate by week, Influenza-like illness (ILI) surveillance in primary health care clinics, 02/03/2020 – 28/11/2021

*Specimens from patients with influenza-like illnesses at 5 sentinel sites in 4 provinces

**SARS-CoV-2 was detected in 149 of 749 (20%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet influenza-like illness (ILI) case definition. These are not included in the epidemiological curve.

 Table 3.
 Number of patients positive for SARS-CoV-2 identified and total number of samples tested by clinic and province,

 Influenza-like illness (ILI) surveillance primary health care clinics, 02/03/2020 – 28/11/2021

Clinic (Province)	SARS-CoV-2 positive	Total samples tested
Agincourt (MP)	62	273
Eastridge (WC)	60	723
Edendale Gateway (KZ)	70	393
Jouberton (NW)	292	1100
Mitchell's Plain (WC)	90	596
Total:	574	3 085

KZ: KwaZulu-Natal; NW: North West; WCP: Western Cape; MP: Mpumalanga (started enrolling on the 10th November 2020)

**SARS-CoV-2 was detected in 149 of 756 (19%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet influenza-like illness (ILI) case definition. These are not included in the table.

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INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE PRIMARY HEALTH CARE CLINICS

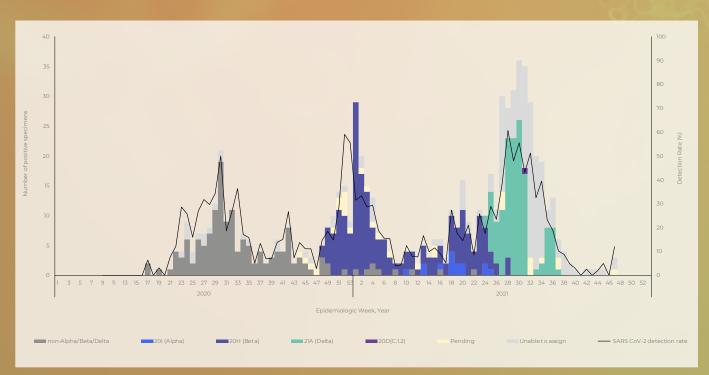


Figure 6. Number and detection rate of laboratory confirmed SARS-CoV-2* cases by variant type (variant PCR/sequencing) and week, Influenza-like illness (ILI) surveillance in primary health care clinics, 02/03/2020 – 28/11/2021

*Specimens are from patients with influenza-like illness at 5 sentinel sites in 4 provinces who met suspected SARS-CoV-2 case definition and met pneumonia (SRI) case definition as well as those that did not meet the ILI case definition.

Unable to assign: no lineage assigned due to poor- sequence quality OR low viral load (ct=>35) OR variant PCR could not assign variant and no sequencing result Pending: outstanding variant results

Table 4. Number of SARS-CoV-2* positive cases by variant (variant PCR and/or sequencing) identified and total number of samples tested by clinic and province, Influenza-like illness (ILI) surveillance primary health care clinics, 02/03/2020 – 28/11/2021

Clinic (Province)	Non-Alpha/ Beta/Delta	201 (Alpha)	20H (Beta)	21A (Delta)	20D (C.1.2)	Pending	Unable to assign	Total SARS- CoV-2 positive
Agincourt (MP)	3	0	29	17	0	7	31	87
Eastridge (WC)	20	Ο	17	7	0	3	14	61
Edendale Gateway (KZ)	27	Ο	22	31	0	3	32	115
Jouberton (NW)	67	15	104	70		17	93	367
Mitchell's Plain (WC)	35	0	27	11	0	1	19	93
Total:	152	15	199	136	1	31	189	723

KZ: KwaZulu-Natal; NW: North West; WCP: Western Cape; MP: Mpumalanga (started enrolling on the 10th November 2020

*Specimens are from patients with influenza-like illness at 5 sentinel sites in 4 provinces who met suspected SARS-CoV-2 case definition and met pneumonia (SRI) case definition as well as those that did not meet the ILI case definition.

