Division of the National Health Laboratory Service

Weekly respiratory pathogens report Week 46 of 2022

<u>Highlights</u>

- The 2022 influenza season started in week 17 (week starting 25 April 2022), and ended in week 42 (week starting 16 October) when the influenza detection rate in the pneumonia surveillance programme dropped below 10% and remained below the seasonal threshold for at least two weeks.
- In 2022 to date, 1 149 influenza cases have been detected from all surveillance programmes. The beginning of the 2022 season was dominated by influenza A(H1N1)pdm09, whereas influenza B Victoria was most commonly detected towards the end of the season. Influenza A(H3N2) circulated throughout the season.
- The 2022 respiratory syncytial virus (RSV) season which started in week 7 (week starting 14 February 2022) when the RSV detection rate among children under five years of age, in the pneumonia surveillance programme, rose above the seasonal threshold. The season ended in week 26. In 2022 to date, 892 respiratory syncytial virus (RSV) cases have been detected and activity remains below threshold in all surveillance programmes. Both RSV A and B have been detected throughout the year.
- In 2022 to date, we detected 95 cases of *Bordetella pertussis;* 81% (77/95) were detected from the Western Cape, 11% (10/95) from Mpumalanga, 5% (5/95) from Gauteng and 3% (3/95) from KwaZulu-Natal. Detection rates for *B. pertussis* cases in influenza-like illness surveillance (ILI) and pneumonia surveillance are similar in November compared to October (3%: October ILI 6/222, pneumonia surveillance 17/652; November ILI 2/60, pneumonia surveillance 10/303), and remain elevated compared to June detection levels (0%, ILI 0/189, pneumonia surveillance 1/800).
- In 2022 to date, a total of 834 COVID-19 cases were detected from all surveillance programmes. Of the 433 hospitalised COVID-19 cases reported, with available data on outcome, 7% (30) died. Since week 32 of 2022, COVID-19 cases have remained relatively stable in ILI and SARI surveillance programmes. Although the detection rate in the viral watch surveillance programme remain high (19-37%) in weeks 43 through 46, the number of samples submitted is low.
- Of the 784 specimens sequenced, 541 were assigned a variant. Omicron was the dominant variant (99%, 536/541); of which 17% (91/536) was Omicron 21K/BA.1, 11% (77/536) was Omicron 21L/BA.2, 0.4% (2/536) was Omicron 21M/BA.3, 30% (159/536) was Omicron 22A/BA.4, 38% (197/536) was Omicron 22B/BA.5, and 0.2% (1/536) was Omicron 22C/BA.2.12.1. Alpha, Delta, C.1.2 (20D) and XAY variants contributed <1% each. The remaining 243/784 (31%) could not be assigned a variant, due to a low viral load or insufficient sample.
- A lower number of specimens was submitted in week 30 (31 July 6 August 2022) due to staff training this likely affected numbers and proportions of viruses detected during this week, therefore trends should be regarded with caution.

Programme Descriptions

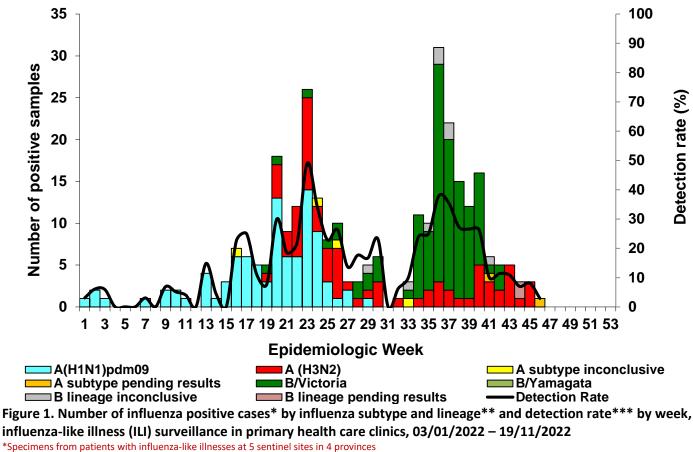
Programme	Influenza-like illness (ILI)	Viral Watch	National syndromic surveillance for pneumonia		
Start year	2012	1984	2009		
Provinces*	KΖ	EC	EC		
	NW	FS	GP		
	WC	GP	KZ		
	MP	LP	MP		
		MP	NW		
		NC	WC		
		NW	we		
		WC			
Type of site	Primary health care clinics	General practitioners	Public hospitals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

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 (ILI in primary health care clinics) 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. For RSV, thresholds from pneumonia surveillance, using data from children aged < 5 years are used to define the start and end of the season.

* EC: Eastern Cape; FS: Free State; GP: Gauteng; KZ: KwaZulu-Natal; LP: Limpopo; MP: Mpumalanga: NC: Northern Cape; NW: North West; WC: Western Cape **Symptoms include ANV 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).***INF: influenza virus; RSV: respiratory syncytial virus; BP: Bordetella pertusis; SARS-CoV-2: severe acute respiratory syndrome coronavirus 2

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**Influenza was detected in two (7%) of 27 specimens from patients who met suspected SARS-CoV-2 or B. pertussis case definition but did not meet influenzalike illness (ILI) case definition. Of which one (50%) was influenza A(H3N2) and one (50%) was influenza B(Victoria). 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

Two dual infections of influenza B(Victoria) + influenza A(H1N1)pdm09 in week 24 and B(Victoria) + influenza A(H3N2) in week 39 not included in the epidemiological curve.

Table 1. Number of laboratory-confirmed influenza* cases by subtype and lineage and total number of samples tested by clinic and province, influenza-like illness (ILI) surveillance in primary health care clinics, 03/01/2022 – 19/11/2022

Clinic (Province)	A(H1N1) pdm09	A(H3N2)	A subtype in- conclusive**	A subtype pending results** *	B/ Victoria	B/ Yamagata	B lineag e in- conclu sive*	B lineage pending results* **	Total sample s
Agincourt (MP)	21	11	0	2	15	0	2	0	271
Eastridge (WC)	11	14	0	0	31	0	1	2	300
Edendale Gateway (KZ)	23	31	0	0	34	0	3	0	546
Jouberton (NW)	24	10	1	0	26	0	1	0	366
Mitchell's Plain (WC)	15	9	3	0	11	0	1	0	261
Total:	94	75	4	2	117	0	8	2	1744

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

*Influenza was detected in two (7%) of 27 specimens from patients who met suspected SARS-CoV-2 or B. pertussis case definition but did not meet influenza-like

illness (ILI) case definition. Of which one (50%) was influenza A(H3N2) and one (50%) was influenza B(Victoria). These are not included in the table.

