
Comparison of COVID-19 Pandemic Waves in 10 Countries in Southern Africa, 2020–2021

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We used publicly available data to describe epidemiology, genomic surveillance, and public health and social measures from the first 3 COVID-19 pandemic waves in southern Africa during April 6, 2020–September 19, 2021. South Africa detected regional waves on average 7.2 weeks before other countries. Average testing volume (244 tests/million/day) increased across waves and was highest in upper-middle-income countries. Across the 3 waves, average reported regional incidence increased (17.4, 51.9, 123.3 cases/1 million population/day), as did positivity of

diagnostic tests (8.8%, 12.2%, 14.5%); mortality (0.3, 1.5, 2.7 deaths/1 million population/day); and case-fatality ratios (1.9%, 2.1%, 2.5%). Beta variant (B.1.351) drove the second wave and Delta (B.1.617.2) the third. Stringent implementation of safety measures declined across waves. As of September 19, 2021, completed vaccination coverage remained low (8.1% of total population). Our findings highlight opportunities for strengthening surveillance, health systems, and access to realistically available therapeutics, and scaling up risk-based vaccination.

As of September 2021, in Africa, 5,650,962 SARS-CoV-2 infections (2.6% of global total) and 135,568 related deaths (3.0% of global total), had been reported (1). However, this number was likely a substantial underestimate of the true number of SARS-CoV-2 infections, given limited surveillance capacity and relatively higher positivity reported in seroprevalence

studies (2–4). The first case in southern Africa, home to ≈14% of the population of Africa (5), was reported on March 5, 2020 in South Africa (6). By September 2021, all countries in southern Africa were experiencing their third COVID-19 pandemic waves.

Although quantitative comparisons of COVID-19 waves have been published, few have

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compared waves in southern Africa (7–9), despite the region experiencing substantial illness and death across waves (10). Furthermore, there has been limited systematic reporting and analysis of public health and social measures (PHSMs) enacted during outbreaks across countries in the region. A comparison of characteristics across waves provides unique insights into reported incidence, mortality, and distribution of variants of concern (VOCs) across geography and time. Population movements between countries in southern Africa, a highly interconnected region, have historically been drivers of HIV and tuberculosis epidemics (11) and could influence COVID-19 wave propagation. To inform public health actions to prevent, detect, and reduce the effects of future COVID-19 pandemic waves across the region, we compared trends in reported testing volume, incidence, mortality, genomic surveillance results, PHSMs, and vaccination coverage across pandemic waves in southern Africa during April 2020–September 2021.

Methods

Data Sources and Data Collection

According to the African Union (<https://au.int>), southern Africa consists of Angola, Botswana, Eswatini, Lesotho, Malawi, Mozambique, Namibia, South Africa, Zambia, and Zimbabwe. We obtained data on testing, incidence, mortality, and vaccination collected during February 7, 2020–September 19, 2021 (final day of data extraction) from the Our World in Data (OWID; <https://ourworldindata.org>) dataset, compiled by Johns Hopkins University (1). We supplemented missing data or errors with data from in-country US Centers for Disease Control and Prevention (CDC) offices, the World Health Organization (WHO), or daily reports from Africa Centers for Disease Control and Prevention (Africa CDC) (12). We excluded still-missing data from indicator computations and computed weekly averages for each indicator to reduce potential bias introduced by missed reports. We based the effective reproduction number on estimates published elsewhere (13). We obtained publicly available SARS-CoV-2 genomic sequencing results from GISAID (<https://www.gisaid.org>) (14); those data were exported on September 19, 2021, and included specimens collected during March 1, 2020–September 6, 2021.

We extracted publicly available PHSM data from the Oxford COVID-19 Government Response Tracker (OxCGRT; <https://covidtracker.bsg.ox.ac.uk>), available during January 1, 2020–September 19, 2021 (15).

OxCGRT contains 23 indicators aggregated into a set of common indices, rated 1–100 to quantify the level of government intervention. All indices, defined on the OxCGRT website, were based on averages of component indicators to provide a measure of how many indicators a government has acted upon and to what degree. We compared the original PHSM stringency, overall government response, containment health, and economic support indices across waves. This activity was reviewed by CDC and conducted consistent with applicable federal laws and CDC policy.

Statistical Analysis

To align with existing analysis of pandemic waves in Africa, we adapted wave definitions published elsewhere (6) (Appendix, <https://wwwnc.cdc.gov/EID/article/28/13/22-0228-App1.pdf>). Different authors independently applied these definitions to determine the wave start, peak, and end weeks (Appendix Table 1); we resolved discrepancies by consensus. We analyzed data in R version 4.01 (The R Foundation for Statistical Computing, <https://www.r-project.org>). We computed averages and maximums across wave periods and countries for reported COVID-19 incidence (7-day average daily cases and peak cases/1 million persons); mortality (7-day average daily deaths and peak deaths/1 million persons and case-fatality ratio [CFR]); testing (7-day average daily tests/1 million persons, 7-day average test positivity, peak 7-day average test positivity, and 7-day average tests per case); and vaccination (total number of persons vaccinated/100 population, total number persons fully vaccinated/100 persons [defined by OWID as total number of persons who received all doses prescribed by the vaccination protocol/100 persons in the total population], and average weekly vaccinations/1 million persons). We computed peak averages as the maximum 7-day average in a period; OWID defines peak 7-day average test positivity as tests conducted per new confirmed case. We computed regional averages for southern Africa by averaging all available country-specific values for each indicator within the wave period. For example, for each 7-day average indicator, we averaged all available country-level 7-day averages to determine overall regional averages, and all available 7-day averages within country-specific wave periods were averaged for regional averages by wave. We conducted 1-way analysis of variance tests to calculate differences in 7-day average cases, deaths, and tests per 1 million persons across waves. We computed genomic surveillance coverage as the total number of sequences submitted to GISAID during that period

divided by the number of cases per 1 million. However, for ease of interpretation, genomic surveillance coverage was reported as its inverse (number of cases/1 million/sequence submitted). Therefore, a country with a higher number of reported cases per 1 million per sample sequenced has lower genomic surveillance coverage than a country with a lower number. We computed medians and interquartile ranges (IQRs) across wave periods for continuous genomic variables and frequencies for categorical genomic variables. We reported genomic sequences using WHO genome labels (16) (Appendix Table 2) and categorized sequences without a WHO label as other lineages (Appendix Table 3). For PHSM data, we computed averages across waves for each index and frequencies for the number of measures mandated at the beginning, peak, end, and throughout the duration of waves.

Results

Burden of COVID-19 in Southern Africa

By September 19, 2021, southern Africa had 3,841,563 SARS-CoV-2 cases, 65.0% of Africa and 1.7% of global totals, and 107,347 COVID-19 deaths, 75.4% of Africa and 2.3% of global totals. South Africa had the highest numbers of cases (75.0%) and deaths (80.3%) among countries in the region. The countries with highest incidence and mortality over the period were Botswana, Namibia, Eswatini, and South Africa (Appendix Figure 1).

Regional Pandemic Wave Propagation Patterns

The earliest start date for the first wave within any country was April 6, 2020 (South Africa); by July 5, 2021, all countries in the region were experiencing a third wave (Figure 1). On average, pandemic waves in the region lasted 16.5 weeks; the first wave, at 19.5 weeks, was the longest, followed by the second, 15.1 weeks, and third, 14.9 weeks (Table 1). Wave durations varied by wave and across countries; the first wave in Angola lasted 30 weeks but the second wave in Zimbabwe lasted 9 weeks. Waves in almost all other countries started an average of 7.2 weeks later than in South Africa, but with some variation: Namibia at 4.0 weeks and Angola at 14.0 weeks later (Table 1).