Unable to assign: no lineage assigned due to poor- sequence quality OR low viral load (ct=>35) OR variant PCR could not assign variant and no sequencing result Pending: outstanding variant results

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INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

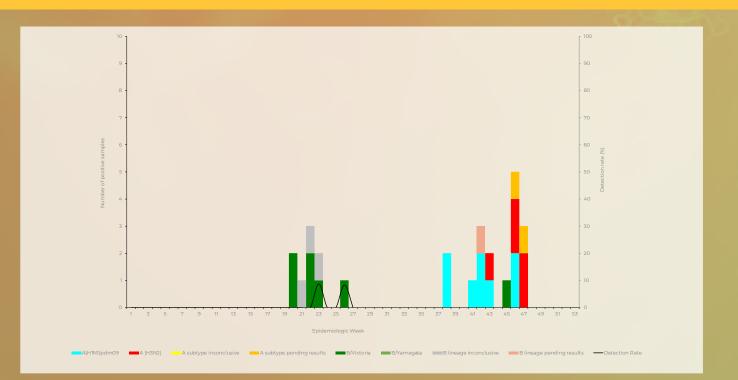


Figure 7. Number oof positive patients* by influenza subtype and lineage and detection rate** by week, ILI surveillance - Viral

Table 5. Number of laboratory confirmed influenza cases by influenza subtype and lineage and total number of samples tested by province, ILI surveillance - Viral Watch, 04/01/2021 – 28/11/2021

Province	A(H1N1) pdm09	A(H3N2)	A subtype inconclusive	A subtype pending results*	B/ Victoria	B/ Yamagata	B lineage inconclusive	B lineage pending results*	Total samples
Eastern Cape		0	0	0		0	0	0	4
Free State	0	0	0	0	Ο	О	О	0	2
Gauteng	5	2	0		5	О	3	0	163
Limpopo	Ο	0	0	0	0	О	О	0	0
Mpumalanga	0	0	0	0	0	О	О	0	3
North West	0	0	0	0	0	О	О	0	2
Northern Cape	0	0	0	0	0	О	О	0	О
Western Cape	2	3	О	1	1	0	0	1	39
Total:	8	5	0	2	7	0	3	1/	215

Inconclusive: insufficient viral load in sample and unable to characterise further *Influenza A subtype or B lineage results are pending

Patients known to have acquired influenza abroad are not included in the table or epidemiological curve.

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INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

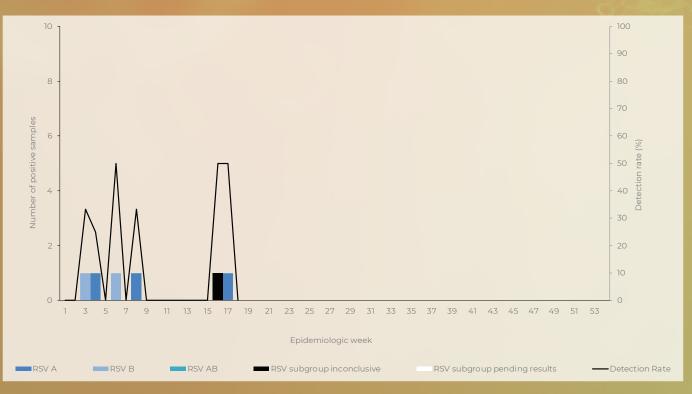


Figure 8. Number of RSV positive cases testing positive for respiratory syncytial virus (RSV)* by subgroup and detection rate by week, ILI surveillance - Viral Watch, 04/01/2021 – 28/11/2021

*Specimens from patients with Influenza-like illnesses at 92 sentinel sites in 8 provinces *RSV results for subgroups are pending

Table 6. Number of RSV positive cases identified and total number of samples tested by province, ILI surveillance - Viral Watch, 04/01/2021 – 28/11/2021

Province	RSV A	RSV B	RSV AB	RSV subgroup inconclusive**	RSV subgroup pending results*	Total samples tested
Eastern Cape	0	0	0	0	0	4
Free State	0	О	0	0	0	2
Gauteng	2	2	0	0	0	163
Limpopo	0	О	0	0	0	0
Mpumalanga	0	0	0	0	0	3
North West	0	0	0	0	0	2
Northern Cape	0	0	0	0	0	О
Western Cape	1	0	0		0	39
Total:	3	2	0	1	0	215

*RSV results for subgroups are pending

**Inconclusive: insufficient viral load in sample and unable to characterise further

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INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

