WEEKLY RESPIRATORY PATHOGENS SURVEILLANCE REPORT



SOUTH AFRICA WEEK 45 2021

CONTENTS

Surveillance programme description	2
Comments	3
Systematic Influenza-like illness (ILI) surveillance Influenza Respiratory syncytial virus SARS-CoV-2	4-9
Influenza-like illness (ILI) Viral Watch Influenza SARS-CoV-2	10-13
National syndromic surveilance for pneumonia Influenza Respiratory syncytial virus SARS-CoV-2	14-19
Summary of laboratory confirmed SARS-CoV-2 cases	20-21
SARS-CoV-2 Testing Methods	22

HIGHLIGHTS: WEEK 45

- · The 2021 influenza season has not yet started although sustained detections of influenza continue in all surveillance programmes. In week 45, transmission is below threshold and impact is low.
- 16 new cases of influenza from Western Cape (n=6), North West (n=7), Kwa-Zulu Natal (n=1) and Mpumalanga (n=2) surveillance sites were detected in week 45. To date, 259 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=33/99, 33%) and pneumonia surveillance (n=46/121, 38%).
- · 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=40/74, 54%) and pneumonia surveillance (n=213/408,
- From 2 March 2020 to date, a total of 2 910 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 2 043 hospitalised COVID-19 cases reported with available data on outcome, 351 (17%) died.
- From 1 January 2021 to date, of the 1192/1291 (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 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.

WEEK **45** 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	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: • 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
Specimens	Any person presenting with an acute (\$14 days) respiratory tract infection or other clinical illness compatible with COVID-19 [§] Oropharyngeal & nasopharyngeal	Any person presenting with an acute (≤14 days) respiratory tract infection or other clinical illness compatible with COVID-19 [®] Throat and/or nasal swabs or	Any person admitted with a physician- diagnosis of suspected COVID-19 and not meeting SRI case definition. Oropharyngeal & nasopharyngeal
collected	swabs	Nasopharyngeal swabs	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	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	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
	1 April 2020 – 31 March 2021: Roche E gene real-time PCR essay (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 2020 – 31 March 2021: Roche E gene real-time PCR essay 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)	gene real-time PCR essay (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)

WEEK **45** 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 season al threshold, Low activity, Moderate activity, High activity, Very high activity. For influenza, thresholds from outpatient influenza like illness (Viral Watch activity) and the context of theProgramme) are used as an indicator of disease transmission in the community and thresholds from pneumonia surveillance are used as an indicator of

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

- ****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

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 259 influenza cases detected in surveillance sites in 2021, the majority (n=101, 39%) were influenza B(Victoria). In week 45, transmission is below threshold and impact is low.

ILI programme: In 2021 to date, specimens from 1 688 patients meeting ILI case definition were received from 4 ILI sites. Influenza was detected in 99 (6%) patients, of which 38 (38%) were influenza A(H1N1)pdm09, 12 (12%) influenza A(H3N2), one (1%) influenza A(inconclusive), 9 (9%) influenza A(pending subtype results), 33 (33%) influenza B(Victoria), four (4%) influenza B(lineage inconclusive) and two (2%) influenza B(lineage pending results). (Fig1, Table1).

Viral Watch programme: In 2021 to date, specimens were received from 196 patients from Viral Watch sites in 5 of the 8 provinces participating in surveillance. Influenza was detected in 17 (9%) patients, of which two (12%) were influenza A(H1N1)pdm09, five (29%) influenza A(pending results), five (29%) influenza B(Victoria), four (24%) influenza B(lineage inconclusive) and one (6%) influenza B(lineage pending results). (Fig7, Table5)

Pneumonia surveillance: Since the beginning of 2021, specimens from 5 400 patients with severe respiratory illness (SRI) were received from the 6 sentinel sites. Influenza was detected in 121 (2%) patients, of which 32 (26%) were influenza A(H1N1)pdm09, 21 (17%) influenza A(H3N2), three (2%) influenza A(subtype inconclusive), 11 (9%) influenza A(pending subtype results), 46 (38%) influenza B(Victoria), seven (6%) were influenza B(lineage inconclusive) and one (1%) influenza B(lineage pending results). (Fig12, Table9)

In addition, influenza was detected in 22 (3%) of 801 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 509 RSV cases detected in 2021, the majority (n=263, 52%) were RSV subgroup A.