Regional and Temporal Variations in Testing

The number of 7-day average daily tests per 1 million persons was higher in the 2 upper-middle-income countries, Namibia (549.0) and South Africa (519.3), where testing data were more available, but lower in low-income countries Malawi (37.9) and Mozambique (51.6) (Table 2, <https://wwwnc.cdc.gov/EID/article/28/13/28-0228-T2.htm>). Testing increased in all 10 countries across successive pandemic waves; the third wave had nearly 3 times (388.0 versus 146.8) the 7-day average daily tests per million persons than did the first wave. There was a statistically significant ($p < 0.05$) mean difference across waves in tests within each country and across all countries. However, 7-day average test/case ratio was highest in the first wave (24.8), followed by the second (17.0) and third (13.5) (Table 2).

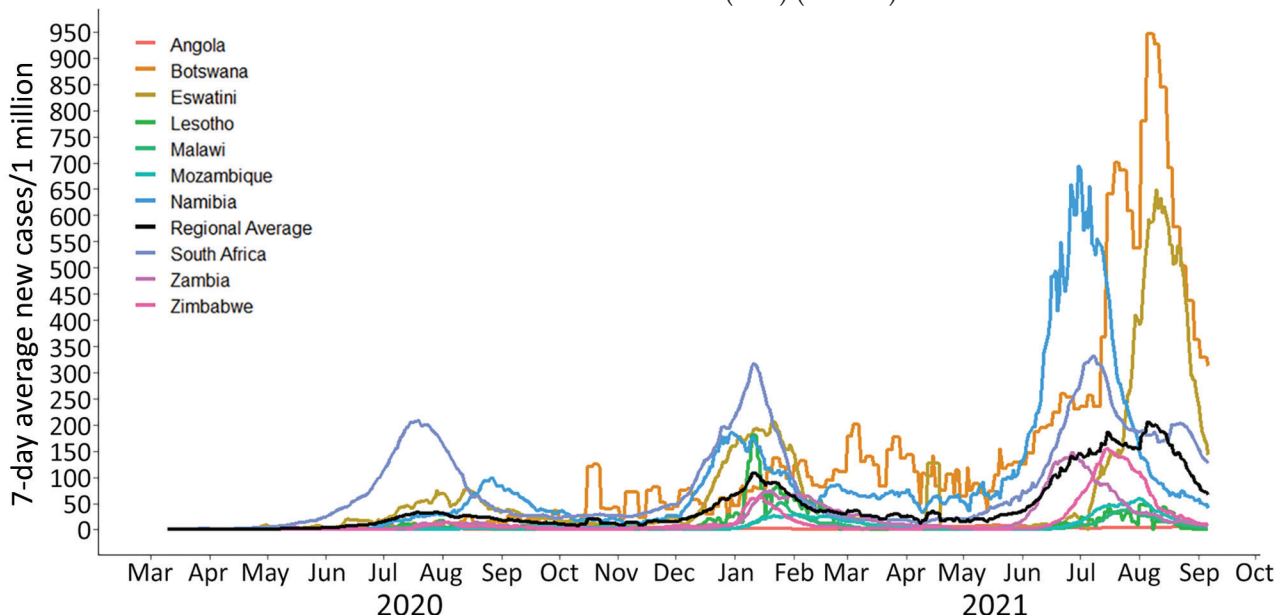


Figure 1. Reported 7-day average of new COVID-19 cases per 1 million population across 10 countries in southern Africa, March 5, 2020–September 17, 2021. Source: Our World in Data (<https://www.ourworldindata.org>), accessed 2021 Sep 20.

Table 1. Total duration of 3 COVID-19 pandemic waves in 10 countries in southern Africa and time since start of wave in South Africa, April 6, 2020–September 19, 2021*

Country	Wave 1		Wave 2		Wave 3		Country average	
	Total duration, wk	Time from start of SA wave, wk	Total duration, wk	Time from start of SA wave, wk	Total duration, wk	Time from start of SA wave, wk	Total duration, wk	Time from start of SA wave, wk
Angola†	30	13	13	20	10	9	17.7	14.0
Botswana	31	5	16	8	17	2	21.3	5.0
Eswatini	15	11	14	4	10	9	13.0	8.0
Lesotho	16	11	12	3	14	5	14.0	6.3
Malawi	15	10	18	5	15	4	16.0	6.3
Mozambique	20	12	19	6	16	3	18.3	7.0
Namibia	20	10	13	0	17	2	16.7	4.0
SA	22	Referent	16	Referent	19	Referent	19.0	Referent
Zambia	15	13	21	3	17	2	17.7	6.0
Zimbabwe	11	12	9	7	14	5	11.3	8.0
Overall average	19.5	10.8	15.1	6.2	14.9	4.6	16.5	7.2

*Appendix Table 1 (<https://wwwnc.cdc.gov/EID/article/28/12/22-0228-App1.pdf>) shows dates of starts, peaks, ends, and period definitions of pandemic waves. SA, South Africa.

†Third wave in Angola had not yet reached its peak as of September 19, 2021.

Temporal Changes in COVID-19 Wave Severity

Average incidence (cases/1 million persons/day) increased across waves, from 17.4 in the first to 5.19 in the second 51.9 and 123.3 in the third. Percentage test positivity increased from 8.8% in the first wave to 12.2% in the second and 14.5% in the third. Mortality (deaths/1 million persons/day) increased from 0.3 in the first wave to 1.5 in the second and 2.7 in the third. CFR increased from 1.9% in the first wave to 2.1% in the second and 2.5% in the third (Table 2).

In an unadjusted analysis that did not control for changes in testing capacity over time, we also found a statistically significant ($p < 0.05$) mean difference across waves in 7-day average daily cases and deaths per 1 million population within each country and the region. However, for some countries the second wave had the highest reported incidence of cases and deaths (Table 2; Figure 2). The second wave in Lesotho had the highest peak 7-day average number of new cases per 1 million persons and the highest peak in deaths per 1 million persons per day in Lesotho, South Africa, and Eswatini (Table 2). Upper middle-income countries South Africa, Namibia, and Botswana had relatively high overall 7-day average numbers of new deaths per 1 million persons compared with low-income countries.

Genomic Surveillance

During the study period, a collective 23,306 SARS-CoV-2 specimen sequences were submitted to GISAID from all 10 countries in southern Africa, most (89.4%) from laboratories in South Africa (Table 3; Appendix Figure 2). Most (18,464, 79.2%) specimens were collected in South Africa, the fewest (18, 0.1%) in Lesotho (Appendix Figure 3). The largest proportion of specimens (43.3%) were collected during the

third wave; the number of sequences submitted increased between the first and second waves in 8/10 countries (Table 4, <https://wwwnc.cdc.gov/EID/article/28/13/28-0228-T4.htm>; Figure 4).

Genomic surveillance coverage (median number of cases/1 million persons/SARS-CoV-2 genome submitted) varied across countries, from 1.02 (IQR 0.94–2.5) in Angola to 211.40 (IQR 210.7–486.4) in Eswatini (Table 4). For the southern Africa region, genomic surveillance coverage was highest before the start of the second wave, median 1.55 cases/1 million persons/SARS-CoV-2 genome submitted. The prevalence of the Beta variant increased from 13.7% in the period before the second wave to 80.6% during the second wave (Table 3). During the third wave, the prevalence of Beta decreased to 14.8% and the prevalence of Delta increased to 73.8%. Beta variant was predominant in the second wave in 8/10 countries and Delta in the third wave in 9/9 countries (Table 4; Figure 3).