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

***Influenza A subtype or B lineage results are pending

Two dual infections of influenza B(Victoria) + influenza A(H1N1)pdm09 from Eastridge (WC) and influenza B(Victoria) + influenza A(H3N2) from Agincourt (MP) indicated in both columns.

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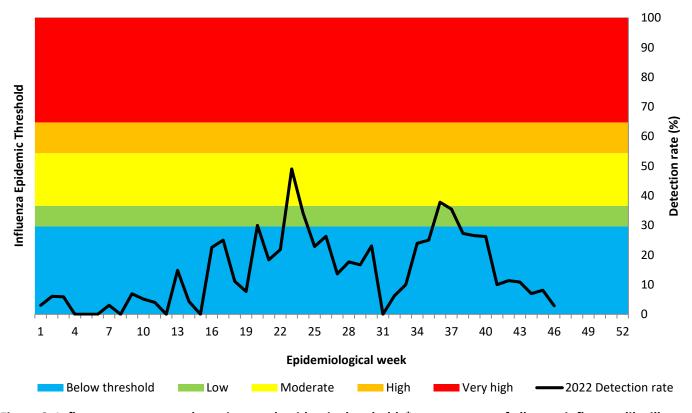


Figure 2. Influenza percentage detections and epidemic thresholds* among cases of all ages, influenza-like illness (ILI) surveillance in primary health care clinics, 03/01/2022 – 19/11/2022 *Thresholds based on 2012-2019 data

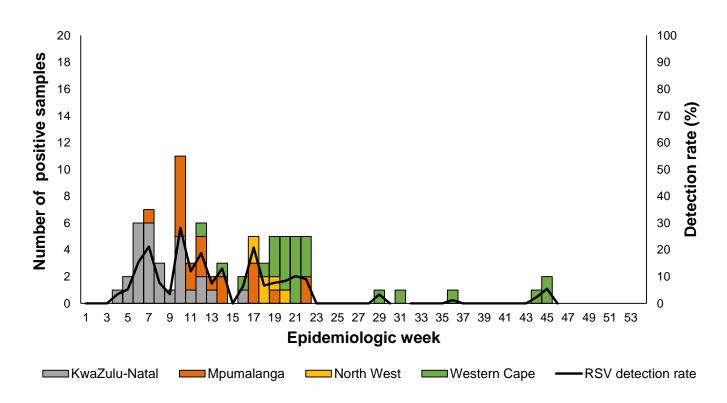


Figure 3. Number of patients testing positive for respiratory syncytial virus* by province and detection rate by week, influenza-like illness (ILI) surveillance in primary health care clinics, 03/01/2022 – 19/11/2022

*RSV was not detected from 27 specimens of patients who met suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet influenza-like illness (ILI) case definition.

**Only reported for weeks with >10 specimens submitted

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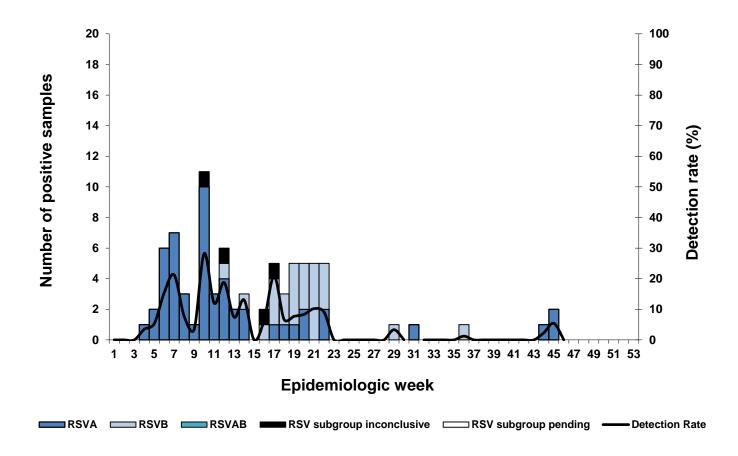


Figure 4. Number of patients testing positive for respiratory syncytial virus* by subgroup and detection rate by week, influenza-like illness (ILI) surveillance in primary health care clinics, 03/01/2022 – 19/11/2022

*RSV was not detected from 27 specimens of patients who met suspected SARS-CoV-2 or *B. pertussis* 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 submitted

RSV AB: Both RSV A and B subgroups identified.

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

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, 03/01/2022 – 19/11/2022

Clinic (Province)	RSVA	RSVB	RSVAB**	RSV subgroup inconclusive* **	RSV subgroup pending** **	Total samples
Agincourt (MP)	18	2	0	1	0	271
Eastridge (WC)	4	10	0	0	0	300
Edendale Gateway (KZ)	26	0	0	3	0	546
Jouberton (NW)	3	3	0	0	0	366
Mitchell's Plain (WC)	1	10	0	0	0	261
Total	52	25	0	4	0	1744

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

*RSV was not detected from 27 specimens of patients who met suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet influenza-like illness (ILI) case definition. These are not included in the table.

**RSV AB: Both RSV A and B subgroups identified

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

****RSV results for subgroups are pending

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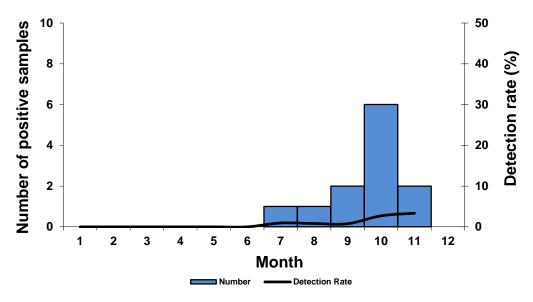


Figure 5. Number of patients testing positive for *B. pertussis** and detection rate by month, influenza-like illness (ILI) surveillance primary health care clinics**, 03/01/2022 – 19/11/2022

*No B. pertussis was detected in 27 specimens of patients who met the suspected SARS-CoV-2 or B. pertussis case definition but did not meet influenza-like illness (ILI) case definition. These are not included in the epidemiological curve.