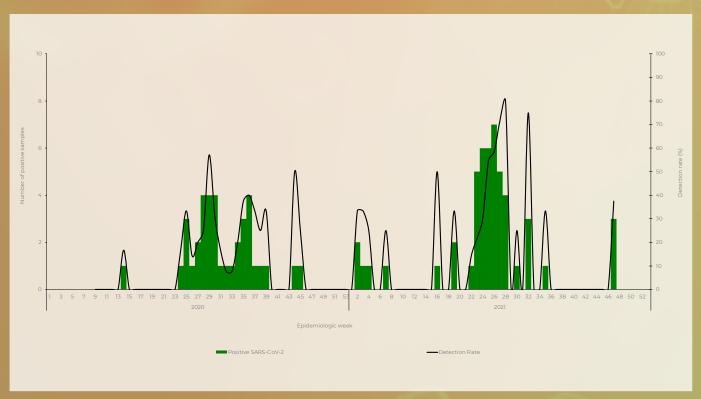


Figure 9. Number of patients testing positive for SARS-CoV-2*, by site and detection rate by week, ILI surveillance - Viral Watch, 02/03/2020 – 28/11/2021

*Specimens from patients with Influenza-like illnesses at 92 sentinel sites in 8 provinces

 Table 7. Number of SARS-CoV-2 positive cases identified and total number tested by province, ILI surveillance - Viral Watch, 02/03/2020 – 28/11/2021

Province	SARS-CoV-2 positive	Total samples tested
Eastern Cape	1	8
Free State		18
Gauteng	67	323
Limpopo	0	2
Mpumalanga		8
North West	0	2
Northern Cape	Ο	2
Western Cape	17	131
Total:	87	494

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WEEK **47** 2021

INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

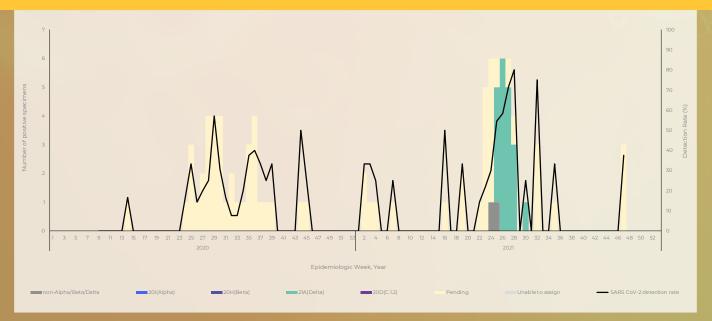


Figure 10. Number and detection rate of laboratory confirmed SARS-CoV-2* cases by variant type (variant PCR/sequencing) and week, ILI surveillance - Viral Watch, 02/03/2020 – 28/11/2021

*Specimens from patients with Influenza-like illnesses at 92 sentinel sites in 8 provinces

Unable to assign: no lineage assigned due to poor- sequence quality OR low viral load (ct=>35) OR variant PCR could not assign variant and no sequencing result Pending: outstanding variant results

Clinic (Province)	Non-Alpha/ Beta/Delta	20I (Alpha)	20H (Beta)	21A (Delta)	20D (C.1.2)	Pending	Unable to assign	Total SARS- CoV-2 positive
Eastern Cape	0	О	0	0	0		0	
Free State	Ο	О	0	0	0		0	
Gauteng	2	О	0	18	О	48	О	67
Limpopo	0	О	0	0	О	О	О	0
Mpumalanga	0	О	0	0	О		О	
North West	0	О	0	0	О	0	0	0
Northern Cape	0	О	0	0	О	0	О	0
Western Cape	0	0	0	-1	О	15	0	17
Total:	2	0	0	19	0	66	0	87

*Specimens from patients with Influenza-like illnesses at 92 sentinel sites in 8 provinces

Unable to assign: no lineage assigned due to poor- sequence quality OR low viral load (ct=>35) OR variant PCR could not assign variant and no sequencing result Pending: outstanding variant results

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NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

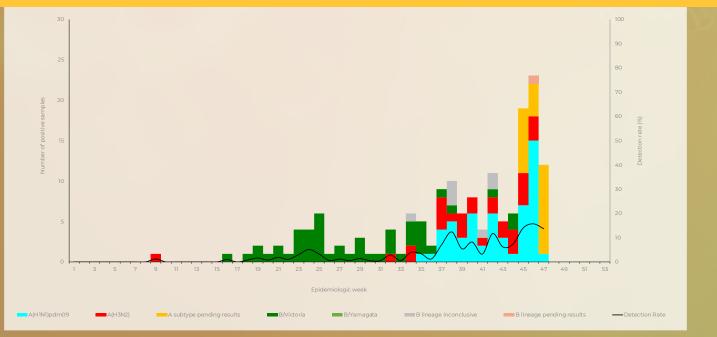


Figure 11. Number of positive influenza positive cases* by influenza subtype and lineage** and detection rate*** by week, pneumonia surveillance public hospitals, 04/01/2021 – 28/11/2021

*Specimens from patients hospitalised with pneumonia at 7 sentinel sites in 5 provinces

**Influenza was detected in six (3%) of 184 specimens, of which two (33%) were influenza A(H1N1)pdm09 and four (67%) were influenza B(Victoria) from patients who met suspected SARS-CoV-2 case definition but did not meet pneumonia (SRI) case definition. These are not included in the epidemiological curve.