PHSMs

PHSM stringency index decreased from the first through the third waves in 8/10 countries (Table 5, <https://wwwnc.cdc.gov/EID/article/28/13/28-0228-T5.htm>; Figure 5). Regionally, average stringency, government response, and economic support indices were highest during the first wave (Table 5). International travel restrictions were the most common PHSM and closing public transport the least common (Table 6). During the first wave, more PHSMs were implemented at the beginning of the wave than at the end, whereas during the second wave, more PHSMs were implemented at the end of the wave than the beginning. For all 3 waves, the most PHSMs were implemented at the peak of the wave (Table 7).

Vaccination Coverage

Countries began SARS-CoV-2 vaccination campaigns after the first wave during February 17 (South Africa) through April 14, 2021 (Zambia) (5). By the time the second wave began, 7/10 countries (excluding Namibia, Lesotho, and Eswatini) had begun vaccinations; all countries had begun vaccinations by the third wave (Table 2). As of September 19, 2021, 10.8% of the population was vaccinated on average across southern Africa and 8.1% fully vaccinated (Table 2). Coverage varied by country: Eswatini had 16.5% and Zambia 1.5% fully vaccinated. Seven-day vaccinations per 1 million persons steadily increased across waves and were 4.2-fold higher during the third wave (1,087.9) than the second (262.1) (Table 2).

Discussion

Among key findings, we found that patterns of wave propagation throughout the region were similar across almost all country waves. In the absence of a representative regional surveillance system for influenza-like illness, surveillance data from South Africa, where waves were first detected, provided an early warning signal for other countries in the region. Although per person volume of testing increased over time in southern Africa, it remained low compared with resource-rich countries and differed among countries, limiting the ability to compare reported incidence and mortality. Genomic sequencing in the region was limited outside of South Africa. In most countries, reported percentage positivity, incidence rates, mortality rates,

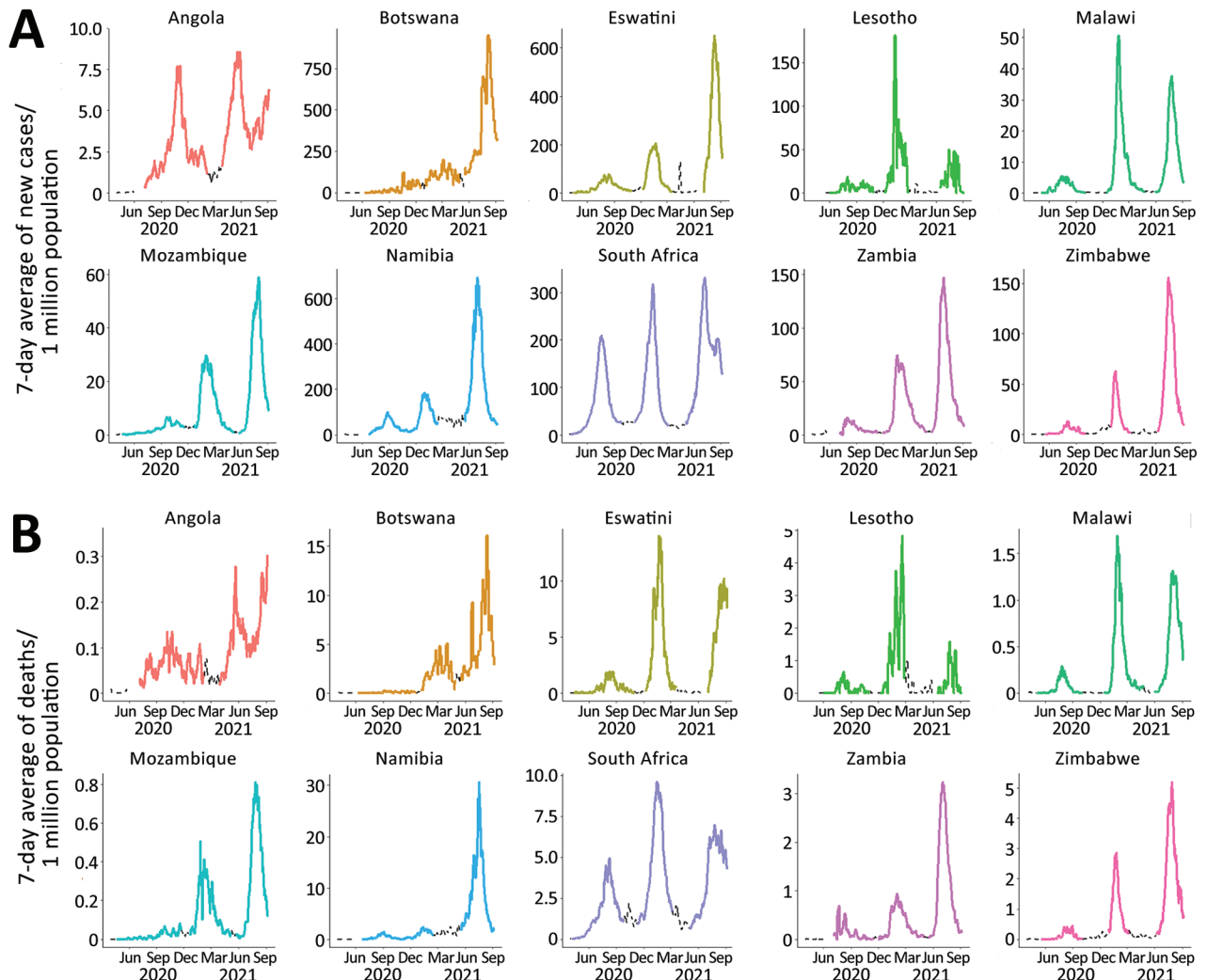


Figure 2. Reported 7-day average new COVID-19 cases (A) and deaths (B) per 1 million persons across pandemic waves in 10 countries in southern Africa, March 5, 2020–September 19, 2021. Colored lines indicate designated wave periods, dashed lines indicate periods between waves. We used differing y-axis scales in this figure to better visualize the wave patterns in each individual country. See Appendix Figure 2 (<https://wwwnc.cdc.gov/EID/article/28/13/22-0228-App1.pdf>) for the same figure placed on corresponding y-axis scales to compare wave magnitudes across countries. Source: Our World in Data (<https://www.ourworldindata.org>), accessed 2021 Sep 20.

and CFRs increased across waves, partly caused by the emergence of more transmissible variants. Stringent PHSM implementation declined over successive waves, and vaccine coverage was low.

Because South Africa accounted for >30% of cases in Africa, average wave patterns were similar between southern Africa and Africa as a whole but with notable regional and intercountry variations (6). Kenya, in eastern Africa, experienced second and third waves before southern Africa. In southern Africa, all waves followed a similar regional pattern: waves were first detected in South Africa, then throughout the remaining interconnected countries an average of 7.2 weeks later. This pattern was less

obvious for Angola, where the second wave started 20 weeks after South Africa (Appendix Table 1). This pattern likely reflects greater testing capacity in South Africa, more sensitive surveillance, and possibly mobility characteristics in the region because South Africa is an international transportation hub. According to phylogenetic analysis, South Africa was determined to be the source of SARS-CoV-2 cases imported to the rest of the region during the first and second pandemic waves (17). Awareness of this pattern is critical for future mitigation efforts; pretravel testing and ongoing sentinel surveillance might be critical for detecting cross-border transmission early, and pandemic surveillance and

Table 3. Overall genomic surveillance comparison across 3 COVID-19 pandemic waves for 10 countries in southern Africa, March 1, 2020–September 6, 2021*