ILI programme: In 2021 to date, 1 688 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). (Fig4, Table2)

Viral Watch programme: In 2021 to date, 196 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, 5 400 specimens were tested and RSV was detected in specimens of 408 (8%) patients. Of which, 213 (52%) were RSV subgroup A, 190 (47%) RSV subgroup B, four (1%) RSV (subgroup inconclusive) and one (0.2%) RSV (subgroup pending). (Fig14, Table10)

In addition, RSV was detected in 21 of 801 (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, 3 002 patients were tested and SARS-CoV-2 was detected in 571 (19%) patients. From 1 January 2021 to date, of the 321/340 (94%) with data on variant type, majority (157/321, 49%) were Beta variant which predominated from weekl to week 24, followed by Delta (139/321, 43%) variant which predominated from week 25 to week 36. (Fig6, Table4)

Viral Watch programme: From March 2020 to date, 475 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. (Fig11, Table8)

Pneumonia surveillance: From March 2020 to date, 9 415 patients with severe respiratory illness (SRI) were tested and SARS-CoV-2 was detected in 2 061 (22%) patients. From 1 January 2021 to date, of the 871/954 (91%) with data on variant type, majority were (425/871, 49%) were Delta variant which dominated from week 22 to week 38 followed by Beta (401/871, 46%) variant which dominated from week 1 to 25. (Fig17, Table12)

In addition, SARS-CoV-2 was detected in 194 of 1004 (19%) specimens from patients who met suspected SARS-CoV-2 case definition but did not meet the pneumonia/ILI surveillance case definitions.

WEEK **45** 2021

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

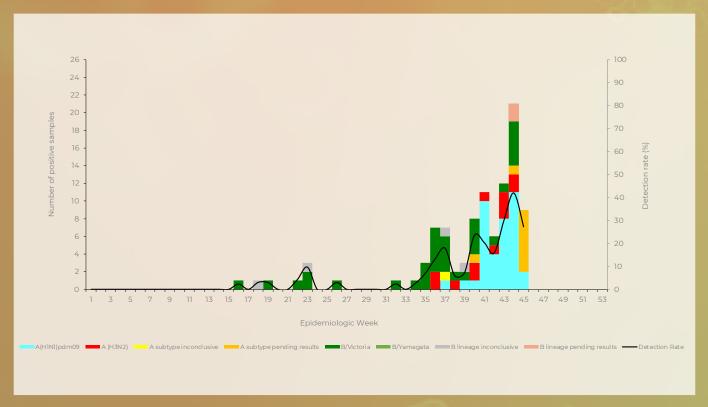


Figure 1. Number of influenza positive cases* by influenza subtype and lineage** and detection rate*** by week, Influenza-like illness (ILI) surveillance in primary health care clinics, 04/01/2021 – 14/11/2021

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, 04/01/2021 - 14/11/2021

Clinic (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
Agincourt (MP)			0	0	0	0	0	0	230
Eastridge (WC)			0		6	0	0	0	231
Edendale Gateway (KZ)	0		0	0	12	0	2	2	233
Jouberton (NW)	35			2	12	0		0	747
Mitchell's Plain (WC)	MIL	2	0	4	3	0	1	0	247
Total:	38	12	1	9	33	0	4	2	1 688

**Influenza was detected in 16 (3%) of 617 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. Dinfluenza A subtype or B lineage results are pending

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WEEK **45** 2021

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

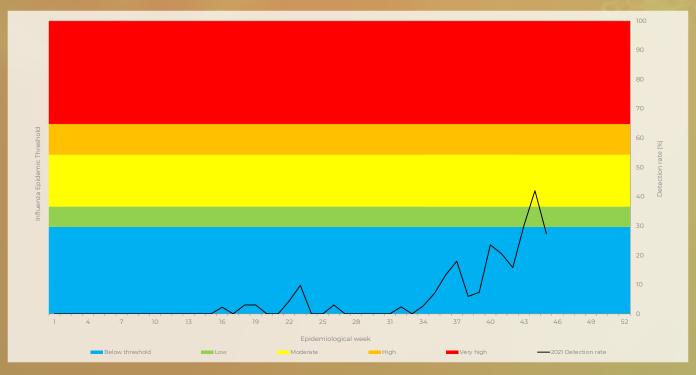


Figure 2. Influenza percentage detections and epidemic thresholds*, Influenza-like illness (ILI) surveillance in primary health care clinics, 04/01/2021 – 14/11/2021