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

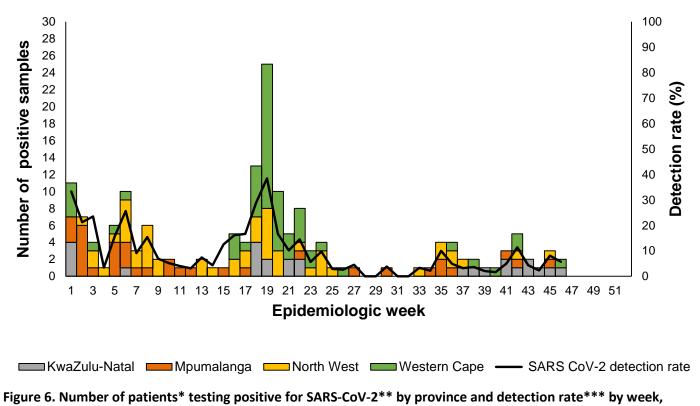
 Table 3. Number of patients testing positive for *B. pertussis** identified and total number of samples tested

 by province, influenza-like illness (ILI) surveillance primary health care clinics, 03/01/2022 – 19/11/2022

Clinic (Province)	<i>B. pertussis</i> Positive	Total samples
Agincourt (MP)	2	266
Eastridge (WC)	6	289
Edendale Gateway (KZ)	2	533
Jouberton (NW)	0	356
Mitchell's Plain (WC)	2	254
Total:	12	1698

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

*No *B. pertussis* was detected in 27 specimens of patients who met the suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet influenza-like illness (ILI) case definition. These are not included in the table.



influenza-like illness (ILI) surveillance in primary health care clinics, 03/01/2022 – 19/11/2022

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

**SARS-CoV-2 was detected in 5 of 27 (19%) specimens from patients who met suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet influenzalike illness (ILI) case definition. These are not included in the epidemiological curve.

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

 Table 4. 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, 03/01/2022 – 19/11/2022

Clinic (Province)	SARS-CoV-2 positive	Total samples tested
Agincourt (MP)	35	271
Eastridge (WC)	12	300
Edendale Gateway (KZ)	26	546
Jouberton (NW)	52	366
Mitchell's Plain (WC)	46	261
Total:	171	1744

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

*SARS-CoV-2 was detected in 5 of 27 (19%) specimens from patients who met suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet influenzalike illness (ILI) case definition. These are not included in the table.

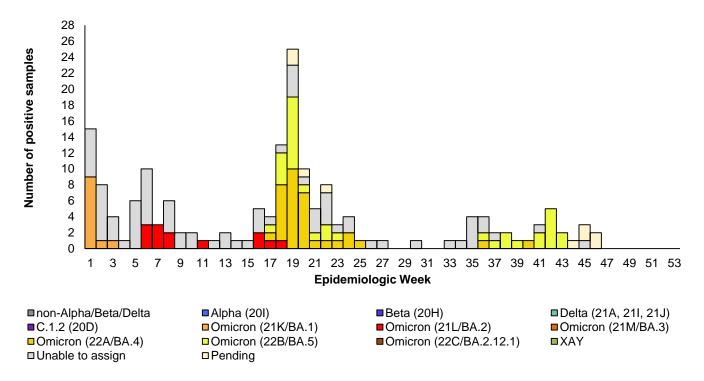


Figure 7. Number and detection rate of laboratory-confirmed SARS-CoV-2* cases by variant type (variant PCR/sequencing) and week, influenza-like illness (ILI) surveillance in primary health care clinics, 03/01/2022 – 19/11/2022

*Specimens are from patients with influenza-like illness at 5 sentinel sites in 4 provinces who met influenza-like illness (ILI), suspected SARS-CoV-2 or *B. pertussis* 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

health care clinics, 03/01/2022 – 19/11/2022											
Clinic (Province)	Delta (21A, 21I, 21J)	Omicron (21K/BA.1)	Omicron (21L/BA.2)	Omicron (21M/BA.3)	Omicron (22A/BA.4)	Omicron (22B/BA.5)	Omicron (22C/ BA.2.12.1)	Unable to assign	Pending	Total SARS- CoV-2 positive	Total samples tested
Agincourt (MP)	0	4	3	0	0	1	0	29	0	37	276

Table 5. Number of cases positive for SARS-CoV-2^{*} 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, 03/01/2022 – 19/11/2022

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

*Specimens are from patients with influenza-like illness at 5 sentinel sites in 4 provinces who met influenza-like illness (ILI), suspected SARS-CoV-2 or *B. pertussis* case definition

**No cases of Alpha, Beta or 20D (C.1.2) variants detected.

Eastridge (WC)

Gateway (KZ) Jouberton

Edendale

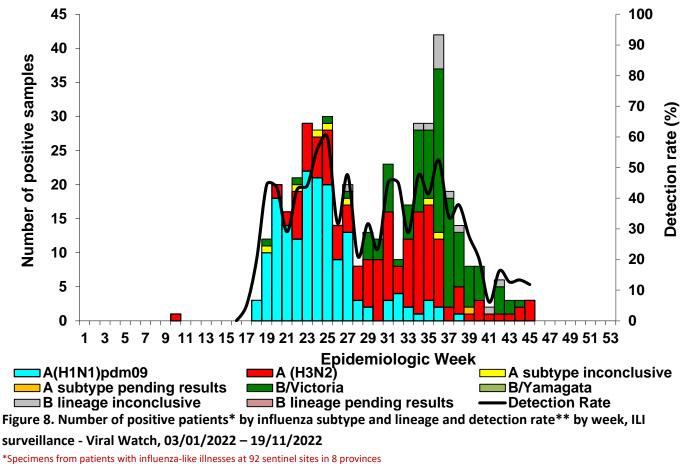
(NW) Mitchell's Plain

(WC)

Total:

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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**Only reported for weeks with >10 specimens submitted. Three-week rolling average of detection rate (current week and two weeks prior) used from week 42 onwards due to low number of specimens collected.