Measure	Before wave 2 start	After wave 2 start, before wave 3 start	After wave 3 start	Overall
Total no. (%) specimens.†	5,543 (23.8)	7,660 (32.9)	10,103 (43.3)	23,306 (100)
Originating country, no. (%)				
Angola	615 (11.1)	264 (3.4)	20 (0.2)	899 (3.9)
Botswana	83 (1.5)	216 (2.8)	799 (7.9)	1098 (4.7)
Eswatini	11 (0.2)	77 (1.0)	34 (0.3)	122 (0.5)
Lesotho	2 (<0.1)	16 (0.2)	0	18 (0.1)
Malawi	16 (0.3)	391 (5.1)	104 (1.0)	511 (2.2)
Mozambique	126 (2.3)	388 (5.1)	66 (0.7)	580 (2.5)
Namibia	19 (0.3)	196 (2.6)	48 (0.5)	263 (1.1)
South Africa	4,013 (72.4)	5,601 (73.1)	8,850 (87.6)	18,464 (79.2)
Zambia	426 (7.7)	182 (2.4)	84 (0.8)	692 (3.0)
Zimbabwe	232 (4.2)	329 (4.3)	98 (1.0)	659 (2.8)
Submitting country, no. (%)				
Botswana	83 (1.5)	216 (2.8)	799 (7.9)	1,098 (4.7)
Germany‡	1 (<0.1)	64 (0.8)	47 (0.5)	112 (0.5)
Malawi	14 (0.3)	8 (0.1)	38 (0.4)	60 (0.3)
South Africa	4,794 (86.5)	6,910 (90.2)	9,135 (90.4)	20,839 (89.4)
Spain‡	13 (0.2)	117 (1.5)	0	130 (0.6)
United Kingdom‡	210 (3.8)	189 (2.5)	0	399 (1.7)
United States‡	0	12 (0.2)	0	12 (0.1)
Zambia	426 (7.7)	144 (1.9)	84 (0.8)	654 (2.8)
Patient sex, no. (%)				
F	2,951 (53.2)	4,053 (52.9)	5,695 (56.4)	12,699 (54.5)
M	2,132 (38.5)	3,274 (42.7)	4,044 (40.0)	9,450 (40.5)
Unknown	460 (8.3)	333 (4.3)	364 (3.6)	1,157 (5.0)
Patient age, y, median (IQR)	37 (27–50)	37 (25–52)	39 (27–54)	38 (26–52)
Genomic surveillance coverage,† median (IQR)	1.55 (0.66–2.79)	3.76 (2.68–4.56)	3.98 (3.37–4.58)	3.48 (2.70–4.44)
Detected SARS-CoV-2 variants, no. (%)§				
Alpha, B.1.1.7 + Q.x	33 (0.6)	158 (2.1)	168 (1.7)	359 (1.5)
Beta, B.1.351 + B.1.351.x	749 (13.7)	6,176 (80.6)	1,493 (14.8)	8,418 (36.2)
Delta, B.1.617.2 + AY.x	0	129 (1.7)	7,454 (73.8)	7,583 (32.6)
Gamma, P.1 + P.1.x	1 (<0.1)	0	1 (<0.1)	2 (<0.1)
Variant of interest	0	1 (<0.1)	0	1 (<0.1)
Variant under monitoring	9 (0.2)	37 (0.5)	174 (1.7)	220 (0.9)
Former variant of interest	2 (<0.1)	1 (<0.1)	1 (<0.1)	4 (<0.1)
Lineages with no WHO label	4,673 (85.3)	1,155 (15.1)	812 (8.0)	6,640 (28.6)
January 2020 strain	14 (0.3)	3 (<0.1)	0	17 (0.1)

*Source: GISAID (<https://www.gisaid.org>), accessed 2021 Sep 20. IQR, interquartile range; World Health Organization.

†Genomic surveillance coverage was defined as the number of reported cases per million per sample sequenced. A country with a higher number of reported cases per million per sample sequenced has lower genomic surveillance coverage than a country with a lower number of reported cases per million per sample sequenced.

‡Germany submitted sequences for specimens collected in Mozambique (n = 1), Namibia (n = 73), Zambia (n = 38). Spain submitted sequences for specimens collected in Mozambique (n = 130). UK submitted sequences for specimens collected in Zimbabwe (n = 401). The United States submitted sequences for specimens collected in South Africa (n = 12).

§Specimens were classified using labels defined by the World Health Organization (Appendix Table 2, <https://wwwnc.cdc.gov/EID/article/28/13/22-0228-App1.pdf>).

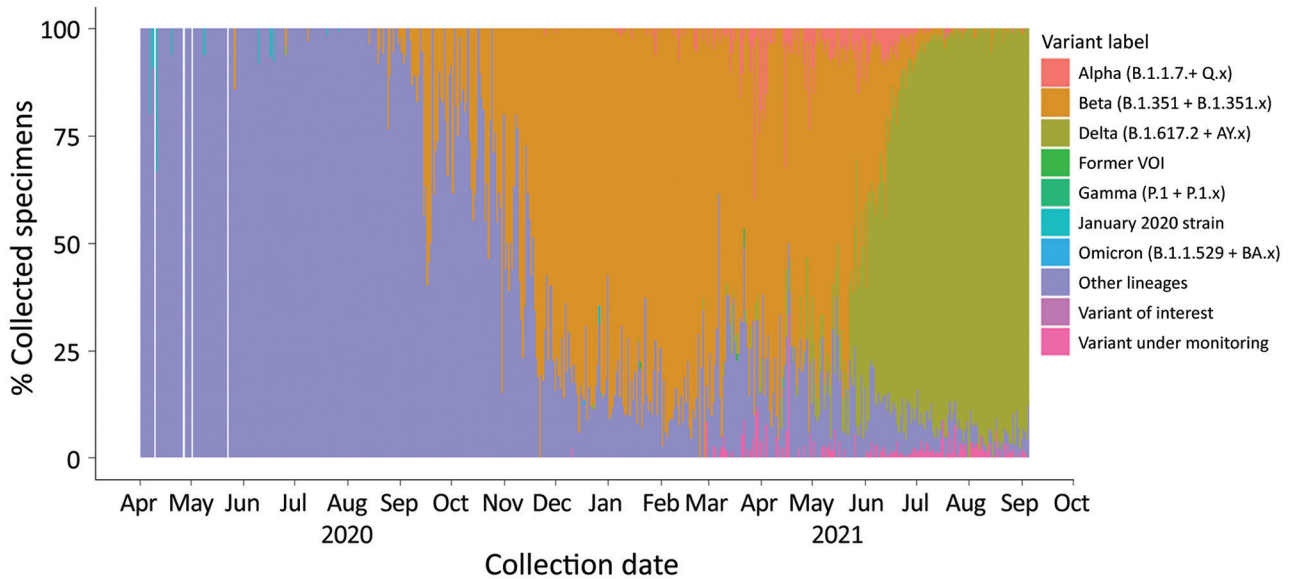


Figure 3. Percentage of SARS-CoV-2 variants among specimens submitted to GISAID in southern Africa, March 1, 2020–September 6, 2021. Definitions of variants are in Appendix Table 2 (<https://wwwnc.cdc.gov/EID/article/28/13/22-0228-App1.pdf>). Source: GISAID (<https://www.gisaid.org>), accessed 2021 Sep 20.

reporting in South Africa can serve as an early warning signal for countries with more limited testing capacity. However, a regional, representative, surveillance system for influenza-like illness and severe acute respiratory illness could improve regional detection and response systems.