*Thresholds based on 2012-2019 data

WEEK **45** 2021

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

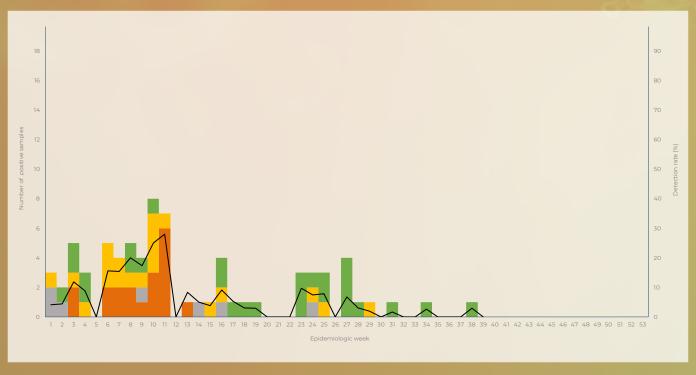


Figure 3. Number of patients testing positive for respiratory syncytial virus* by province and detection rate** by week, Influenza-like illness (ILI) surveillance in primary health care clinics, 04/01/2021 – 14/11/2021

**RSV was detected from 15 of 617 (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.

WEEK **45** 2021

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

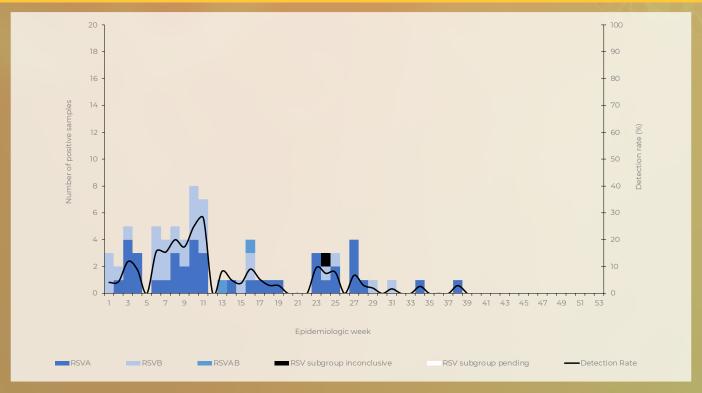


Figure 4. Number of patients testing positive for respiratory syncytial virus** by subgroup and detection rate by week,

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 - 14/11/2021

Clinic (Province)	RSVA	RSVB	RSVAB	RSV subgroup inconclusive	RSV subgroup pending*	Total samples
Agincourt (MP)	11	7		0	0	230
Eastridge (WC)	23			0	0	231
Edendale Gateway (KZ)		6	0	0	0	233
Jouberton (NW)	2	17	0		0	747
Mitchell's Plain (WC)	3	0	0	0	0	247
Total	40	31	2	1	0	1 688

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

These are not included in the table

WEEK **45** 2021

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

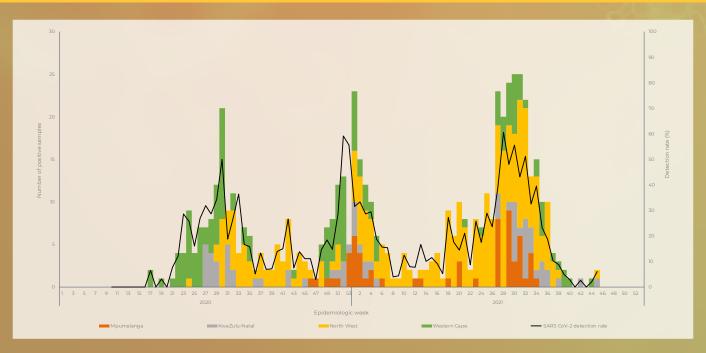


Figure 5. Number of patients testing positive for SARS-CoV-2* by province and detection rate by week, Influenza-like illness (ILI) surveillance in primary health care clinics, 02/03/2020 - 14/11/2021

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 – 14/11/2021

Clinic (Province)	SARS-CoV-2 positive	Total samples tested		
Agincourt (MP)	62	260		
Eastridge (WC)	60	720		
Edendale Gateway (KZ)	70	378		
Jouberton (NW)	289	1070		
Mitchell's Plain (WC)	90	576		
Total:	571	3 002		

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

^{*}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.