***Only reported for weeks with >10 specimens submitted

Inconclusive: insufficient viral load in sample and unable to characterise further

Table 9. Number of laboratory confirmed influenza cases by subtype and lineage** and total number of samples tested by hospital, pneumonia surveillance public hospitals, 04/01/2021 – 28/11/2021

Hospital (Province)	A(H1N1) pdm09	A(H3N2)	A subtype inconclusive	A subtype pending results***	B/ Victoria	B/ Yamagata	B lineage inconclusive	B lineage pending results***	Total samples
Edendale (KZ)	٦	2	0		11	Ο	0	1	890
Helen Joseph- Rahima Moosa (GP)	23	14	2		17	0	5	0	1514
Klerksdorp- Tshepong (NW)	12			7	4	Ο	2	0	902
Mapulaneng- Matikwana (MP)	3		0	0		Ο	О	0	550
Red Cross (WC)	8	7	0	10	12	0	О	0	741
Mitchell's Plain (WC)	7	3	Ο		2	Ο	Ο	0	871
Tintswalo (MP)		12	2	3	1	0	Ο	0	248
Total:	55	29	5	23	48	0	7	1	5 716

CP: Gauteng; KZ: KwaZulu-Natal; NW: North West; MP: Mpumalanga (Tintswalo started enrolling on the 10th Feb 2021); WC: Western Cape Inconclusive: insufficient viral load in sample and unable to characterise further

**Influenza was detected in six (3%) of 184 specimens, of which two (33%) were influenza A(H1N1)pdm09 and four (67%) were influenza B(Victoria) from patients who met suspected SARS-CoV-2 case definition but did not meet pneumonia (SRI) case definition. These are not included in the table.

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NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

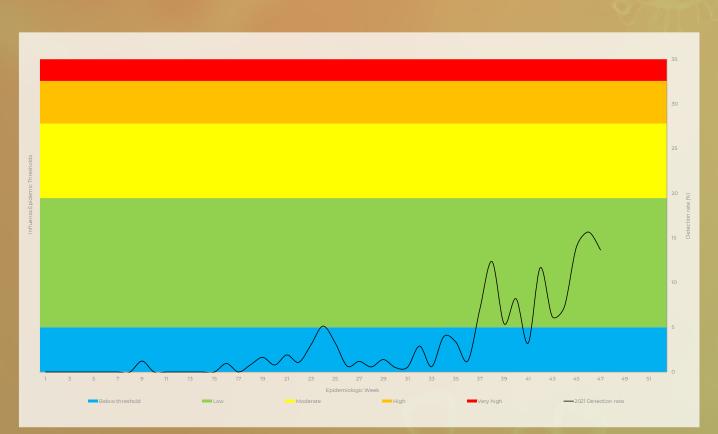


Figure 12. Influenza percentage detections and epidemic thresholds*, pneumonia surveillance public hospitals, 04/01/2021 – 28/11/2021

*Thresholds based on 2010-2019 data

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NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

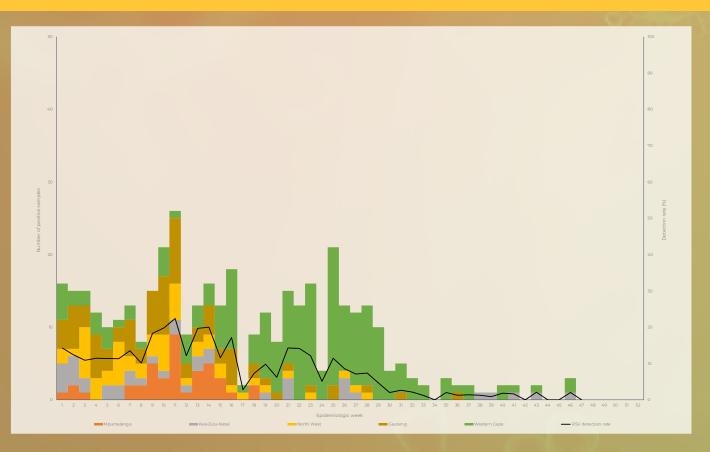


Figure 13. Number oof patients testing positive for respiratory syncytial virus* by province and detection rate by week, pneumonia surveillance public hospitals, 04/01/2021 – 28/11/2021

*RSV was detected in six of 184 (3%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet pneumonia (SRI) case definition. These are not included in the epidemiological curve.