Although weekly population-level numbers of tests increased, testing per case, an indicator of sufficient coverage in high-transmission periods, decreased across waves, and the region never achieved

the WHO-recommended target of 1,000 tests/1 million persons (18). The region’s average testing volume per person was low compared with resource-rich countries: ≈240 tests/1 million persons/day in southern Africa versus >3,000 tests/1 million/day in the United States (4). Even Namibia and South Africa, despite relatively higher testing volumes, were below the WHO target for testing. This target might be unreachable for most countries in this region unless test accessibility for the general population is

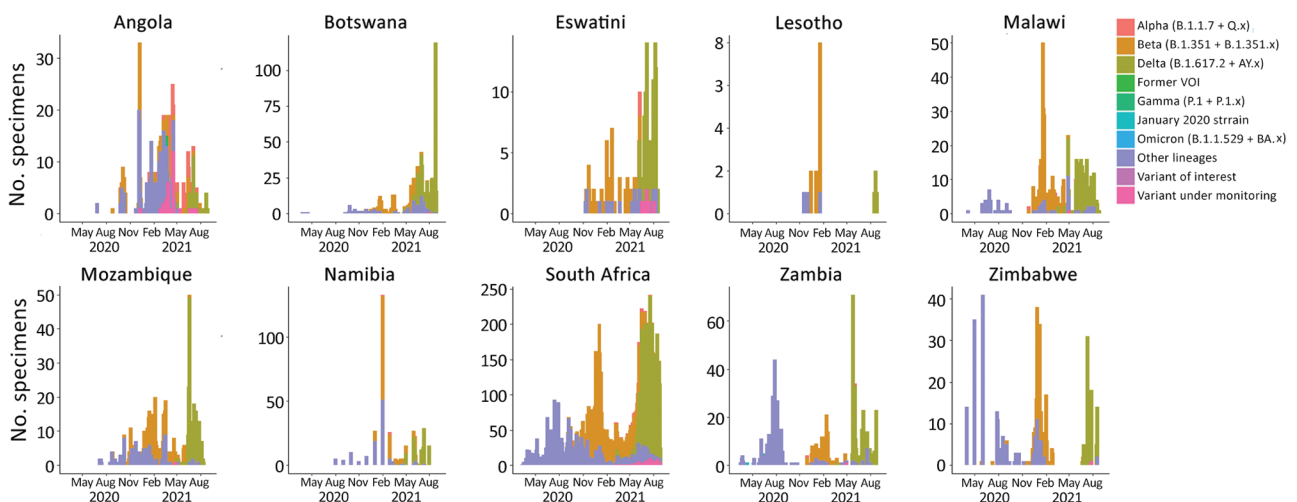


Figure 4. Counts of SARS-CoV-2 variants (World Health Organization classifications) in 10 countries in southern Africa, March 1, 2020–September 6, 2021. Definitions of variants are in Appendix Table 2 (<https://wwwnc.cdc.gov/EID/article/28/13/22-0228-App1.pdf>). We used differing y-axis scales used in this figure to better visualize genomic sampling patterns in each individual country. See Appendix Figure 3 for the same figure placed on corresponding y-axis scales to compare wave magnitudes across countries. Source: GISAID (<https://www.gisaid.org>), accessed 2021 Sep 20.

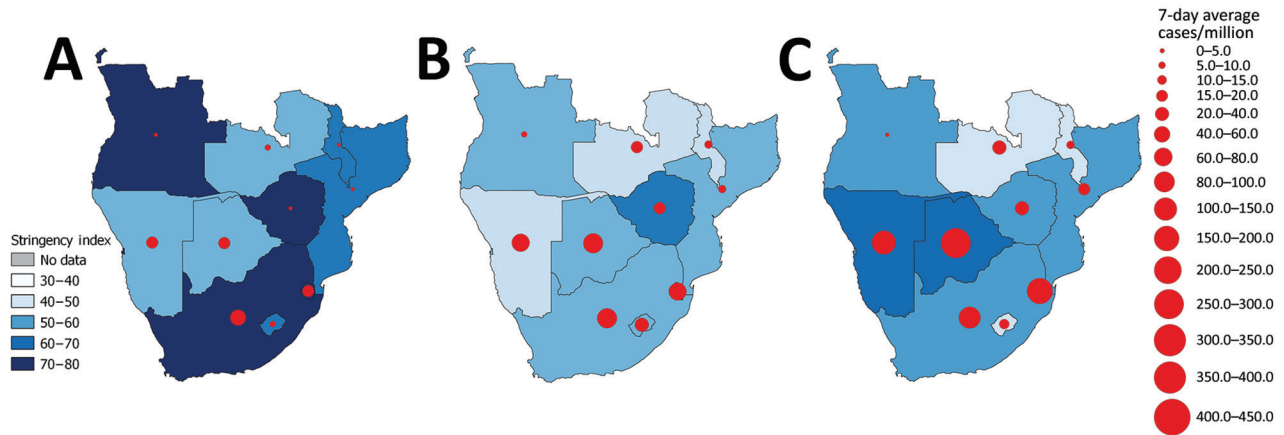


Figure 5. Comparison of public health and social measure stringency and 7-day average new COVID-19 cases per million across 3 COVID-19 pandemic waves in 10 countries in southern Africa, April 6, 2020–July 17, 2021. Source: GISAIID (<https://www.gisaid.org>), accessed 2021 Sep 20.

substantially improved. Increasing availability and feasibility of COVID-19 self-tests, as recommended by Africa CDC (19), might increase testing and improve public health mitigation efforts (20,21).

In this resource-constrained region, testing volumes should be expanded on the basis of need and be designed to collect data to address key objectives for public health response. These data include diagnosing admissions, classifying excess deaths because of COVID-19, defining the timing of pandemic waves, monitoring circulating variants, and informing guidance for work, school, and social engagements. Data gathered from serosurveillance and postmortem activities might also help address these objectives (3,4,22).

Our ability to directly compare SARS-CoV-2 case and death counts in the region using publicly available data was limited by changes over time in test types and availability, low likelihood of diagnosis (4), and various and changing testing strategies. Sustaining COVID-19 sentinel surveillance systems in the community and among high-risk populations (23), including through targeted use of antigen rapid diagnostic tests (24), and improving standard reporting throughout the region to ensure appropriate local epidemiologic evaluations and responses (25), could be considered. These data-gathering systems could be coordinated through a regional body such as the recently established Africa CDC Southern Africa Regional Collaborating Centre (26).

Table 6. Most frequent public health and social measure types implemented across COVID-19 pandemic waves for 10 countries in southern Africa, January 1, 2020–September 19, 2021*

Wave	Most common measures	Least common measures
Wave 1		
Beginning	Workplace closing, cancel public events, restrictions on gatherings, international travel	Public transport closings
Peak	School closing, workplace closing, cancel public events, restrictions on gatherings, international travel	Public transport closings
End	School closing, workplace closing, cancel public events, restrictions on gatherings, international travel	Public transport closings
Duration	School closing, workplace closing, cancel public events, international travel	Public transport closings
Wave 2		
Beginning	Workplace closing, cancel public events, restrictions on gatherings, international travel	Public transport closings
Peak	School closing, workplace closing, cancel public events, restrictions on gatherings, international travel	Movement restrictions
End	School closing, cancel public events, restrictions on gatherings, international travel	Close public transport, movement restrictions
Duration	School closing, cancel public events, restrictions on gatherings, international travel	Public transport closings
Wave 3		
Beginning	Restrictions on gatherings, international travel	Movement restrictions
Peak	Workplace closing, cancel public events, restrictions on gatherings, international travel	Public transport closings, movement restrictions
End	Workplace closing, international travel	Movement restrictions
Duration	International travel	School closings

*Public health and social measures, as defined and measured by Oxford COVID-19 Government Response Tracker (<https://www.bsg.ox.ac.uk/research/research-projects/covid-19-government-response-tracker>), accessed 2021 Sep 20.

