WEEKLY RESPIRATORY **PATHOGENS SURVEILLANCE** REPORT

SOUTH AFRICA WEEK 40 2021

NATIONAL INSTITUTE FOR

COMMUNICABLE DISEASES

CONTENTS

Surveillance programme description	2
Comments	3
Systematic Influenza-like illness (ILI) surveillance Influenza Respiratory syncytial virus SARS-CoV-2	4-8
Influenza-like illness (ILI) Viral Watch Influenza SARS-CoV-2	9-13
National syndromic surveilance for pneumonia Influenza Respiratory syncytial virus SARS-CoV-2	14-16
Summary of laboratory confirmed SARS-CoV-2 cases	17-18
SARS-CoV-2 Testing Methods	22

HIGHLIGHTS: WEEK 40

• Eight new cases of influenza from Gauteng (n=6) and North West (n=2) surveillance sites were detected in week 40. To date, 143 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 B(Victoria) was the most commonly detected lineage in both influenza-like illness (ILI) surveillance (n=21/34, 62%) and pneumonia surveillance (n=45/82, 55%). The influenza season has not started.
 RSV activity remains below seasonal threshold in both influenza-like illness (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=40/74, 54%) and pneumonia surveillance (n=206/402, 51%).

 From 2 March 2020 to 10 October 2021, a total of 2 882 COVID-19 cases were detected from all surveillance programmes. A sustained decline in detection rate has been noted in both ILI programme and pneumonia surveillance. Of the 1 969 hospitalised COVID-19 cases reported with available data on outcome, 339 (17%) died.

From 1 January 2021 to date, of the 1192/1298 (92%) SARS-CoV-2 positive cases with variant type results, Delta (425/871, 49%) and Beta (157/321, 49%), were the most detected variants in pneumonia surveillance and in ILI, respectively. Delta variant dominated both programmes (from week22, week starting 31st May 2021 until to date). Beta variant predominated from week 47 of 2020 to week 21 of 2021.

PAGE 1

WEEK **40** 2021

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 Suspected SARS-CoV-2 	ILI: An acute respiratory illness with a temperature (≥38°C) and cough, & onset ≤10 days Suspected SARS-CoV-2	 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:		
	Any person presenting with an acute (≤14 days) respiratory tract infection or other clinical illness compatible with COVID-19 ^β	Any person presenting with an acute (<14 days) respiratory tract infection or other clinical illness compatible with COVID-19 ^g	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 April 2020 – 31 March 2021: Roche E gene real-time PCR essay (Corman et al., Euro Surv 2020) 	 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) 	 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) 1 April 2021 to date: AllplexTM SARS- Development of the program of the		
	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)	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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PAGE 2

WEEK **40** 2021

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: Bordetello 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 influenza 2021 season has not started. 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 143 influenza cases detected in surveillance sites in 2021, the majority (n=87, 61%) were influenza B(Victoria).

ILI programme: In 2021 to date, specimens from 1 454 patients meeting ILI case definition were received from 4 ILI sites. Influenza was detected in 34 (2%) patients, of which two (6%) were influenza A(H1N1)pdm09, three (9%) influenza A(H3N2), one (3%) influenza A(inconclusive), one (3%) influenza A(pending subtype results), 21 (62%) influenza B(Victoria), three (9%) influenza B(lineage inconclusive) and three influenza B(lineage pending results).

Viral Watch programme: In 2021 to date, specimens were received from 180 patients from Viral Watch sites in 5 of the 8 provinces participating in surveillance. Influenza was detected in 11 (6%) patients, of which two (18%) were influenza A(H1N1) pdm09, five (45%) were influenza B(Victoria), and four (36%) influenza B(lineage inconclusive).

Pneumonia surveillance: Since the beginning of 2021, specimens from 4 901 patients with severe respiratory illness (SRI) were received from the 6 sentinel sites. Influenza was detected in 82 (2%) patients, of which 14 (17%) were influenza A(H1N1)pdm09, 12 (15%) influenza A(H3N2), seven (9%) influenza A(pending subtype results), 45 (55%) influenza B(Victoria) and four (5%) were influenza B(lineage inconclusive).

In addition, influenza B(Victoria) was detected in 16 of 745 (2%) patients who met suspected SARS-CoV-2 case definition but did not meet the pneumonia/ILI surveillance case definitions.

Respiratory syncytial virus

In 2021 to date, RSV detection has been reported from all surveillance programmes, activity remains below seasonal threshold. Of the 480 RSV cases detected in 2021, the majority (n=244, 51%) were RSV subgroup A.