^{**}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 table.

WEEK **45** 2021

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

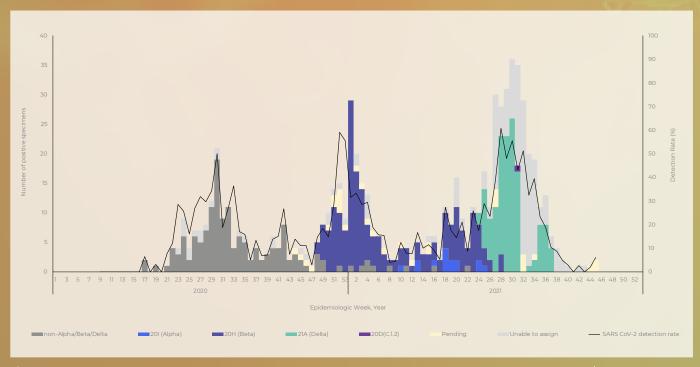


Figure 6. Number and detection rate of laboratory confirmed SARS-CoV-2* cases by variant type (variant PCR/sequencing) and week, Influenza-like illness (ILI) surveillance in primary health care clinics, 02/03/2020 – 14/11/2021

*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 – 14/11/2021

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	0	17	7	О	2	14	61
Edendale Gateway (KZ)	27	0	22	31	О	4	30	115
Jouberton (NW)	67	15	104	73		14	89	364
Mitchell's Plain (WC)	35	0	27	11	О	1	19	93
Total:	152	15	199	139	1	28	183	720

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

WEEK **45** 2021

INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

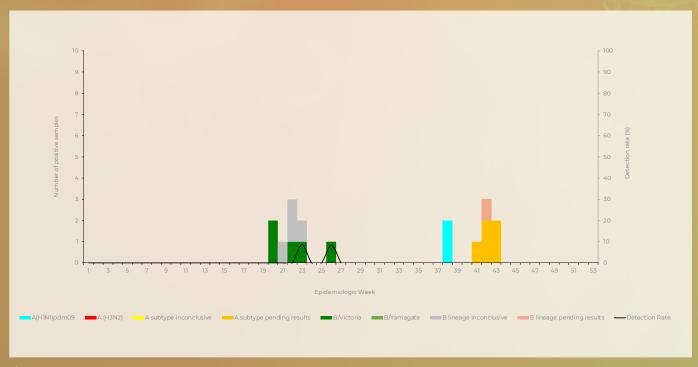


Figure 7. Number of positive patients* by influenza subtype and lineage and detection rate** by week, ILI surveillance - Viral Watch, 04/01/2021 – 14/11/2021

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

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 – 14/11/2021

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	Ο	0	0	0	0	0	О	2
Gauteng	2	О	0	4		0	4	0	154
Limpopo	О	О	0	0	О	0	0	0	0
Mpumalanga	О	О	0	0	0	0	0	0	3
North West	О	0	0	0	0	0	0	0	2
Northern Cape	0	0	0	0	О	0	0	0	0
Western Cape	О	0	0	1	О	0	0	О	31
Total:	2	0	0	5	5	0	4	1/	196

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

From 04 January 2021 to date, no patients were tested for influenza at the time of entry into South Africa following travel abroad.

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

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PAGE **10**

^{*}Specimens from patients with Influenza-like illnesses at 90 sentinel sites in 8 provinces

^{*} Only reported for weeks with >10 specimens submitted.