Table 7. Number of public health and social measures implemented across COVID-19 pandemic waves by type for 10 countries in southern Africa, January 1, 2020–September 19, 2021*

Intervention	Wave 1, n = 10				Wave 2, n = 10				Wave 3, n = 10†			
	Start	Peak	End	Duration	Start	Peak	End	Duration	Start	Peak	End	Duration
School closings	9	10	10	10	9	10	10	10	4	7*	2	1
Workplace closings	10	10	10	10	10	10	9	9	9	9*	10	6
Canceled public events	10	10	10	10	10	10	10	10	9	9*	9	7
Restrictions on gatherings	10	10	10	9	10	10	10	10	10	9*	9	8
Public transport closings	7	6	4	4	2	5	5	0	5	5*	7	2
Stay-at-home requirements	9	9	8	7	7	9	10	6	8	7*	8	7
Movement restrictions	9	9	7	6	6	4	5	2	3	5*	4	2
International travel	10	10	10	10	10	10	10	10	10	9*	10	9

*Public health and social measures, as defined and measured by Oxford COVID-19 Government Response Tracker <https://www.bsg.ox.ac.uk/research/research-projects/covid-19-government-response-tracker>), accessed 2021 Sep 20.

†For peak of wave 3, n = 9.

Genomic sequencing varied across countries and was limited outside South Africa. Low sequencing limits detection of new VOCs, posing regional and global health security risks. Africa CDC and WHO are strengthening genomic surveillance by establishing a continentwide laboratory network, leveraging existing surveillance systems, to better detect variant evolution (27). To improve sequencing of SARS-CoV-2 and other endemic and epidemic pathogens, systematic in-country genomic surveillance could be built and sustained in the region by adopting sequencing targets such as weekly targets based on incidence and estimated prevalence of variants in line with Africa CDC guidelines (28–30). In southern Africa, Beta variant was predominant in the second wave and Delta in the third.

Across the region, the third COVID-19 wave had the highest 7-day average percentage positivity, daily cases, deaths per 1 million population, and CFR. Increases in reported incidence and mortality at a time of increasing percentage positivity occurred at least partly because of the emergence of more transmissible variants across waves. However, the connection between high testing volume and reported incidence and mortality rates per person in upper-middle-income countries Namibia and South Africa might reflect better testing capacity contributing to improved accuracy of identifying cases and classifying cause of death, leading to higher reported overall incidence and mortality rates (31).

Neither emergence of more transmissible variants nor improved testing capacity can fully explain the increase in CFR over time, an observation that has been previously reported for Africa (32–34). Possible explanations for this increase include increased strain on limited critical care capacity as transmission and hospitalizations increased (6,34,35); health systems with minimal critical care resources are not optimized for managing critically ill COVID-19 cases. A recent prospective cohort analysis found that mortality

among critically ill hospitalized patients was 48.2% in Africa, higher than the estimated 31.5% global average (32,34). Other explanations might include delays in healthcare-seeking behavior by patients, improved differential testing and reporting (i.e., relatively fewer tests among persons who are not ill but more among very ill persons), improvements in classifying COVID-19–related deaths, and declining ability to protect vulnerable populations from SARS-CoV-2 exposure. However, increased CFR in the region suggests the need for improved health systems and access to newer therapeutics for high-risk patients, such as antivirals molnupiravir (36) and nirmatrelvir (37).

The increased incidence, mortality, and CFR during the third wave were not universal across countries. Lesotho reported highest average incidence rates during its second wave, and Lesotho and South Africa reported highest average mortality rates during the second waves. Eswatini also reported a lower CFR in its third wave than in its second. Possible explanations for those patterns include development of natural immunity to severe disease (4), improved outbreak response and service delivery (38), or incomplete data analysis because the third wave was not yet complete when we collected data.

Declining stringency in adherence to PHSMs in the region likely occurred as governments acknowledged sociopolitical, cultural, and economic context, rather than just epidemiologic data, to determine appropriate restrictions (39). Decreasing acceptance of and adherence to PHSMs has been observed in 4 countries in the region, in part because of negative effects on livelihoods and lack of access to health services (23). To improve adherence, PHSMs could be introduced, adapted, and lifted based on situational assessments in each country and considering community feedback (25,40,41). Given likely challenges in implementing and enforcing stringent PHSMs in the future, policymakers could consider targeting new measures towards persons at highest risk for severe disease.

On average, 8.1% of the population in the region was fully vaccinated as of September 19, 2021, compared with 46.7 in Morocco, 53.9 in the United States, and 63.6 in Israel (5). Vaccine coverage in southern Africa faced challenges including low domestic manufacturing capacity, donations of vaccines near their expiration dates, vaccine hoarding by high-income countries, and low vaccine uptake (42,43), highlighting the need to expand equitable access to vaccines and regional vaccine manufacturing capacity (44). Considering the WHO-recommended target that 70% of the population be fully vaccinated by mid-2022 might be unrealistic for the region (45) and likely high SARS-CoV-2 seropositivity (4), vaccination campaigns targeting populations in the region at highest risk for death, such as persons who are elderly or have chronic underlying conditions (46), might be effective in reducing severe disease and emergence of VOCs (47). To expand access to COVID-19 vaccinations, particularly for immunosuppressed persons, some countries in Africa (e.g., Zambia) have integrated COVID-19 vaccination services into existing health delivery platforms and clinics (e.g., HIV clinics); bringing vaccine access closer to home might aid in uptake (48).

We used publicly available datasets, each with data quality challenges. The OWID dataset missed some daily reports, so we requested coauthor data validation from country officials and Africa CDC to ensure reliability of the data. However, missing data from OWID limited our ability to compare pandemic waves between countries, especially those outside South Africa. OWID uses date reported, rather than specimen collection date, meaning that waves might have appeared to begin and end later in countries with time lags between testing and reporting. We assumed standard WHO definitions were used for reporting COVID-19 cases and deaths in the OWID dataset. We did not account for changes in test availability and testing strategies over time, which limited consideration of potential differences in those indicators among countries. The GISAID dataset varied in representativeness because some countries submitted very limited specimens, so we reported genomic surveillance results at a country level to highlight variability among countries. The OxCGRT dataset includes safety and control measures mandated by governments but not the extent of adherence to the measures, which might better correlate with transmission. Regional trends might be more influenced by data reported by an individual country, particularly South Africa, which provided most OWID and GISAID data. Our data were also

extracted while the third wave was ongoing in the region, although it was declining except in Angola, where the third wave had not yet peaked by September 19, 2021. Despite those limitations, by soliciting data reviews from representatives for each country, reporting results at a country level, and computing regional indicators averaging country rates adjusted for population size and daily variation, we have compiled a reasonable description of the pandemic situation across southern Africa.

By September 19, 2021, southern Africa had experienced 3 waves of COVID-19, almost all first detected in South Africa, and with successively higher reported percentages of positivity, incidence, mortality, and CFRs. Increased incidence and mortality could be partly explained by the emergence of more transmissible SARS-CoV-2 variants and improved testing capacity and surveillance. Increasing CFRs warrants further research and highlights opportunities for strengthening health systems and increasing access to feasible therapeutics for high-risk persons. Testing volume increased across waves but varied by country and remained low compared with resource-rich countries. Genomic surveillance capacity was limited, although South Africa played a key role in supporting other countries. Stringent PHSM implementation declined over time, indicating a decrease in feasibility. Vaccination coverage remained very low; scale-up, especially among high-risk persons, should be considered. Coordinated regional solutions could be considered to strengthen and sustain sentinel surveillance systems, genomic surveillance capacity, risk-based vaccination, and tailored public health mitigation to better detect, prevent, and reduce the severity of future COVID-19 waves and other outbreaks in southern Africa.