ILI programme: In 2021 to date, 1 454 specimens from patients meeting the ILI case definition were tested and RSV was detected in 74 (5%) patients. Of which, 40 (54%) were RSV subgroup A, 31 (42%) RSV subgroup B, two (3%) RSV subgroup

A and B and one (1%) was RSV (subgroup inconclusive).

Viral Watch programme: In 2021 to date, 180 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).

Pneumonia surveillance: Since the beginning of 2021, 4 901 specimens were tested and RSV was detected in specimens of 402 (8%) patients. Of which, 206 (51%) were RSV subgroup A, 183 (46%) RSV subgroup B, four (1%) RSV subgroup A and B, eight (2%) RSV (subgroup inconclusive and one (0.2%) RSV (subgroup pending results).

In addition, RSV was detected in 21 of 745 (3%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet the pneumonia/ILI surveillance case definitions.

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

ILI programme: From March 2020 to date, 2 765 patients were tested and SARS-CoV-2 was detected in 566 (21%) patients. From 1 January 2021 to date, of the 321/339 (95%) with data on variant type, majority (157/321, 49%) were Beta variant which predominated from week1 to week 24, followed by Delta (139/321, 43%) variant which predominated from week 25 to week 36.

Viral Watch programme: From March 2020 to date, 459 patients presenting with ILI were tested and SARS-CoV-2 was detected in 84 (18%) patients. From 1 January 2021, of the 21/84 (25%) with data on variant type, majority were (19/21, 90%) were Delta variant which dominated from week 25 to week 30.

Pneumonia surveillance: From March 2020 to date, 8 917 patients with severe respiratory illness (SRI) were tested and SARS-CoV-2 was detected in 2 042 (23%) patients. From 1 January 2021 to date, of the 871/959 (91%) with data on variant type, majority were (425/871, 49%) were Delta variant which dominated from week 22 to week38 followed by Beta (401/871, 46%) variant which dominated from week 1 to 25.

In addition, SARS-CoV-2 was detected in 190 of 948 (20%) 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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PAGE 3

WEEK 40 2021

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

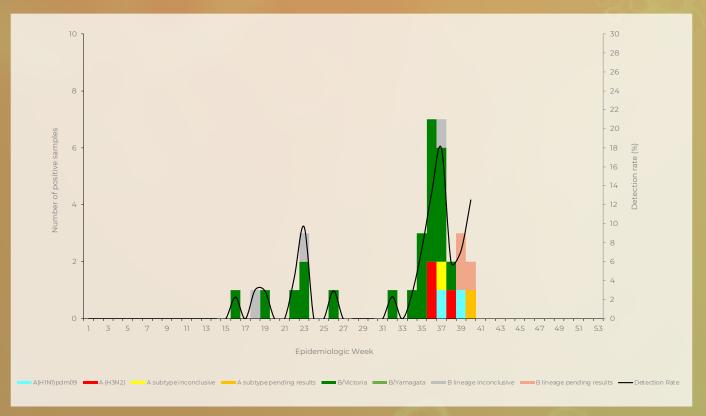


Figure 1. Number of influenza positive cases* by influenza subtype and lineage** and detection rate*** by week, Influenza-like illness (ILI) surveillance in primary health care clinics, 04/01/2021 – 10/10/2021

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

**Influenza B (Victoria) was detected in 12 (2%) of 577 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.

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

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

Table 1. Cumulative 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 – 10/10/2021

Clinic (Province)	A(H1N1) pdm09	A(H3N2)	A subtype inconclusive	A subtype pending results ^g	B/ Victoria	B/ Yamagata	B lineage inconclusive	B lineage pending results [®]	Total samples
Agincourt (MP)	О	0	0	0	0	0	0	0	198
Eastridge (WC)	О	2	0	0	5	Ο	0		213
Edendale Gateway (KZ)	0		Ο	0		0	2	0	184
Jouberton (NW)	2	О			10	0			683
Mitchell's Plain (WC)	0	0	0	0	2	0	0	1	176
Total:	2	3	1	1	21	0	3	3	1 454

KZ: KwaZulu-Natal; NW: North West; WC: Western Cape; MP: Mpumalanga Inconclusive: insufficient viral load in sample and unable to characterise further

**Influenza B (Victoria) was detected in 12 (2%) of 577 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. #influenza A subtype or B lineage results are pending