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NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

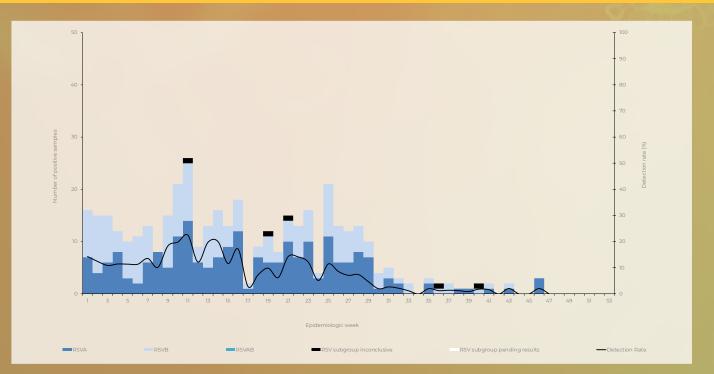


Figure 14. Number of patients testing positive for respiratory syncytial virus* by subgroup and detection rate by week, pneumonia surveillance public hospitals, 04/01/2021 – 28/11/2021

Table 10: Number of patients positive for respiratory syncytial virus subgroups** by subgroups identified and total number of

Hospital (Province)	RSVA	RSVB	RSVAB	RSV subgroup inconclusive	RSV subgroup pending*	Total samples
Edendale (KZ)	10	29	0	0	О	890
Helen Joseph-Rahima Moosa (GP)	37	42	О		Ο	1514
Klerksdorp-Tshepong (NW)	8	45	О		Ο	902
Mapulaneng-Matikwana (MP)	21	7	О	Ο	О	550
Red Cross (WC)	88	51	О	2	Ο	741
Mitchell's Plain (WC)	42	13	О		Ο	871
Tintswalo (MP)	10	3	О	О	О	248
Total:	216	190	0	5	0	5 716

GP: Gauteng; KZ: KwaZulu-Natal; NW: North West; MP: Mpumalanga (Tintswalo started enrolling on the 10th Feb 2021); WC: Western Cape Inconclusive: insufficient viral load in sample and unable to characterise further

RSV AB: Both RSV A and B subgroup identified *RSV results for subgroups are pending

**RSV was detected in six of 184 (3%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet pneumonia (SRI) case definition. These are not included in the table

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NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

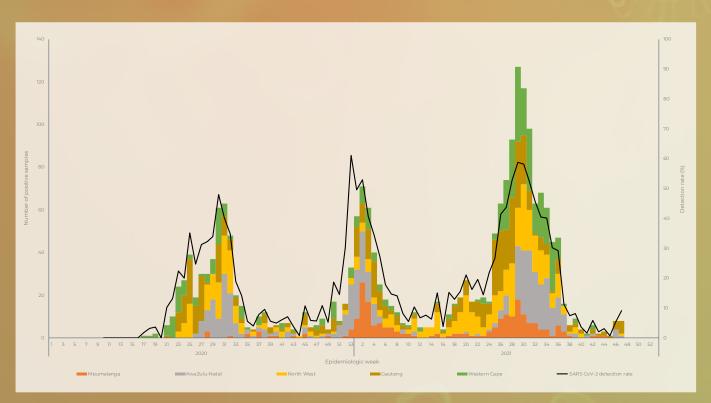


Figure 15. Number of patients testing positive for SARS-CoV-2*⁸ province and detection rate by week, pneumonia surveillance public hospitals, 02/03/2020 – 28/11/2021

*Specimens from patients hospitalized with pneumonia at 6 sentinel sites in 5 provinces

*#SARS-CoV-2 was detected in 44 of 255 (17%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet pneumonia (SRI) case definition. These are not included in the epidemiological curve.