WEEK 45 2021

INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

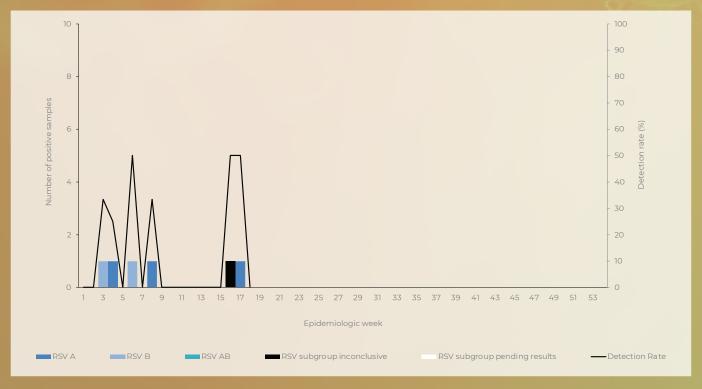


Figure 8. Number of RSV positive cases testing positive for respiratory syncytial virus (RSV)* by subgroup and detection rate by week, ILI surveillance - Viral Watch, 04/01/2021 - 14/11/2021

Table 6. Number of RSV positive cases identified and total number of samples tested by province, ILI surveillance - Viral Watch, 04/01/2021 – 14/11/2021

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	154
Limpopo	0	О	0	О	0	0
Mpumalanga	0	0	0	О	0	3
North West	0	0	0	О	0	2
Northern Cape	0	0	0	О	0	0
Western Cape		0	0		0	31
Total:	3	2	0	1	0	196

^{*}RSV results for subgroups are pending

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

^{*}RSV results for subgroups are pending

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

WEEK **45** 2021

INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

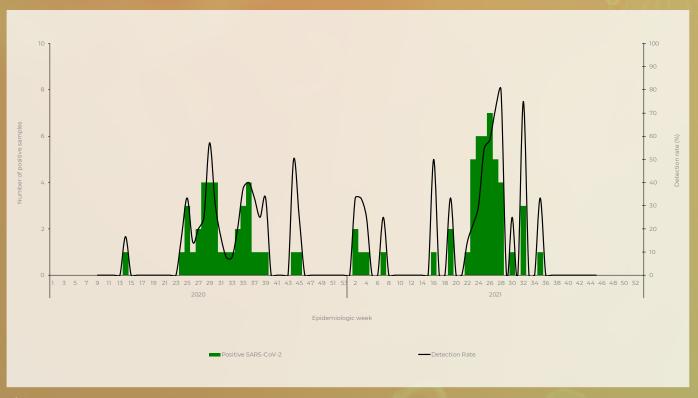


Figure 9. Number of patients testing positive for SARS-CoV-2*, by site and detection rate by week, ILI surveillance - Viral Watch, 02/03/2020 - 14/11/2021

Table 7. Number of SARS-CoV-2 positive cases identified and total number tested by province, ILI surveillance - Viral Watch, 02/03/2020 – 14/11/2021

Province	SARS-CoV-2 positive	Total samples tested
Eastern Cape		8
Free State		16
Gauteng	65	314
Limpopo	0	2
Mpumalanga		8
North West	0	2
Northern Cape	0	2
Western Cape	16	123
Total:	84	475

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

WEEK **45** 2021

INFLUENZA-LIKE ILLNESS (ILI) SURVEILLANCE VIRAL WATCH

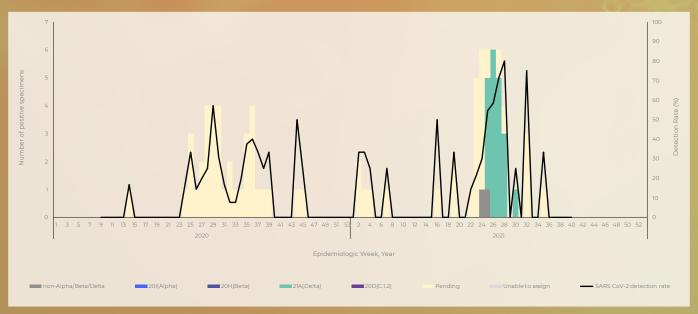


Figure 10. Number and detection rate of laboratory confirmed SARS-CoV-2* cases by variant type (variant PCR/sequencing) and week, ILI surveillance - Viral Watch, 02/03/2020 - 14/11/2021