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PAGE 4

WEEK **40** 2021

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

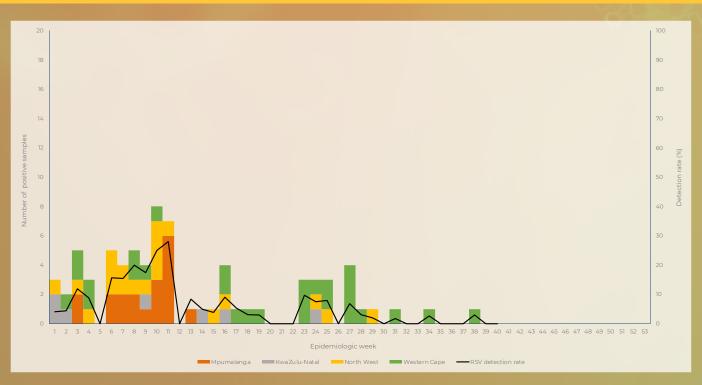


Figure 2. Number of patients testing positive for respiratory syncytial virus* by province and detection rate** by week, Influenzalike illness (ILI) surveillance in primary health care clinics, 04/01/2021 – 10/10/2021

**RSV was detected from 15 of 577 (3%) 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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PAGE 5

WEEK 40 2021

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

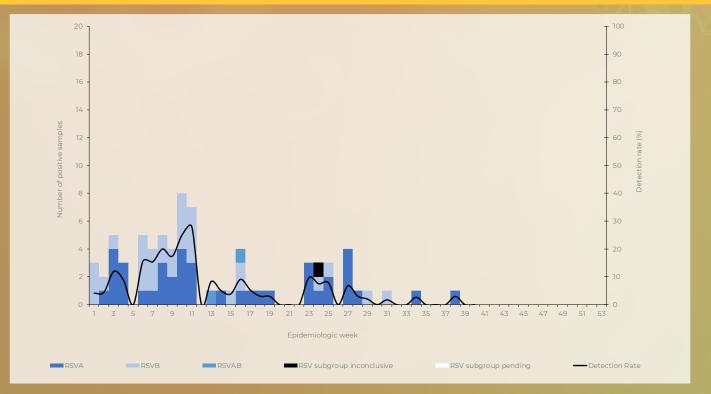


Figure 3. Number of patients testing positive for respiratory syncytial virus** by subgroup and detection rate by week,

 Table 2. Cumulative 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 –
 10/10/2021

Clinic (Province)	RSVA	RSVB	RSVAB	RSV subgroup inconclusive	RSV subgroup pending*	Total samples
Agincourt (MP)	11	7		0	Ο	198
Eastridge (WC)	23			О	Ο	213
Edendale Gateway (KZ)		6	0	Ο	Ο	184
Jouberton (NW)	2	17	0		Ο	683
Mitchell's Plain (WC)	3	0	0	0	О	176
Total	40	31	2	1	0	1 454

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 577 (3%) 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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PAGE 6

WEEK **40** 2021

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

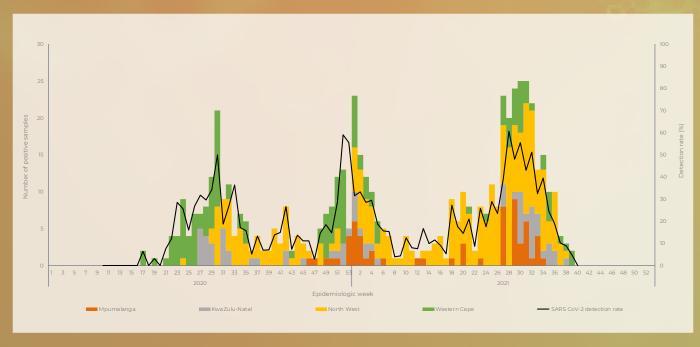


Figure 4. Number of patients testing positive for SARS-CoV-2* by province and detection rate by week, Influenza-like illness (ILI) surveillance in primary health care clinics, 02/03/2020 – 10/10/2021

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

**SARS-CoV-2 was detected in 148 of 709 (21%) 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. Cumulative 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 – 10/10/2021

Clinic (Province)	SARS-CoV-2 positive	Total samples tested
Agincourt (MP)	62	228
Eastridge (WC)	59	702
Edendale Gateway (KZ)	68	327
Jouberton (NW)	288	1003
Mitchell's Plain (WC)	89	505
Total:	566	2 765

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

**SARS-CoV-2 was detected in 148 of 709 (21%) 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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PAGE 7