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Comparison of COVID-19 Pandemic Waves in 10 Countries in Southern Africa, 2020–2021

Appendix

Appendix Table 1. Dates for starts, ends and peaks of COVID-19 pandemic waves in 10 Southern African countries, 06 April 2020 to 19 September 2021*

Country	First wave			Second wave			Third wave†	
	Start	Peak	End	Start	Peak	End	Start	Peak
Angola	06 July 2020 (week 28)	25 Oct 2020 (week 43)	07 Feb 2021 (week 58)	29 March 2021 (week 66)	30 May 2021 (week 74)	04 July 2021 (week 79)	05 July 2021 (week 80)	-
Botswana	11 May 2020 (week 20)	18 Oct 2020 (week 42)	20 Dec 2020 (week 51)	04 Jan 2021 (week 54)	07 March 2021 (week 62)	02 May 2021 (week 70)	17 May 2021 (week 73)	08 Aug 2021 (week 84)
Eswatini	22 June 2020 (week 26)	16 Aug 2020 (week 33)	11 Oct 2020 (week 41)	07 Dec 2020 (week 50)	24 Jan 2021 (week 56)	21 March 2021 (week 64)	05 July 2021 (week 80)	15 Aug 2021 (week 85)
Lesotho	22 June 2020 (week 26)	02 Aug 2020 (week 31)	18 Oct 2020 (week 42)	30 Nov 2020 (week 49)	10 Jan 2021 (week 54)	28 Feb 2021 (week 61)	07 June 2021 (week 76)	01 Aug 2021 (week 83)
Malawi	15 June 2020 (week 25)	12 July 2020 (week 28)	04 Oct 2020 (week 40)	14 Dec 2020 (week 51)	24 Jan 2021 (week 56)	25 April 2021 (week 69)	31 May 2021 (week 75)	25 July 2021 (week 82)
Mozambique	29 June 2020 (week 27)	20 Sept 2020 (week 38)	22 Nov 2020 (week 47)	21 Dec 2020 (week 52)	31 Jan 2021 (week 57)	09 May 2021 (week 71)	24 May 2021 (week 74)	01 Aug 2021 (week 83)
Namibia	15 June 2020 (week 25)	23 Aug 2020 (week 34)	08 Nov 2020 (week 45)	09 Nov 2020 (week 46)	27 Dec 2020 (week 52)	14 Feb 2021 (week 59)	17 May 2021 (week 73)	27 July 2021 (week 78)
South Africa	06 April 2020 (week 15)	19 July 2020 (week 29)	13 Sept 2020 (week 37)	09 Nov 2020 (week 46)	10 Jan 2021 (week 54)	07 March 2021 (week 62)	03 May 2021 (week 71)	04 July 2021 (week 79)
Zambia	06 July 2020 (week 28)	02 Aug 2020 (week 31)	25 Oct 2020 (week 43)	30 Nov 2020 (week 49)	17 Jan 2021 (week 55)	02 May 2021 (week 70)	17 May 2021 (week 73)	27 June 2021 (week 78)
Zimbabwe	29 June 2020 (week 27)	02 Aug 2020 (week 31)	20 Sept 2020 (week 38)	28 Dec 2020 (week 53)	10 Jan 2021 (week 54)	07 March 2021 (week 62)	07 June 2021 (week 76)	18 July 2021 (week 81)

Source: Our World in Data (OWID), accessed 20 September 2021

*We used Salyer et al.'s start week for a country's first wave. Otherwise, 'rising numbers of COVID-19 cases' were classified by examining fold-increases in weekly cases per million, where the first week of at least 1-fold sequential week-by-week increases indicated the start week of a wave. Peak weeks were defined as a local maximum preceded by sequential week-by-week increases and followed by a sequential week-by-week decline. End weeks were defined as the first week of a local minimum following a sequential week-by-week decline. To align with global epidemiologic reporting, we used WHO epidemiologic weeks: start dates are the first day of that week (Monday); and end dates are the last day of that week (Sunday). Given the selected definition for start, peak and end weeks, wave periods used for this analysis and therefore resultant statistics may differ from those used in-country by Southern African governments that may have applied a different definition for wave start and end weeks.

†We used 19 Sept 2021 (week 90), the final date of data extraction for our study, as the end date of the third wave for all southern Africa countries although the wave was ongoing in Angola

Appendix Table 2. World Health Organization (WHO) label classification of SARS-CoV-2 variants by Pango lineage

WHO label	Pango lineage
Alpha	B.1.1.7 + Q.x
Beta	B.1.351 + B.1.351.x
Gamma	P.1 + P.1.x
Delta	B.1.617.2 + AY.x
Variants of interest (VOI)	C.37 (Lambda); B.1.621 (Mu)
Variants under monitoring (VUM)	B.1.427; B.1.429; R.1; B.1.466.2; B.1.1.318; B.1.1.519; C.36.3; B.1.214.2; B.1.1.523; B.1.619; B.1.620; C.1.2; B.1.617.1 (Kappa); B.1.526 (Iota); B.1.525 (Eta); B.1.429 (Epsilon)
Former variant of interest (VOI)*	P.2 (Zeta); P.3 (Theta)
Other lineages	All other lineages
January 2020 strain	A

Source: WHO variant label classification (<https://www.who.int/en/activities/tracking-SARS-CoV-2-variants>), accessed 20 September 2021

*No longer a VUM or VOI

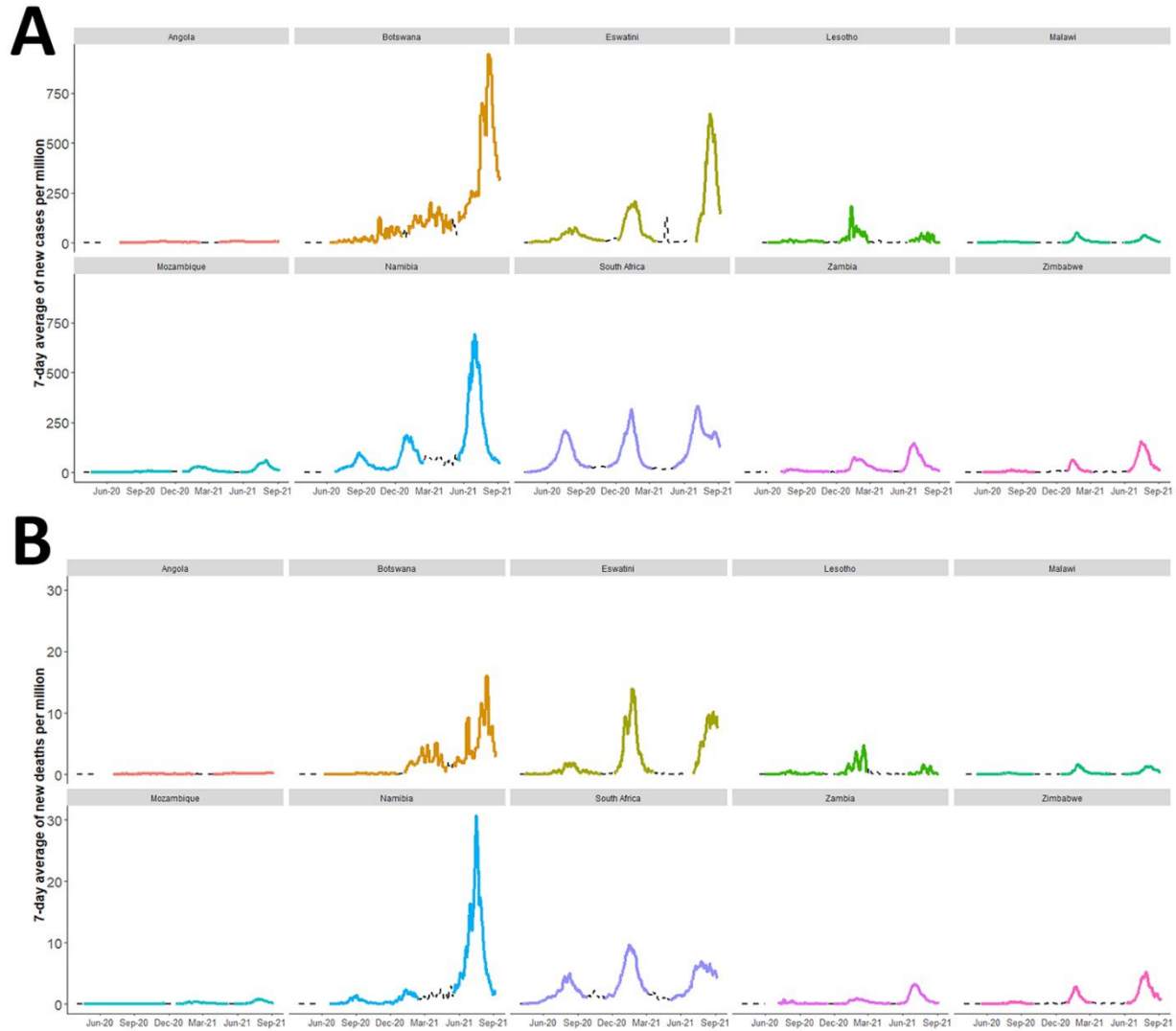
Appendix Table 3. Distribution of SARS-CoV-2 lineages representing $\geq 1\%$ of other lineages classification, 01 March 2020–19 September 2021