Table 11. Number of patients positive for SARS-CoV-2** and total number of samples tested by hospital, pneumonia surveillance public hospitals, 02/03/2020 – 28/11/2021

Hospital (Province)	SARS-CoV-2 positive	Total samples tested
Edendale (KZ)	465	1 737
Helen Joseph-Rahima Moosa (GP)	504	2 360
Klerksdorp-Tshepong (NW)	477	1 495
Mapulaneng-Matikwana (MP)	167	852
Red Cross (WC)	61	1 699
Mitchell's Plain (WC)	352	1 341
Tintswalo (MP)	52	247
Total:	2 078	9 731

GP: Gauteng; KZ: KwaZulu-Natal; NW: North West; MP: Mpumalanga; WC: Western Cape

**SARS-CoV-2 was detected in 44 of 255 (17%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet pneumonia (SRI) case definition. These are not included in the table.

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NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

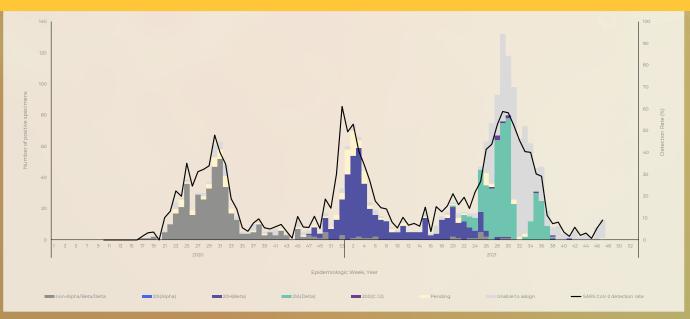


Figure 16. Number and detection rate of laboratory confirmed SARS-CoV-2 cases* by variant type (variant PCR/sequencing), pneumonia surveillance public hospitals, 02/03/2020 – 28/11/2021

Specimens are from hospitalized patients at 7 sentinel sites in 5 provinces who met suspected SARS-CoV-2 case definition and met pneumonia (SRI) case definition as wel as those that did not meet the SRI case definition. **Unable to assign:** no lineage assigned due to poor- sequence quality OR low viral load (ct=>35) OR variant PCR could not assign variant and no sequencing result

Pending: outstanding variant results

Table 12. Number of SARS-CoV-2 positive cases* by variant (variant PCR and/or sequencing) identified and total number of samples tested by hospital, pneumonia surveillance public hospitals, 02/03/2020 – 28/11/2021

Hospital (Province)	Non-Alpha/ Beta/Delta	20I (Alpha)	20H (Beta)	21A (Delta)	20D (C.1.2)	Pending	Unable to assign	Total SARS- CoV-2 positive
Edendale (KZ)	102		75	83	2	59	151	472
Helen Joseph-Rahima Moosa (GP)	134	5	120	101	5	20	125	510
Klerksdorp-Tshepong (NW)	130	9	113	90	2	13	120	477
Mapulaneng- Matikwana (MP)	16	0	90	29	Ο	17	50	202
Red Cross (WC)	15	О	5	7	0	7	27	61
Mitchell's Plain (WC)	50	0	50	96	0	14	143	353
Tintswalo (MP)	0	1	12	15	0	5	19	52
Total:	447	16	465	422	9	133	635	2 127

CP: Gauteng; KZ: KwaZulu-Natal; NW: North West; MP: Mpumalanga (Tintswalo started enrolling on the 10th Feb 2021); WC: Western Cape *Specimens are from hospitalized patients at 7 sentinel sites in 5 provinces who met suspected SARS-CoV-2 case definition and met pneumonia (SRI) case definition as well as those that did not meet the SRI case definition.

Unable to assign: no lineage assigned due to poor- sequence quality OR low viral load (ct=>35) OR variant PCR could not assign variant and no sequencing result **Pending:** outstanding variant results

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SUMMARY OF LABORATORY CONFIRMED SARS-COV-2 CASES

Table 13. Characteristics of individuals with laboratory-confirmed SARS-CoV-2, enrolled in influenza-like illness(ILI) and pneumonia surveillance programmes, South Africa, 2 March 2020 - 28 November 2021