WEEK **40** 2021

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

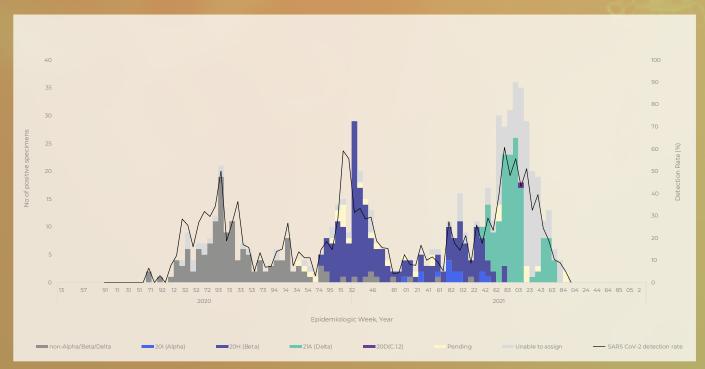


Figure 5. Number and detection rate of laboratory confirmed SARS-CoV-2* cases by variant type (variant PCR/sequencing) and week, Influenza-like illness (ILI) surveillance in primary health care clinics, 02/03/2020 – 10/10/2021

*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. Cumulative 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 – 10/10/2021

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
Agincourt (MP)	3	0	29	17	0	7	31	87
Eastridge (WC)	20	Ο	17	7	0	3	13	60
Edendale Gateway (KZ)	27	Ο	22	31	О	3	29	112
Jouberton (NW)	67	15	104	73		14	89	363
Mitchell's Plain (WC)	35	0	27	11	О	3	16	92
Total:	152	15	199	139	1	30	178	714

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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PAGE 8

WEEK 40 2021

INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

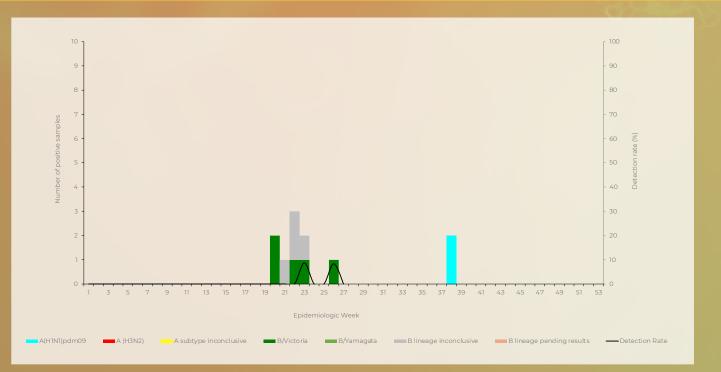


Figure 6. Number of positive patients* by influenza subtype and lineage and detection rate** by week, ILI surveillance - Viral

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 - 10/10/2021

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	1	0	0	0	1
Free State	О	О	0	Ο	0	О	О	0	Ο
Gauteng	О	2	0	0		О	4	0	146
Limpopo	Ο	О	0	0	Ο	О	О	0	0
Mpumalanga	0	О	0	0	0	О	О	Ο	3
North West	0	О	0	0	0	О	О	0	2
Northern Cape	0	0	О	Ο	Ο	0	0	О	Ο
Western Cape	0	0	О	0	0	0	0	0	28
Total:	0	2	0	0	5	0	4	0 /	180

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

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PAGE 9

WEEK 40 2021

INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

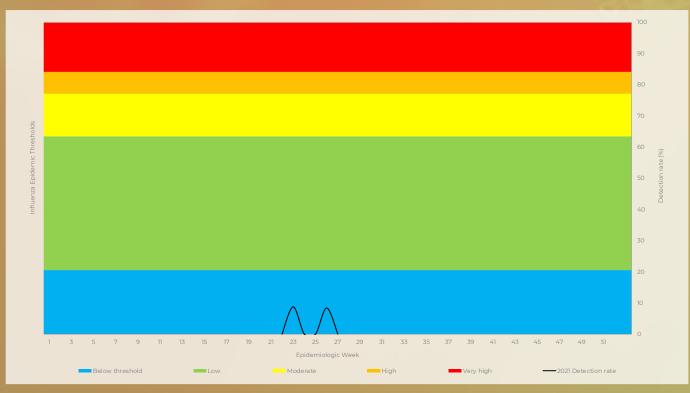


Figure 7. Influenza percentage detections and epidemic thresholds*, ILI surveillance - Viral Watch, 04/01/2021 - 10/10/2021