Lineage classification in GISAID	Number of sequences	Other lineages category sequences, %
None*	1,337	20.1
B.1	1,031	15.5
B.1.1	571	8.6
B.1.1.448	467	7.0
C.1	383	5.8
B.1.1.54	297	4.5
B.1.237	202	3.0
C.16	187	2.8
B.1.1.273	149	2.2
B.1.1.412	129	1.9
B.1.381	121	1.8
B.1.1.528	89	1.3
AE.1	83	1.3
B.1.1.57	75	1.1
B.1.1.117	73	1.1
B.1.1.34	71	1.1
B.1.1.52	71	1.1
B.1.1.111	66	1.0
Total†	5402	81.4

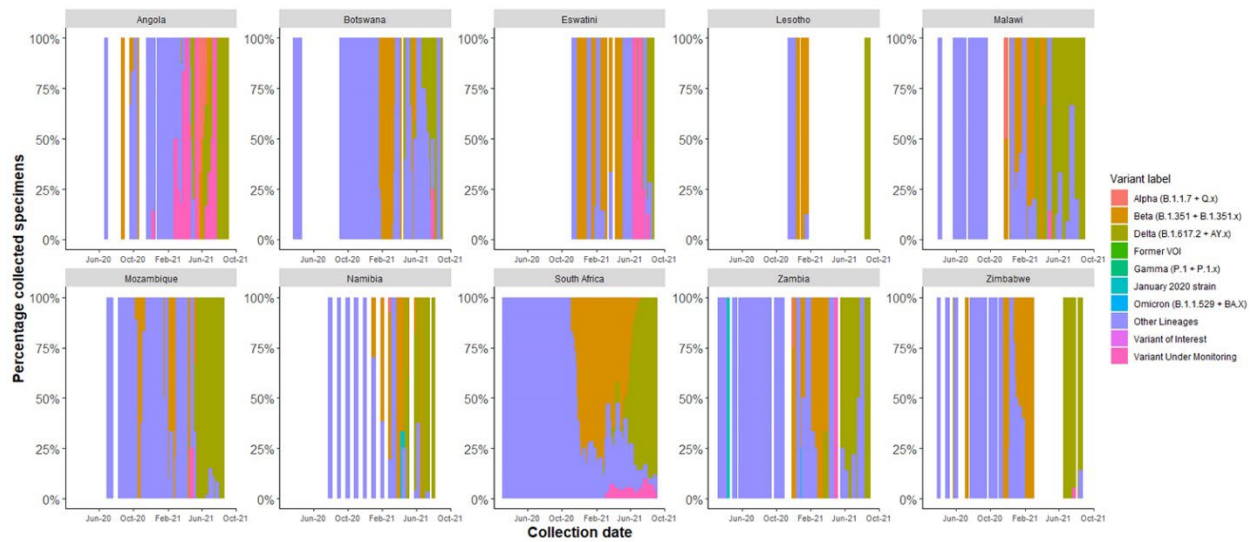
Source: Global Initiative on Sharing Avian Influenza Data (GISAID), accessed 20 September 2021

*No lineage was specified in the GISAID database, although a clade was specified for these specimens.

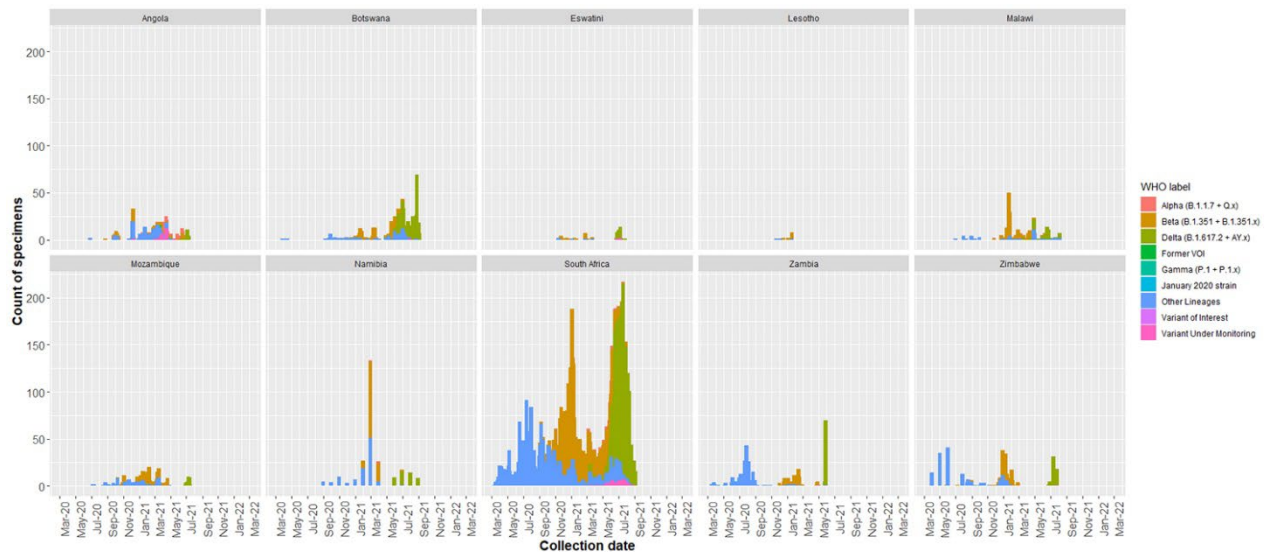
†There were a total of 6,640 sequences during the period. The displayed 5,402 sequences represented 81.4% total lineages that composed at least 1.0% each of the other lineages classification



Appendix Figure 1. Reported 7-day average A) new COVID-19 cases and COVID-19 deaths /1 million persons across pandemic waves in 10 southern Africa countries, 05 March 2020 – 19 September 2021. Colored lines indicate designated wave periods, dashed lines indicate interwave periods. Corresponding Y axes scales were used in this figure to better visualize the comparison of wave magnitudes across countries. Source: Our World in Data (OWID).



Appendix Figure 2. Percentage of collected SARS-CoV-2 specimens by country submitting to GISAID across 10 southern Africa countries, 1 March 2020 – 6 September 2021 (collection date). Source: Global Initiative on Sharing Avian Influenza Data (GISAID).



Appendix Figure 3. Counts of SARS-CoV-2 variants in 10 southern African countries, 1 March 2020 – 6 September 2021 (collection date). Variants were classified according to World Health Organization labels (Appendix Table 2). Corresponding Y axes scales were used in this figure to better visualize the comparison of genomic sampling magnitudes across countries. Source: Global Initiative on Sharing Avian Influenza Data (GISAID).