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

Three dual infections from GP (one influenza A(H3N2) + influenza A(H1N1)pdm09 in week 17, one influenza B(lineage inconclusive) + influenza A(H1N1)pdm09 in week 23, and one influenza A(H3N2) + influenza B(Victoria) in week 37) not included in the epidemiological curve.

Table 6. Number of laboratory-confirmed influenza cases by influenza subtype and lineage and total number of samples tested by province, ILI surveillance - Viral Watch, 03/01/2022 – 19/11/2022

Province	A(H1N1) pdm09	A(H3N2)	A subtype inconclusiv e	A subtype pending results*	B/Victor ia	B/Yamag ata	B lineage inconclus ive	B lineage pending results*	Total samples
Eastern Cape	20	8	0	0	10	0	1	0	60
Free State	7	0	0	0	0	0	0	0	8
Gauteng	85	50	4	1	62	0	7	0	840
Limpopo	2	2	1	0	1	0	0	0	8
Mpumalanga	7	2	0	0	4	0	3	0	37
North West	3	0	0	0	0	0	0	0	6
Northern Cape	0	0	0	0	0	0	0	0	0
Western Cape	41	87	2	0	36	0	2	0	359
Total:	165	149	7	1	113	0	13	0	1318

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

**Influenza A subtype or B lineage results are pending

Three dual infections from GP (one influenza A(H3N2) + influenza A(H1N1)pdm09 in week 17, one influenza B(lineage inconclusive) + influenza A(H1N1)pdm09 in week 23, and one influenza A(H3N2) + influenza B(Victoria) in week 37) indicated in both columns.

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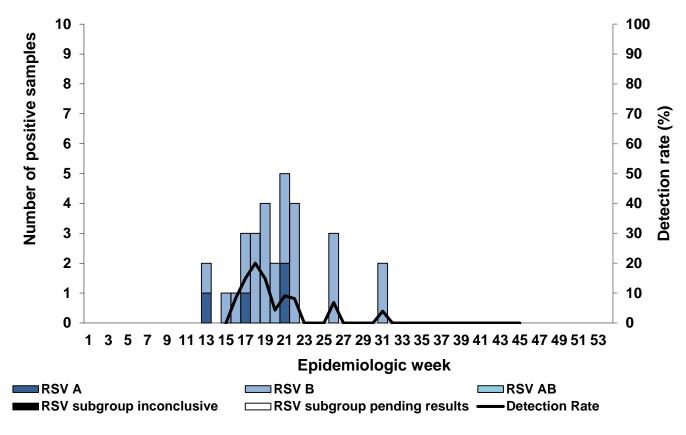


Figure 9. Number of RSV positive cases testing positive for respiratory syncytial virus (RSV)* by subgroup and detection rate** by week, ILI surveillance - Viral Watch, 03/01/2022 – 19/11/2022

*Specimens from patients with Influenza-like illnesses at 92 sentinel sites in 8 provinces **Only reported for weeks with >10 specimens submitted.

Table 7. Number of RSV positive cases identified and total number of samples tested by province, ILI surveillance - Viral Watch, 03/01/2022 – 19/11/2022

Province	RSV A	RSV B	RSV AB*	RSV subgroup inconclusive **	RSV subgroup pending results***	Total samples tested
Eastern Cape	0	1	0	0	0	60
Free State	0	0	0	0	0	8
Gauteng	4	13	0	0	0	840
Limpopo	0	0	0	0	0	8
Mpumalanga	0	0	0	0	0	37
North West	0	0	0	0	0	6
Northern Cape	0	0	0	0	0	0
Western Cape	0	12	0	0	0	359
Total:	4	26	0	0	0	1318

*RSV AB: Both RSV A and B subgroup identified

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

***RSV results for subgroups are pending

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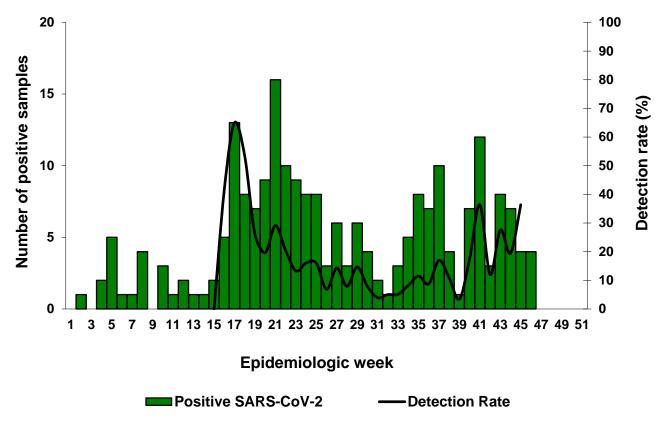


Figure 10. Number of patients testing positive for SARS-CoV-2*, by site and detection rate** by week, ILI surveillance - Viral Watch, 03/01/2022 – 19/11/2022

*Specimens from patients with influenza-like illnesses at 92 sentinel sites in 8 provinces **Only reported for weeks with >10 specimens submitted.

Table 8. Number of SARS-CoV-2 positive cases identified and total number tested by province, ILI surveillance- Viral Watch, 03/01/2022 – 19/11/2022

Province	SARS-CoV-2 positive	Total samples tested
Eastern Cape	4	60
Free State	0	8
Gauteng	155	840
Limpopo	1	8
Mpumalanga	3	37
North West	0	6
Northern Cape	0	0
Western Cape	62	359
Total:	225	1318

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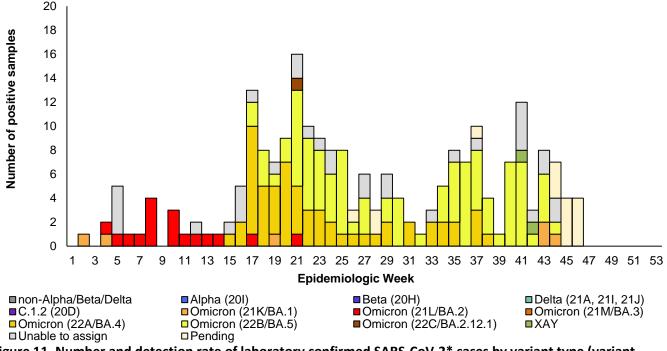