*Thresholds based on 2010-2019 data



WEEK 40 2021

INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

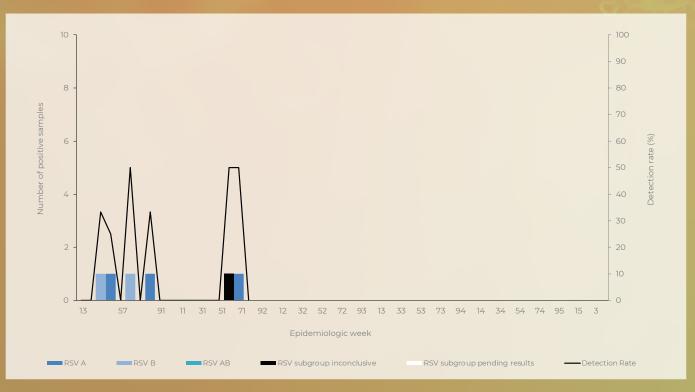


Figure 8. Number of RSV positive cases testing positive for respiratory syncytial virus (RSV)* by subgroup and detection rate by week, ILI surveillance - Viral Watch, 04/01/2021 – 10/10/2021

*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 – 10/10/2021

Province	RSV A	RSV B	RSV AB	RSV subgroup inconclusive	RSV subgroup pending results*	Total samples tested
Eastern Cape	0	0	0	0	0	41
Free State	О	О	0	0	0	О
Gauteng	2	2	О	Ο	0	146
Limpopo	О	О	0	0	0	0
Mpumalanga	О	0	0	О	0	3
North West	0	0	0	0	0	2
Northern Cape	0	Ο	О	0	0	О
Western Cape		о	0	1	0	28
Total:	3	2	0	1	0	180

*RSV results for subgroups are pending

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

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PAGE 11

WEEK 40 2021

INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

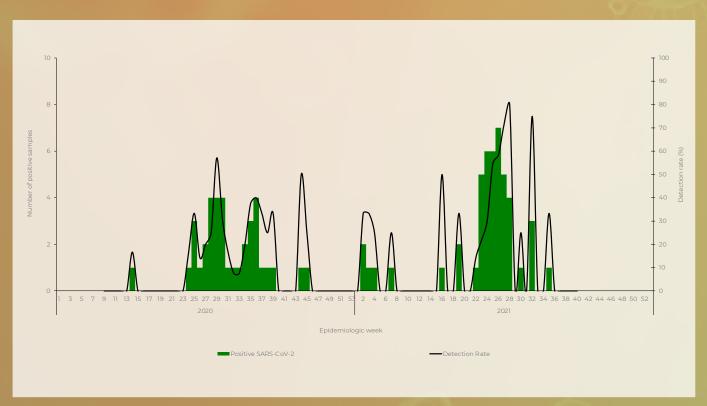


Figure 9. Number of patients testing positive for SARS-CoV-2*, by site and detection rate by week, ILI surveillance - Viral Watch, 04/01/2021 – 10/10/2021

*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, 04/01/2021 – 10/10/2021

Province	SARS-CoV-2 positive	Total samples tested				
Eastern Cape		5				
Free State		14				
Gauteng	65	306				
Limpopo	Ο	2				
Mpumalanga		8				
North West	Ο	2				
Northern Cape	Ο	2				
Western Cape	16	120				
Total:	84	459				

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PAGE **12**

WEEK 40 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, 04/01/2021 – 10/10/2021

*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

 Table 8. 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, 04/01/2021 - 10/10/2021

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
Eastern Cape	0	0	0	0	0	1	0	1 - N
Free State	0	О	0	0	Ο		0	
Gauteng	2	О	0	18	О	45	0	65
Limpopo	0	О	0	0	О	О	0	0
Mpumalanga	0	О	0	0	О		0	
North West	0	О	0	0	О	0	0	0
Northern Cape	0	О	0	Ο	О	0	0	0
Western Cape	0	0	0		О	1	0	16
Total:	2	0	0	19	0	63	0	84

*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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PAGE **13**

WEEK 40 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

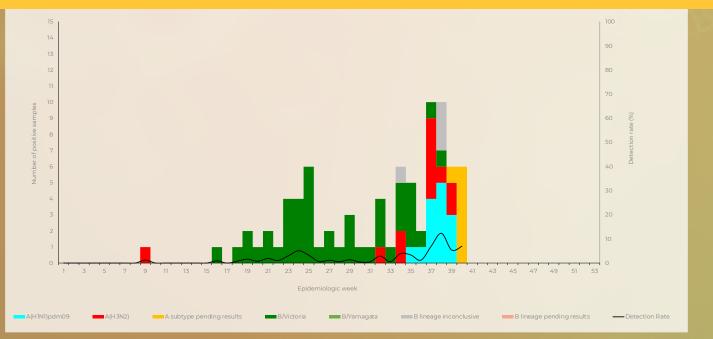


Figure 11. Number of positive influenza positive cases* by influenza subtype and lineage** and detection rate*** by week, pneumonia surveillance public hospitals, 04/01/2021 – 10/10/2021