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, 02/03/2020 – 14/11/2021

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
Eastern Cape	0	0	0	О	0		0	
Free State	О	0	0	О	0		0	
Gauteng	2	0	0	18	0	45	0	65
Limpopo	О	0	0	О	0	0	0	0
Mpumalanga	О	0	0	О	0		0	
North West	О	0	0	О	0	0	0	0
Northern Cape	О	0	О	О	0	0	0	0
Western Cape	О	0	0	1	0	15	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

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

WEEK **45** 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

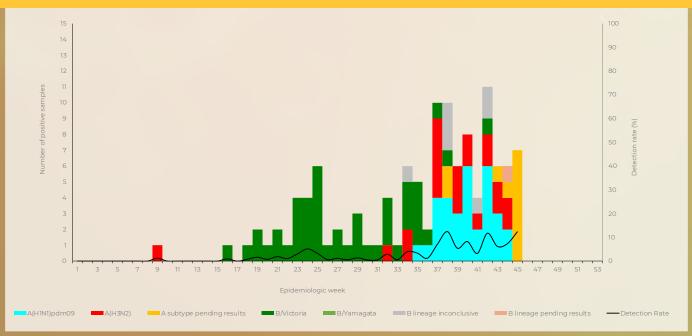


Figure 11. Number of positive influenza positive cases* by influenza subtype and lineage** and detection rate*** by week, pneumonia surveillance public hospitals. 04/01/2021 – 14/11/2021

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 - 14/11/2021

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)	0	1	0	- 1	9	О	0	1	837
Helen Joseph- Rahima Moosa (GP)	22	13		2	17	0	5	Ο	1433
Klerksdorp- Tshepong (NW)	6		0	5	4	0	2	0	863
Mapulaneng- Matikwana (MP)	2	0	0	2		0	О	0	535
Red Cross (WC)		3	0		12	0	О	0	690
Mitchell's Plain (WC)		3	0	0	2	0	Ο	0	817
Tintswalo (MP)	0	0	2	0	1	0	0	0	225
Total:	32	21	3	11	46	0	7	ī	5 400

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

**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.

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

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TOLL-FREE NUMBER 0800 029 999

PAGE **14**

^{*}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(HINI)pdm09 and four (67%) were influenza B(Victoria) from patients who me 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

WEEK **45** 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA



Figure 12. Influenza percentage detections and epidemic thresholds*, pneumonia surveillance public hospitals, 04/01/2021 – 14/11/2021

*Thresholds based on 2010-2019 data

WEEK **45** 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

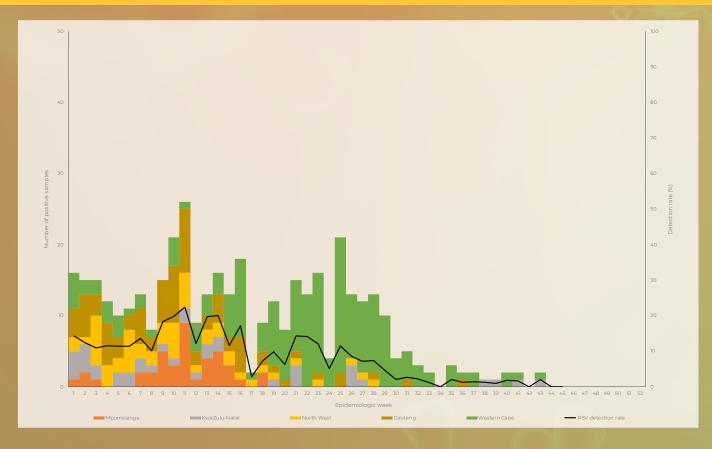


Figure 13. Number of patients testing positive for respiratory syncytial virus* by province and detection rate by week, pneumonia surveillance public hospitals, 04/01/2021 – 14/11/2021

*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.