Figure 11. Number and detection rate of laboratory confirmed SARS-CoV-2* cases by variant type (variant PCR/sequencing) and week, ILI surveillance - Viral Watch, 03/01/2022 – 19/11/2022

^{*}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 ($C_t \ge 35$) **OR** variant PCR could not assign variant and no sequencing result **Pending**: outstanding variant results

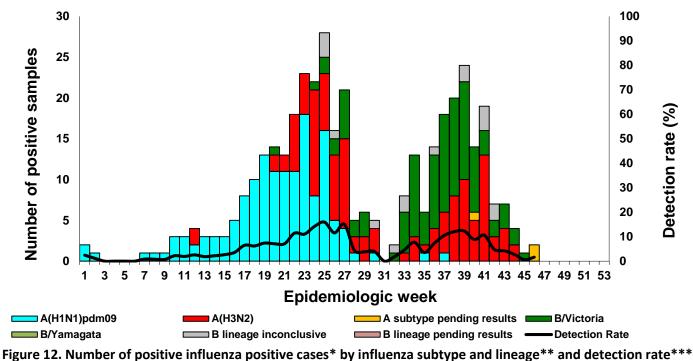
Table 9. Number of SARS-CoV-2* positive cases by variant (variant PCR and/or sequencing) identified and total
number of samples tested by province, ILI surveillance - Viral Watch, 03/01/2022 – 19/11/2022

Clinic (Province)	Omicron (21K/BA. 1)	Omicron (21L/BA. 2)	Omicron (21M/BA .3)	Omicron (22A/BA. 4)	Omicron (22B/BA. 5)	Omicron (22C/ BA.2.12. 1)	ХАҮ	Unable to assign	Pending	Total SARS- CoV-2 positive	Total samples tested
Eastern Cape	0	1	0	2	1	0	0	0	0	4	60
Free State	0	0	0	0	0	0	0	0	0	0	8
Gauteng	5	8	0	46	57	1	2	24	12	155	840
Limpopo	0	0	0	0	0	0	0	1	0	1	8
Mpumalanga	0	0	0	1	2	0	0	0	0	3	37
North West	0	0	0	0	0	0	0	0	0	0	6
Northern Cape	0	0	0	0	0	0	0	0	0	0	0
Western Cape	1	8	0	9	33	0	0	8	3	62	359
Total:	6	17	0	58	93	1	2	33	15	225	1318

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

**No cases of Alpha, Beta or 20D (C.1.2) variants detected.

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



by week, pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022

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

*Specimens from patients hospitalised with pneumonia at 11 sentinel sites in 6 provinces

**Influenza was not detected in 16 specimens from patients who met suspected the SARS-CoV-2 or B. pertussis 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

One dual infection of influenza B(Victoria) + influenza A(H3N2) in week 24 not included in the epidemiological curve.

Table 10. Number of laboratory confirmed influenza cases by subtype and lineage* and total number of samples tested by hospital, pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022

Hospital (Province)	A(H1N1)p dm09	A(H3N2)	A subtype inconclusive	A subtype pending results***	B/Victoria	B/Yamagata	B lineage inconclusive	B lineage pending results***	Total samples
Edendale (KZ)	26	16	1	0	11	0	2	0	896
Helen Joseph-Rahima Moosa (GP)	30	11	1	0	14	0	2	0	1313
Klerksdorp-Tshepong (NW)	28	5	0	1	16	0	3	0	561
Livingstone (EC)	9	11	2	0	7	0	2	0	515
Mapulaneng- Matikwana (MP)	11	18	1	2	13	0	1	0	564
Mitchell's Plain (WC)	5	13	1	0	5	0	3	0	709
Red Cross (WC)	9	19	2	0	14	0	1	0	1252
Tambo Memorial (GP)	0	7	0	0	3	0	1	0	71
Tembisa (GP)	7	2	0	0	12	0	1	0	337
Tintswalo (MP)	18	19	1	0	4	0	0	0	354
Tygerberg (WC)	4	4	1	0	3	0	0	0	167
Total:	147	125	10	3	102	0	16	0	6739

EC: Eastern Cape (Livingstone started enrolling on the 3rd of May 2022); GP: Gauteng (Tembisa started enrolling on the 10th of March 2022 and Tambo Memorial on the 21st of September 2022); KZ: KwaZulu-Natal; NW: North West; MP: Mpumalanga; WC: Western Cape (Tygerberg started enrolling on the 20th April 2022)

*Influenza was not detected in 16 specimens from patients who met suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet pneumonia (SRI) case definition. These are not included in the table.

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

***Influenza A subtype or B lineage results are pending

One dual infection of influenza B(Victoria) + influenza A(H3N2) in week 24 from Tintswalo (MP) indicated in both columns.

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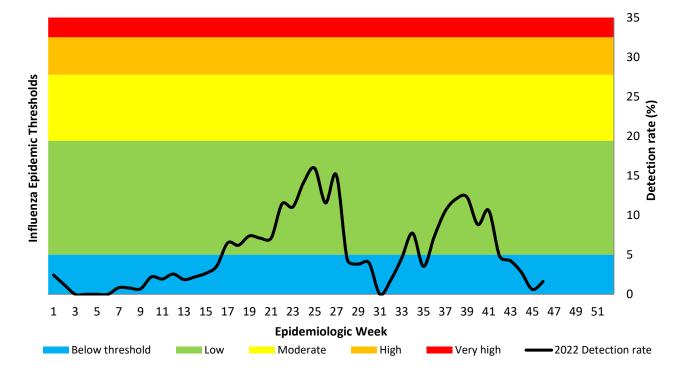
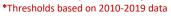


Figure 13. Influenza percentage detections and epidemic thresholds* among cases of all ages, pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022



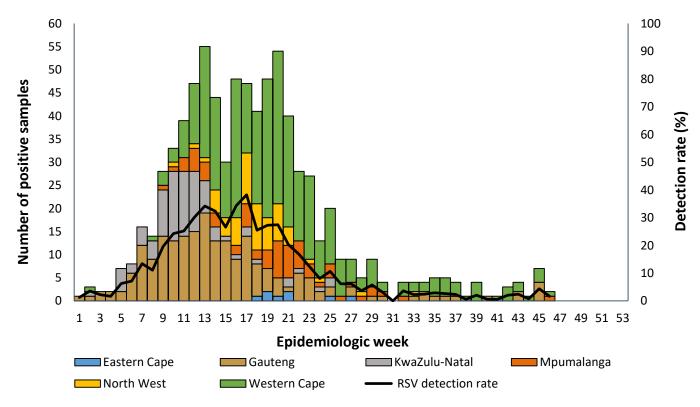


Figure 14. Number of patients (all ages) testing positive for respiratory syncytial virus* by province and detection rate by week, pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022

Specimens from patients hospitalised with pneumonia at 11 sentinel sites in 6 provinces.