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

**Influenza B(Victoria) was detected from four of 168 (2%) 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.

***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 – 10/10/2021

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)	0	0	0	0	9	0	0	0	757
Helen Joseph- Rahima Moosa (GP)	12	9	0	6	17	0	3	Ο	1275
Klerksdorp- Tshepong (NW)		0	Ο		4	0		0	799
Mapulaneng- Matikwana (MP)		0	Ο	0		0	Ο	0	480
Red Cross (WC)	0		0	0	12	Ο	0	0	635
Mitchell's Plain (WC)	0	2	Ο	0		0	Ο	0	759
Tintswalo (MP)	0	0	0	0	1	0	0	0	196
Total:	14	12	0	7	45	0	4	0	4 901

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 B(Victoria) was detected from four of 168 (2%) 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.
***influenza A subtype or B lineage results are pending

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PAGE **14**

WEEK 40 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

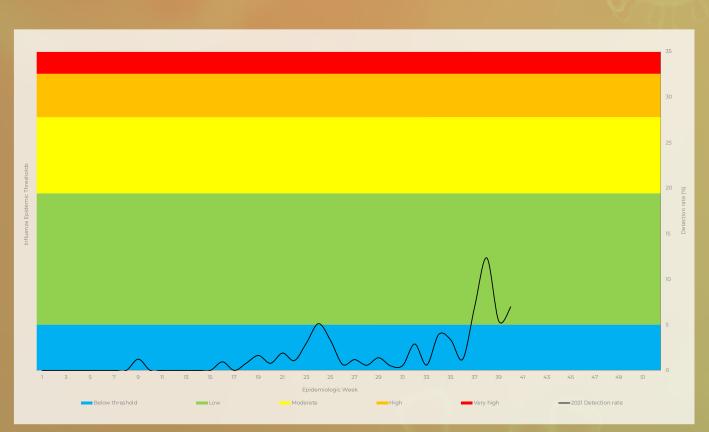


Figure 12. Influenza percentage detections and epidemic thresholds*, pneumonia surveillance public hospitals, 04/01/2021 – 10/10/2021

*Thresholds based on 2010-2019 data

www.nicd.ac.za TOLL-FREE NUMBER 0800 029 999

PAGE **15**

WEEK **40** 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

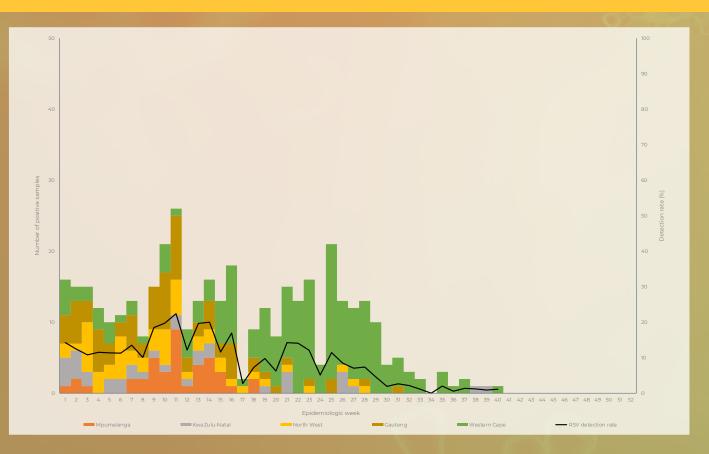


Figure 13. Number of patients testing positive for respiratory syncytial virus* by province and detection rate by week, pneumonia surveillance public hospitals, 04/01/2021 – 10/10/2021

*RSV was detected in six of 168 (4%) 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.