WEEK **45** 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

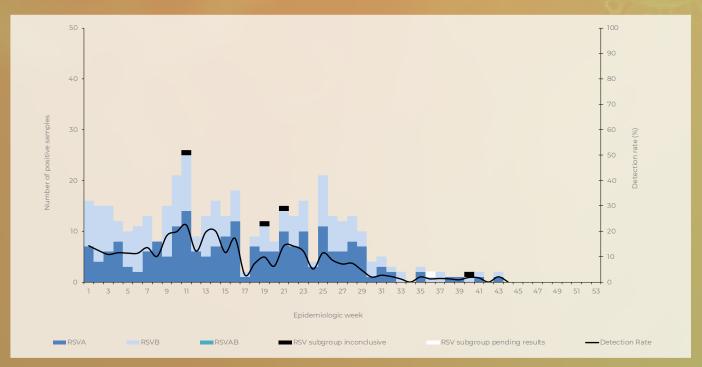


Figure 14. Number of patients testing positive for respiratory syncytial virus* by subgroup and detection rate by week, pneumonia surveillance public hospitals, 04/01/2021 – 14/11/2021

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 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

Table 10: Number of patients positive for respiratory syncytial virus subgroups** by subgroups identified and total number of samples tested by hospital, pneumonia surveillance public hospitals, 04/01/2021 – 14/11/2021

Hospital (Province)	RSVA	RSVB	RSVAB	RSV subgroup inconclusive	RSV subgroup pending*	Total samples
Edendale (KZ)	9	29	0	0		837
Helen Joseph-Rahima Moosa (GP)	37	42	О	0	0	1433
Klerksdorp-Tshepong (NW)	8	45	0		О	863
Mapulaneng-Matikwana (MP)	21	7	0	О	0	535
Red Cross (WC)	87	51	0	2	О	690
Mitchell's Plain (WC)	41	13	0		О	817
Tintswalo (MP)	10	3	0	0	0	225
Total:	213	190	0	4	1	5 400

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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TOLL-FREE NUMBER **0800 029 999**

PAGE **17**

WEEK **45** 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

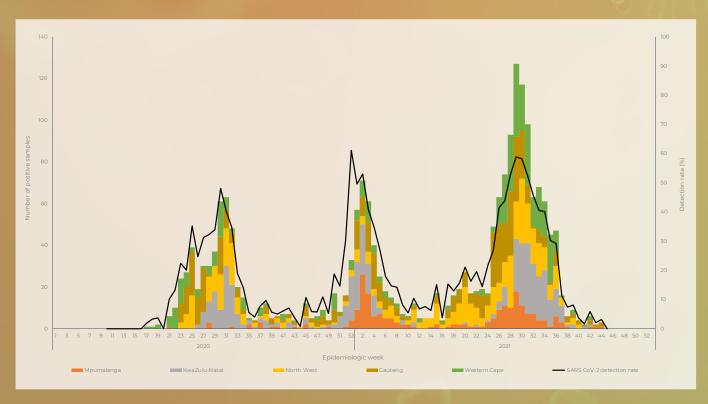


Figure 15. Number of patients testing positive for SARS-CoV-2*[§] by province and detection rate by week, pneumonia surveillance public hospitals, 02/03/2020 – 14/11/2021

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

*SARS-CoV-2 was detected in 45 of 255 (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 = 14/11/2021

Hospital (Province)	SARS-CoV-2 positive	Total samples tested	
Edendale (KZ)	460	1 684	
Helen Joseph-Rahima Moosa (GP)	493	2 279	
Klerksdorp-Tshepong (NW)	476	1 456	
Mapulaneng-Matikwana (MP)	167	837	
Red Cross (WC)	61	1648	
Mitchell's Plain (WC)	352	1 287	
Tintswalo (MP)	52	224	
Total:	2 061	9 415	

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

**SARS-CoV-2 was detected in 45 of 255 (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.

WEEK **45** 2021

NATIONAL SYNDROMIC SURVEILLANCE FOR PNEUMONIA

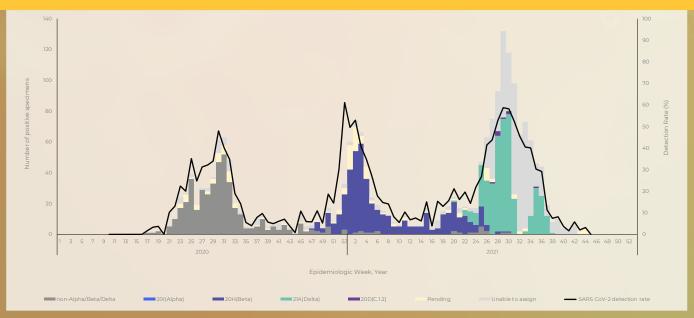


Figure 16. Number and detection rate of laboratory confirmed SARS-CoV-2 cases* by variant type (variant PCR/sequencing), pneumonia surveillance public hospitals, 02/03/2020 – 14/11/2021