*RSV was not detected in 16 specimens from patients who met suspected SARS-CoV-2 or B. pertussis case definition but did not meet pneumonia (SRI) case definition.

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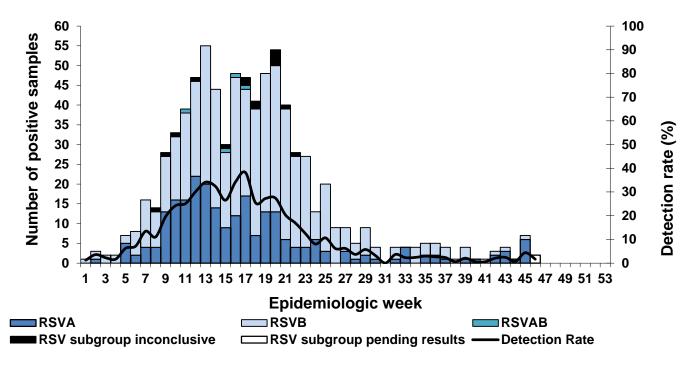


Figure 15. Number of patients (all ages) testing positive for respiratory syncytial virus* by subgroup and detection rate by week, pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022

Specimens from patients hospitalised with pneumonia at 11 sentinel sites in 6 provinces.

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

RSV AB: Both RSV A and B subgroup identified

RSV subgroup pending: RSV results for subgroups are pending

*RSV was not detected in 16 specimens from patients who met suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet pneumonia (SRI) case definition. These are not included in the epidemiological curve.

Table 11. Number of patients (all ages) positive for respiratory syncytial virus subgroups* by subgroups identified and total number of samples tested by hospital, pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022

Hospital (Province)	RSVA	RSVB	RSVAB**	RSV subgroup inconclusive** *	RSV subgroup pending** **	Total samples
Edendale (KZ)	86	1	0	2	0	896
Helen Joseph-Rahima Moosa (GP)	47	155	3	1	0	1313
Klerksdorp-Tshepong (NW)	30	31	1	0	0	561
Livingstone (EC)	1	6	0	1	0	515
Mapulaneng-Matikwana (MP)	18	25	0	0	1	564
Mitchell's Plain (WC)	10	67	0	0	0	709
Red Cross (WC)	46	212	0	8	1	1252
Tambo Memorial (GP)	0	0	0	0	0	71
Tembisa (GP)	0	2	0	0	0	337
Tintswalo (MP)	4	15	0	3	0	354
Tygerberg (WC)	0	4	0	0	0	167
Total:	242	518	4	15	2	6739

EC: Eastern Cape (Livingstone started enrolling on the 3rd of May 2022); GP: Gauteng (Tembisa started enrolling on the 10th of March 2022 and Tambo Memorial on the 21st of September 2022); KZ: KwaZulu-Natal; NW: North West; MP: Mpumalanga; WC: Western Cape (Tygerberg started enrolling on the 20th April 2022)

*RSV was not detected in 16 specimens from patients who met suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet pneumonia (SRI) case definition. These are not included in the table.

**RSV AB: Both RSV A and B subgroup identified

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

****RSV results for subgroups are pending

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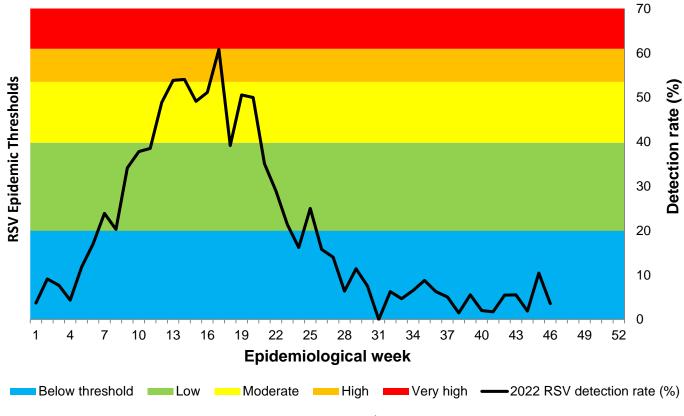
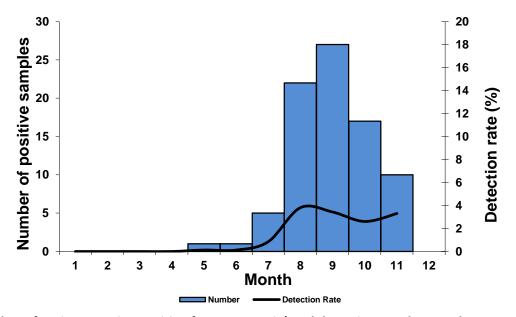
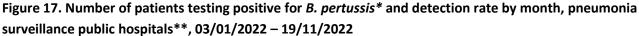


Figure 16. RSV percentage detections and epidemic thresholds* among children aged < 5 years, pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022 *Thresholds based on 2010-2019 data





*No *B. pertussis* was detected in 16 specimens of patients who met the suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet Pneumonia Surveillance case definition. These are not included in the epidemiologic curve. *Specimens from patients hospitalised with pneumonia at 11 sentinel sites in 6 provinces.