www.nicd.ac.za TOLL-FREE NUMBER 0800 029 999

PAGE **16**

WEEK **40** 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

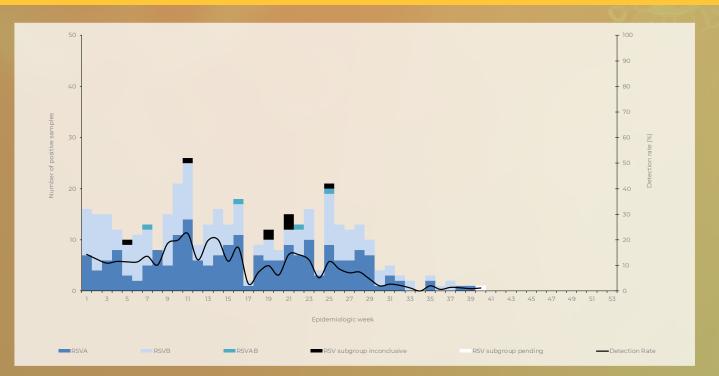


Figure 14. Number of patients testing positive for respiratory syncytial virus* by subgroup and detection rate by week, pneumonia surveillance public hospitals, 04/01/2021 – 10/10/2021

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)	7	29	0	Ο	0	757
Helen Joseph-Rahima Moosa (GP)	37	42	0	О	О	1275
Klerksdorp-Tshepong (NW)	8	44	0	2	О	799
Mapulaneng-Matikwana (MP)	20	7		0	О	480
Red Cross (WC)	84	47	2	5	О	635
Mitchell's Plain (WC)	40	11				759
Tintswalo (MP)	10	3	0	0	0	196
Total:	206	183	4	8		4 901

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 168 (4%) 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

TOLL-FREE NUMBER 0800 029 999 www.nicd.ac.za

WEEK 40 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

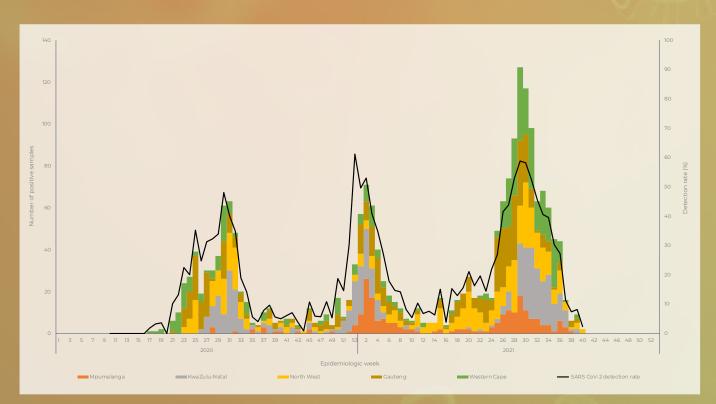


Figure 15. Number of patients testing positive for SARS-CoV-2*⁶ by province and detection rate by week, pneumonia surveillance public hospitals, 02/03/2020 – 10/10/2021

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

*#SARS-CoV-2 was detected in 42 of 239 (18%) 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 – 10/10/2021

Hospital (Province)	SARS-CoV-2 positive	Total samples tested		
Edendale (KZ)	458	1 600		
Helen Joseph-Rahima Moosa (GP)	483	2 120		
Klerksdorp-Tshepong (NW)	476	1 392		
Mapulaneng-Matikwana (MP)	165	782		
Red Cross (WC)	61	1 599		
Mitchell's Plain (WC)	348	1 228		
Tintswalo (MP)	51	196		
Total:	2 042	8 917		

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

**SARS-CoV-2 was detected in 42 of 239 (18%) 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.

www.nicd.ac.za TOLL-FREE NUMBER 0800 029 999

PAGE **18**

WEEK 40 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

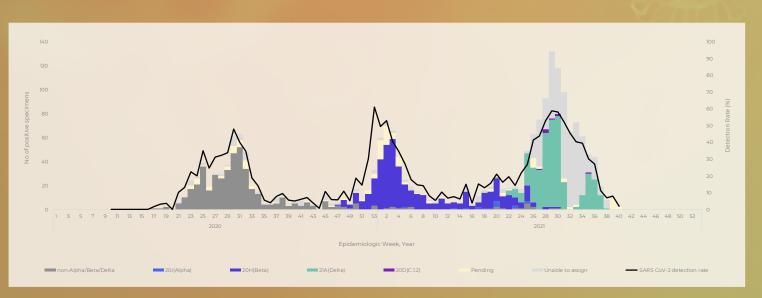


Figure 16. Number and detection rate of laboratory confirmed SARS-CoV-2 cases* by variant type (variant PCR/sequencing),