Characteristic	Influenza-like illness (ILI), public-sector, n=723 (%)	Pneumonia, n=2 127 (%)		
Age group				
0-9	53/723 (7)	139/2127 (7)		
10-19	53/723 (7)	15/2127 (1)		
20-39	338/723 (47)	389/2127 (18)		
40-59	224/723 (31)	791/2127 (37)		
60-79	53/723 (7)	707/2127 (33)		
≥80	2/723 (<1)	77/2127 (4)		
Sex-female	437/723 (60)	1300/2127 (61)		
Province*				
Gauteng	N/A	510/2127 (24)		
KwaZulu-Natal	115/723 (16)	472/2127 (22)		
Mpumalanga**	87/723 (12)	254/2127 (12)		
North West	367/723 (51)	477/2127 (22)		
Western Cape	154/723 (21)	415/2127 (19)		
Race				
Black	545/721 (76)	1685/2119. (79)		
Coloured	145/721 (20)	327/2119 (16)		
Asian/Indian	4/721 (1)	52/2119 (2)		
White	22/721 (3)	44/2119 (2)		
Other	5/721 (1)	11/2119 (<1)		
Variant ^{\$\$}				
Non-Alpha/Beta/Delta	152/537 (28)	447/1506 (30)		
20I(Alpha)	15/537 (3)	16/1506 (1)		
20H(Beta)	199/537 (37)	463/1506 (31)		
21A(Delta)	139/537 (26)	425/1506 (28)		
20D(C.1.2)	1/537 (0.2)	8/1506 (1)		
Pending results	28/537 (5)	147/1506 (10)		

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Characteristic	Influenza–like illness (ILI), public-sector, n=723 (%)	Pneumonia, n=2 127 (%)		
Presentation				
Fever	564/721 (78)	976/2119 (46)		
Cough	702/721 (97)	2025/2119 (96)		
Shortness of breath	263/721 (37)	1659/2119 (78)		
Chest pain	317/721 (44)	863/2119 (41)		
Diarrhoea	54/721 (8)	137/2119 (6)		
Underlying conditions				
Hypertension ^{\$}	17/691 (2)	249/1896 (13)		
Cardiac	2/721 (<1)	49/2119 (2)		
Lung disease	0/721 (0)	3/2119 (<1)		
Diabetes	17/721 (2)	438/2119 (21)		
Cancer	2/721 (<1)	13/2119 (1)		
Tuberculosis	11/721 (2)	58/2119 (3)		
HIV-infection	135/721 (19)	475/2119 (22)		
Other ***	12/721 (2)	90/2119 (4)		
SARS-CoV-2 Vaccine				
Pfizer-BioNTech (1st dose)	12/433 (3)	83/1230 (7)		
Pfizer-BioNTech (2nd dose)	2/433 (0.2)	11/1230 (1)		
Johnson & Johnson	14/433 (3)	14/1230 (1)		
Unknown	0/433 (0)	5/1230 (0.4)		
Management				
Oxygen therapy	9/721 (1)	1658/2119 (78)		
ICU admission	N/A	50/2119 (2)		
Ventilation	N/A	56/2119 (3)		
Outcome***				
Died	0/718 (0)	355/2054 (17)		

*ILI surveillance not conducted in Gauteng province

arted enrolling on the 10th November 2020 and an additional SARI site started enrolling on the 10th Februar

****Outcome includes patients who are still bespitalised by

^{\$}Data on hypertension was not collected on all cases.

Note: Children may be over-represented amongst hospitalised patients due to the inclusion of a large paediatric hospital in Cape Town.

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METHODS

SARS-CoV-2 Testing

March 2020 – March 2021: SARS-CoV-2 was detected using the Roche E gene real-time PCR assay (Corman et al. Euro Surveillance 2020) with cycle threshold (Ct) <40 interpreted as positive for SARS-CoV-2. From April 2021 to date the laboratory changed to the Allplex[™] SARS-CoV-2/FluA/FluB/ RSV kit (Seegene Inc., Seoul, South Korea), with positivity assigned if the PCR cycle threshold (Ct) was <40 for ≥1 gene targets (N, S or RdRp).

A confirmed SARS-CoV-2 case is a person of any age enrolled in surveillance with laboratory confirmation of SARS-CoV-2 infection by PCR. Only positive SARS-CoV-2 specimens on PCR are further tested to determine variant/lineage type by variant PCR or genomic sequencing.

Variant PCR

Allplex[™] SARS-CoV-2 Variants I PCR detects Alpha and Beta/ Gamma variants. The assay was conducted on all SARS-CoV-2-positive samples from 1 March 2020 – 30 June 2021.

Allplex[™] SARS-CoV-2 Variants II PCR detects Delta variant and distinguishes Beta from Gamma. The assay was conducted on SARS-CoV-2-positive samples from 1 Jan to 30 June 2021.

Extraction: Total nucleic acids were extracted from 200µl NP/ OP samples in universal or viral transport medium using a MagNA Pure 96 automated extractor and DNA/Viral NA Small Volume v2.0 extraction kit (Roche Diagnostics, Mannheim, Germany).