Table 12. Number of patients testing positive for *B. pertussis** identified and total number of samples tested by hospital and province, pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022

Hospital (Province)	<i>B. pertussis</i> Positive	Total samples	
Edendale (KZ)	1	880	
Helen Joseph-Rahima Moosa (GP)	1	1297	
Klerksdorp-Tshepong(NW)	0	551	
Livingstone (EC)	0	501	
Mapulaneng-Matikwana (MP)	6	546	
Mitchell's Plain (WC)	13	689	
Red Cross (WC)	55	1228	
Tambo Memorial (GP)	3	66	
Tembisa (GP)	1	333	
Tintswalo (MP)	2	352	
Tygerberg (WC)	1	162	
Total:	83	6605	

EC: Eastern Cape (Livingstone started enrolling on the 3rd of May 2022); GP: Gauteng (Tembisa started enrolling on the 10th of March 2022 and Tambo Memorial on the 21st of September 2022); KZ: KwaZulu-Natal; NW: North West; MP: Mpumalanga; WC: Western Cape (Tygerberg started enrolling on the 20th April 2022)

*No *B. pertussis* was detected in 16 specimens of patients who met the suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet the pneumonia (SRI) case definition. These are not included in the table.

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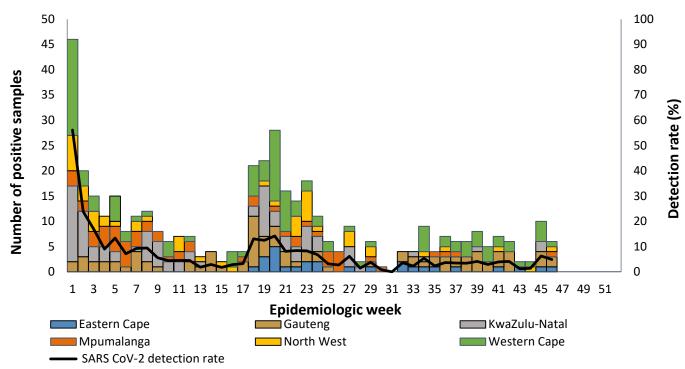


Figure 18. Number of patients testing positive for SARS-CoV-2^{**} by province and detection rate by week, pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022

*Specimens from patients hospitalized with pneumonia at 11 sentinel sites in 6 provinces. **SARS-CoV-2 was detected in 6 of 16 (38%) specimens from patients who met suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet pneumonia (SRI) case definition. These are not included in the epidemiological curve.

Table 13. Number of patients positive for SARS-CoV-2* and total number of samples tested by hospital, pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022

Hospital (Province)	SARS-CoV-2 positive	Total samples tested	
Edendale (KZ)	90	896	
Helen Joseph-Rahima Moosa (GP)	64	1313	
Klerksdorp-Tshepong (NW)	49	561	
Livingstone (EC)	27	515	
Mapulaneng-Matikwana (MP)	36	564	
Mitchell's Plain (WC)	57	709	
Red Cross (WC)	52	1252	
Tambo Memorial (GP)	5	71	
Tembisa (GP)	21	337	
Tintswalo (MP)	20	354	
Tygerberg (WC)	6	167	
Total:	427	6739	

EC: Eastern Cape (Livingstone started enrolling on the 3rd of May 2022); GP: Gauteng (Tembisa started enrolling on the 10th of March 2022 and Tambo Memorial on the 21st of September 2022); KZ: KwaZulu-Natal; NW: North West; MP: Mpumalanga; WC: Western Cape (Tygerberg started enrolling on the 20th April 2022)

*SARS-CoV-2 was detected in 6 of 16 (38%) specimens from patients who met suspected SARS-CoV-2 or *B. pertussis* case definition but did not meet pneumonia (SRI) case definition. These are not included in the table.

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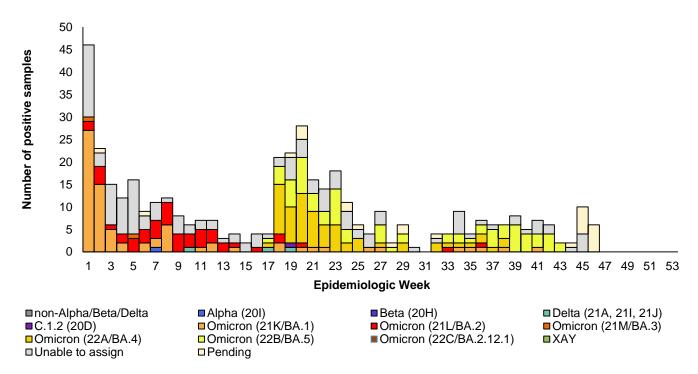


Figure 19. Number and detection rate of laboratory-confirmed SARS-CoV-2 cases* by variant type (variant PCR/sequencing), pneumonia surveillance public hospitals, 03/01/2022 – 19/11/2022

*Specimens are from hospitalized patients at 11 sentinel sites in 6 provinces who met the pneumonia (SRI), suspected SARS-CoV-2 or *B. pertussis* case definition Unable to assign: no lineage assigned due to poor- sequence quality OR low viral load (Ct235) OR variant PCR could not assign variant and no sequencing result Pending: outstanding variant results

Table 14. 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, 03/01/2022 – 19/11/2022

Hospital (Province)	Delta (21A, 21I, 21J)	Omicron (21K/BA. 1)	Omicron (21L/BA. 2)	Omicron (21M/B A.3)	Omicron (22A/BA .4)	Omicron (22B/BA .5)	Omicron (22C/ BA.2.12. 1)	Unable to assign	Pending	Total SARS- CoV-2 positive	Total samples tested
Edendale (KZ)	1	24	13	1	3	21	0	27	3	94	905
Helen Joseph- Rahima Moosa (GP)	0	8	9	0	13	12	0	16	5	64	1313
Klerksdorp- Tshepong (NW)	0	11	2	1	5	8	0	18	4	49	561
Livingstone (EC)	0	2	3	0	8	6	0	6	2	27	515
Mapulaneng- Matikwana (MP)	0	6	8	0	4	1	0	18	1	38	571
Mitchell's Plain (WC)	0	13	1	0	15	6	0	17	5	57	709
Red Cross (WC)	0	4	6	0	13	12	0	13	4	52	1252
Tambo Memorial (GP)	0	0	0	0	1	3	0	0	1	5	71
Tembisa (GP)	2	2	0	0	4	5	0	7	1	21	337
Tintswalo (MP)	0	4	4	0	1	1	0	10	0	20	354
Tygerberg (WC)	0	1	1	0	1	2	0	1	0	6	167
Total:	3	75	47	2	68	77	0	133	26	433	6755