*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

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 – 10/10/2021

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	57	145	465
Helen Joseph-Rahima Moosa (GP)	134	5	120	101	4	19	101	484
Klerksdorp-Tshepong (NW)	130	9	112	94	2	13	116	476
Mapulaneng- Matikwana (MP)	16	0	90	29	0	17	47	199
Red Cross (WC)	15	О	5	7	О	7	27	61
Mitchell's Plain (WC)	50	0	49	96	О	14	139	348
Tintswalo (MP)	0	1	12	15	Ο	5	18	51
Total:	447	16	463	425	8	132	593	2084

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

TOLL-FREE NUMBER 0800 029 999 www.nicd.ac.za

PAGE 19

WEEK **40** 2021

SUMMARY OF LABORATORY CONFIRMED SARS-COV-2 CASES

Table 10. Characteristics of individuals with laboratory-confirmed SARS-CoV-2, enrolled in influenza-like illness (ILI) and pneumonia surveillance programmes, South Africa, 2 March 2020 - 10 October 2021

Characteristic	Influenza–like illness (ILI), public-sector, n=714 (%)	Pneumonia, n=2 084 (%)		
Age group				
0-9	52/714(7)	138/2084 (6)		
10-19	53/714 (7)	15/2084 (1)		
20-39	334/714 (47)	376/2084 (18)		
40-59	220/714 (31)	784/2084 (38)		
60-79	53/714 (7)	695/2084 (33)		
≥80	2/714 (<1)	76/2084 (4)		
Sex-female	431/714 (60)	1272/2084 (61)		
Province*				
Gauteng	N/A	484/2084 (23)		
KwaZulu-Natal	112/714 (16)	465/2084 (22)		
Mpumalanga**	87/714 (12)	250/2084 (12)		
North West	363/714 (51)	476/2084 (23)		
Western Cape	152/714 (21)	409/2084 (20)		
Race				
Black	537/707 (76)	1640/2070 (79)		
Coloured	139/707 (20)	325/2070 (16)		
Asian/Indian	4/707 (1)	50/2070 (2)		
White	22/707 (3)	41/2070 (2)		
Other	5/707 (1)	14/2070 (<1)		
Presentation				
Fever	555/707 (79)	961/2070 (46)		
Cough	688/707 (97)	1985/2070 (96)		
Shortness of breath	256/707 (36)	1620/2070 (78)		
Chest pain	308/707 (44)	842/2070 (41)		
Diarrhoea	54/707 (8)	132/2070 (6)		

www.nicd.ac.za TOLL-FREE NUMBER 0800 029 999

PAGE **20**

WEEK **40** 2021

Characteristic	Influenza-like illness (ILI), public-sector, n=714 (%)	Pneumonia, n=2 084 (%)
Underlying conditions		
Hypertension ^{\$}	17/690 (2)	249/1895 (13)
Cardiac	2/707 (<1)	47/2070 (2)
Lung disease	0/707 (0)	3/2070 (<1)
Diabetes	17/707 (2)	433/2070 (21)
Cancer	2/707 (<1)	13/2070 (1)
Tuberculosis	11/707 (2)	53/2070 (3)
HIV-infection	132/707 (19)	452/2070 (22)
Other ***	12/704 (2)	90/2054 (4)
Management		
Oxygen therapy	9/707 (1)	1595/2070 (77)
ICU admission	N/A	46/2070 (2)
Ventilation	N/A	53/2070 (3)
Outcome***		
Died	0/700 (0)	339/1969 (17)

**Mpumalanga (ILI site started enrolling on the 10th November 2020 and an additional SARI site started enrolling on the 10th February 2021)

***Chronic lung, liver and kidney disease, organ transplant, pregnancy, malnutrition, obesity, tracheostomy, prematurity, seizure, stroke, anaemia, asplenia, burns, Systemic lupus erythemate

⁵Data 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. Of the 339 patients who died, three were in <20 age group, 25 in the 20-39 year age group, 113 in the 40-59 year age group, and 196 were ≥60 years; 194/339 (57%) were female.

Of the 1159 hospitalised cases with COVID19 vaccine status, 96 (8%) received vaccine. 11 (11%) received Johnson & Johnson, 74 (77%) received the first dose of Pfizer-BioNTech, 3 (3%) Astra Zeneca and 8 (8%) could not remember the vaccine type received. 18 (19%) of the 96 cases had full vaccination (n=11, J&J and n=7, Pfizer-BioNTech). Cases age varied from 35 years to >60 years old and the majority (72%, 69/96) were >60 years old. Females were a majority 57% (55/96) and 11% (11/96) cases were HIV infected.

www.nicd.ac.za TOLL-FREE NUMBER 0800 029 999



WEEK **40** 2021

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 COVIDSeq 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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PAGE **22**