SARS-CoV-2 genomic surveillance

SARS-CoV-2 Whole-Genome Sequencing and Genome Assembly

RNA Extraction

RNA was extracted either manually or automatically in batches, using the QIAamp viral RNA mini kit (QIAGEN, CA, USA) or the Chemagic 360 using the CMG-1049 kit (PerkinElmer, MA, USA). A modification was done on the manual extractions by adding 280 μ l per sample, in order to increase yields. 300 μ l of each sample was used for automated magnetic bead-based extraction using the Chemagic 360. RNA was eluted in 60 μ l of the elution buffer. Isolated RNA was stored at -80 °C prior to use.

PCR and Library Preparation

Sequencing was performed using the Illumina COVIDSeq protocol (Illumina Inc., CA, USA) or nCoV-2019 ARTIC network sequencing protocol v3 (https://artic.network/ncov-2019). These are amplicon-based next-generation sequencing approaches. Briefly, for the nCoV-2019 ARTIC network sequencing protocol, the first strand synthesis was carried out on extracted RNA samples using random hexamer primers from the SuperScript IV reverse transcriptase synthesis kit (Life Technologies, CA, USA) or LunaScript RT SuperMix Kit (New England Biolabs (NEB), MA, USA). The synthesized cDNA was amplified using multiplex polymerase chain reactions (PCRs) using ARTIC nCoV-2019 v3 primers. For the COVIDSeq protocol, the first strand synthesis was carried out using random hexamer primers from Illumina and the synthesized cDNA underwent two separate multiplex PCR reactions.

For Illumina sequencing using the nCoV-2019 ARTIC network sequencing protocol, the pooled PCR products underwent bead-based tagmentation using the Nextera Flex DNA library preparation kit (Illumina Inc., CA, USA). The adapter-tagged amplicons were cleaned up using AmpureXP purification beads (Beckman Coulter, High Wycombe, UK) and amplified using one round of PCR. The PCRs were indexed using the Nextera CD indexes (Illumina Inc., CA, USA) according to the manufacturer's instructions. For COVIDSeg sequencing protocol, pooled PCR amplified products were processed for tagmentation and adapter ligation using IDT for Illumina Nextera UD Indexes. Further enrichment and cleanup was performed as per protocols provided by the manufacturer (Illumina Inc., CA, USA). Pooled samples from both COVIDSeq protocol and nCoV-2019 ARTIC network protocol were quantified using Qubit 3.0 or 4.0 fluorometer (Invitrogen Inc., MA, USA) using the Qubit dsDNA High Sensitivity assay according to manufacturer's instructions. The fragment sizes were analyzed using TapeStation 4200 (Invitrogen Inc., MA, USA). The pooled libraries were further normalized to 4nM concentration and 25 µl of each normalized pool containing unique index adapter sets were combined in a new tube. The final library pool was denatured and neutralized with 0.2 N sodium hydroxide and 200 mM Tris-HCL (pH7), respectively. 1.5 pM sample library was spiked with 2% PhiX. Libraries were loaded onto a 300-cycle NextSeg 500/550 HighOutput Kit v2 and run on the Illumina NextSeq 550 instrument (Illumina Inc., CA, USA).

Assembly, Processing and Quality Control of Genomic Sequences

Raw reads from Illumina sequencing were assembled using the Exatype NGS SARS-CoV-2 pipeline v1.6.1, (https://sarscov-2.exatype.com/). The resulting consensus sequence was further manually polished by considering and correcting indels in homopolymer regions that break the open reading frame (probably sequencing errors) using Aliview v1.27, (http:// ormbunkar.se/aliview/) (Larsson, 2014). Mutations resulting in mid-gene stop codons and frameshifts were reverted to wild type. All assemblies determined to have acceptable quality (defined as having at least 1 000 000 reads and at least 40 % 10 X coverage) were deposited on GISAID (https://www.gisaid. org/) (Elbe & Buckland-Merrett, 2017; Shu & McCauley, 2017).

Classification of Lineage, Clade and Associated Mutations

Assembled genomes were assigned lineages using the 'Phylogenetic Assignment of Named Global Outbreak Lineages' (PANGOLIN) software suite (https://github.com/ hCoV-2019/pangolin) (Rambaut et al., 2020), a tool used for dynamic SARS-CoV-2 lineage classification. The SARS-CoV-2 genomes in our dataset were also classified using the clade classification proposed by NextStrain (https://nextstrain.org/), a tool built for real-time tracking of the pathogen evolution (Hadfield et al., 2018).

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