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 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 – 14/11/2021

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	55	149	467
Helen Joseph-Rahima Moosa (GP)	134	5	120	101	4	25	105	494
Klerksdorp-Tshepong (NW)	130	9	112	94	2	12	117	476
Mapulaneng- Matikwana (MP)	16	0	90	29	0	18	49	202
Red Cross (WC)	15	0	5	7	0	7	27	61
Mitchell's Plain (WC)	50	0	49	96	0	15	144	354
Tintswalo (MP)	0	1	12	15	0	6	18	52
Total:	447	16	463	425	8	138	609	2 106

GP: 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

WEEK **45** 2021

SUMMARY OF LABORATORY CONFIRMED SARS-COV-2 CASES

Table 13. Characteristics of individuals with laboratory-confirmed SARS-CoV-2, enrolled in influenza-like illness

Characteristic	Influenza-like illness (ILI), public-sector, n=720 (%)	Pneumonia, n=2 106 (%)		
Age group				
0-9	53/720 (7)	138/2106 (7)		
10-19	53/720 (7)	15/2106 (1)		
20-39	336/720 (47)	386/2106 (18)		
40-59	223/720 (31)	786/2106 (37)		
60-79	53/720 (7)	703/2106 (33)		
≥80	2/720 (<1)	78/2106 (4)		
Sex-female	435/720 (60)	1286/2106 (61)		
Province*				
Gauteng	N/A	494/2106 (23)		
KwaZulu-Natal	115/720 (16)	467/2106 (22)		
Mpumalanga**	87/720 (12)	254/2106 (12)		
North West	364/720 (51)	476/2106 (23)		
Western Cape	154/720 (21)	415/2106 (20)		
Race				
Black	542/714 (76)	1668/2097 (79)		
Coloured	141/714 (20)	327/2097 (16)		
Asian/Indian	4/714 (1)	52/2097 (2)		
White	22/714 (3)	41/2097 (2)		
Other	5/714 (1)	9/2097 (<1)		
Variant ^{\$\$}				
Non-Alpha/Beta/Delta	152/535 (28)	447/1497 (30)		
201(Alpha)	15/535 (3)	16/1497 (1)		
20H(Beta)	199/535 (37)	463/1497 (31)		
21A(Delta)	139/535 (26)	425/1497 (28)		
20D(C.1.2)	1/535 (0.2)	8/1497 (1)		
Pending results	29/535 (5)	138/1497 (9)		

WEEK **45** 2021

Characteristic	Influenza-like illness (ILI), public-sector, n=720 (%)	Pneumonia, n=2 106 (%)		
Presentation				
Fever	559/714 (78)	971/2097 (46)		
Cough	695/714 (97)	2010/2097 (96)		
Shortness of breath	260/714 (36)	1647/2097 (79)		
Chest pain	312/714 (44)	856/2097 (41)		
Diarrhoea	54/714 (8)	135/2097 (6)		
Underlying conditions				
Hypertension ^s	17/691 (2)	249/1896 (13)		
Cardiac	2/714 (<1)	49/2097 (2)		
Lung disease	0/714 (0)	3/2097 (<1)		
Diabetes	17/714 (2)	438/2097 (21)		
Cancer	2/714 (<1)	13/2097 (1)		
Tuberculosis	11/714 (2)	57/2097 (3)		
HIV-infection	135/714 (19)	467/2097 (22)		
Other ***	12/714 (2)	90/2097 (4)		
Management				
Oxygen therapy	9/714 (1)	1652/2097 (79)		
ICU admission	N/A	50/2097 (2)		
Ventilation	N/A	55/2097 (3)		
Outcome***				
Died	0/711 (0)	351/2043 (17)		



WEEK **45** 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 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, (https://sars-cov-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).