EC: Eastern Cape (Livingstone started enrolling on the 3rd of May 2022); GP: Gauteng (Tembisa started enrolling on the 10th of March 2022 and Tambo Memorial on the 21st of September 2022); KZ: KwaZulu-Natal; NW: North West; MP: Mpumalanga; WC: Western Cape (Tygerberg started enrolling on the 20th April 2022) *Specimens are from hospitalized patients at 11 sentinel sites in 6 provinces who met the pneumonia (SRI), suspected SARS-CoV-2 or *B. pertussis* case definition **One case of Alpha variant from Helen Joseph-Rahima Moosa (GP), no cases of Beta variant and one case of 20D (C.1.2) variant detected from Edendale (KZ). **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 individuals with laboratory-confirmed SARS-CoV-2

Table 15: Characteristics of individuals with laboratory-confirmed SARS-CoV-2, enrolled in influenza-like illness (ILI) and pneumonia surveillance programmes, South Africa, 03/01/2022 – 19/11/2022

Characteristic	Influenza–like illness (ILI), public- sector, n=176 (%)	Pneumonia, public-sector, n=433 (%)
Age group (years)		
0-9	39/176 (22)	112/433 (26)
10-19	16/176 (9)	9/433 (2)
20-39	46/176 (26)	115/433 (27)
40-59	56/176 (32)	100/433 (23)
60-79	17/176 (10)	81/433 (19)
≥80	2/176 (1)	16/433 (4)
Sex-female	109/176 (62)	224/433 (52)
Province*	0 (176 (0)	27 (422 (5)
Eastern Cape	0/176 (0)	27/433 (6)
Gauteng	0/176 (0)	90/433 (21)
KwaZulu-Natal	28/176 (16)	94/433 (22)
Mpumalanga	37/176 (21)	58/433 (13)
North West	53/176 (30)	49/433 (11)
Western Cape Race	58/176 (33)	115/433 (27)
Black	104/176 (59)	319/433 (74)
Coloured	43/176 (24)	78/433 (18)
Asian/Indian		2/433 (0)
White	0/176 (0) 15/176 (9)	
Other	15/176 (9) 14/176 (8)	19/433 (4)
Variant	14/176 (8)	15/433 (3)
Non-Alpha/Beta/Delta	0/176 (0)	0/433 (0)
Alpha(20I) Bota(20H)	0/176 (0)	1/433 (0)
Beta(20H)	0/176 (0)	0/433 (0)
Delta(21A, 21I, 21J)	0/176 (0)	3/433 (1)
C.1.2(20D)	0/176 (0)	1/433 (0)
Omicron (21K/BA.1)	11/176 (6)	75/433 (17)
Omicron (21L/BA.2)	13/176 (7)	47/433 (11)
Omicron (21M/BA.3)	0/176 (0)	2/433 (0)
Omicron (22A/BA.4)	33/176 (19)	68/433 (16)
Omicron (22B/BA.5)	33/176 (19)	77/433 (18)
Omicron (22C/ BA.2.12.1)	0/176 (0)	0/433 (0)
XAY	0/176 (0)	0/433 (0)
Unable to assign**	77/176 (44)	133/433 (31)
Pending results***	9/176 (5)	26/433 (6)
Presentation		470 (400 (40)
Fever	116/163 (71)	170/420 (40)
Cough	162/164 (99)	394/420 (94)
Shortness of breath	61/159 (38)	281/414 (68)
Chest pain	66/159 (42)	162/414 (39)
Diarrhoea	20/159 (13)	40/414 (10)
Underlying conditions		(()
Hypertension	31/161 (19)	75/415 (18)
Cardiac	3/176 (2)	15/433 (3)
Lung disease	0/161 (0)	1/415 (0)
Diabetes	9/161 (6)	48/415 (12)
Cancer	0/176 (0)	4/433 (1)
Tuberculosis - Previous	1/176 (1)	4/433 (1)
Tuberculosis - Current	2/176 (1)	50/433 (12)
HIV-infection	21/176 (12)	159/433 (37)
Other ****	6/158 (4)	42/412 (10)
SARS-CoV-2 Vaccine		
Pfizer-BioNTech (1 st dose)	25/176 (14)	41/433 (9)
Pfizer-BioNTech (2 nd dose)	23/176 (13)	34/433 (8)
Johnson & Johnson (1 st dose)	18/176 (10)	29/433 (7)
Johnson & Johnson (2 nd dose)	3/176 (2)	2/433 (0)
Unknown	18/176 (10)	23/433 (5)
No vaccine	93/176 (53)	325/433 (75)
Management		
Oxygen therapy	0/163 (0)	238/411 (58)
ICU admission	0/163 (0)	3/411 (1)
Ventilation	0/163 (0)	36/411 (9)
Outcome****		
Died	0/163 (0)	30/400 (8)

*ILI surveillance not conducted in Gauteng or Eastern Cape province **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 results: outstanding variant results

****Choine includes patients who are still hospitalised, have been discharged or referred, and those who died

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

Of the 30 patients who died, 8 were in the 20-39-year age group, 10 were in 40-59 age group and 12 were ≥60 years; 19/30 (63%) were female.

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 (C_t) <40 interpreted as positive for SARS-CoV-2. From April 2021 to date the laboratory changed to the AllplexTM SARS-CoV-2/FluA/FluB/RSV kit (Seegene Inc., Seoul, South Korea), with positivity assigned if the PCR cycle threshold (C_t) 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

AllplexTM 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.

AllplexTM 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 µl per sample, in order to increase yields. 300 µl of each sample was used for automated magnetic bead-based extraction using the Chemagic 360. RNA was eluted in 60 µl 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 COVIDSeq sequencing protocol, pooled PCR amplified products were processed for tagmentation and adapter ligation using IDT for Illumina Nextera UD Indexes. Further enrichment and clean-up 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 Ta peStation 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 NextSeq 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, (<u>https://sars-cov-2.exatype.com/</u>). 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, (<u>http://ormbunkar.se/aliview/</u>) (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 (<u>https://www.gisaid.org/</u>) (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 (<u>https://github.com/hCoV-2019/pangolin</u>) (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 (<u>https://nextstrain.org/</u>), a tool built for real-time tracking of the pathogen evolution (Hadfield et al., 2018).