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Reporting Summary

Nature Research wishes to improve the reproducibility of the work that we publish. This form provides structure for consistency and transparency in reporting. For further information on Nature Research policies, see our <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

Sta	tist	ics							
For a	all sta	tistical ana	alyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.						
n/a	Conf	firmed	med						
\boxtimes		The exact :	sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement						
\boxtimes		4 statemei	nt on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly						
\boxtimes	1 1		istical test(s) used AND whether they are one- or two-sided mon tests should be described solely by name; describe more complex techniques in the Methods section.						
\boxtimes		A descripti	tion of all covariates tested						
\boxtimes		A descripti	on of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons						
	\boxtimes	A full desci AND variat	cription of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) tion (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)						
\boxtimes			ypothesis testing, the test statistic (e.g. F , t , r) with confidence intervals, effect sizes, degrees of freedom and P value noted use as exact values whenever suitable.						
	X F	For Bayesia	an analysis, information on the choice of priors and Markov chain Monte Carlo settings						
\boxtimes	F	For hierard	hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes						
\boxtimes	Estimates of effect sizes (e.g. Cohen's <i>d</i> , Pearson's <i>r</i>), indicating how they were calculated								
	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.								
Sof	twa	are and	d code						
Polic	y info	ormation a	bout <u>availability of computer code</u>						
Data collection		llection	From March 2020 to August 2020.						
Data analysis Genome Detective,		alysis	Genome Detective,						

For manuscripts utilizing custom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors and reviewers. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Research guidelines for submitting code & software for further information.

Data

Policy information about availability of data

All manuscripts must include a <u>data availability statement</u>. This statement should provide the following information, where applicable:

- Accession codes, unique identifiers, or web links for publicly available datasets
- A list of figures that have associated raw data
- A description of any restrictions on data availability

All sequence data has been deposited in the GISAID (assembled genomes) and the short read archive (for short reads). On GISAID, all of the accession numbers are given in the supplementary data S2 file. On the SRA, the bio-project Accession: PRJNA636748 ID: 636748

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Please select the o	ne below that is the best fit for y	our research. If you are not sure, read the appropriate sections before making your selection.			
Life sciences	Life sciences Behavioural & social sciences Ecological, evolutionary & environmental sciences				
For a reference copy of	the document with all sections, see <u>natur</u>	e.com/documents/nr-reporting-summary-flat.pdf			
Life scier	nces study desi	gn			
All studies must dis	sclose on these points even whe	n the disclosure is negative.			
Sample size	All available genomes from SARS-CoV-2 from South Africa that were produced and available in public database were used in the analysis. At the time of writing, 1365 genomes passed the quality control.				
Data exclusions	Supplementary figure S9 show the data exclusion process. In summary, Curation of South Africa dataset from all available South African genomes available on GISAID as at 15th September 2020, showing the initial number of genomes (n=1409), how many were excluded at each cleaning step and the final number of genomes (n=1365). Genomes were excluded if < 90% of coverage AND/OR have sequencing quality problem. In total, 16 genomes were excluded due to low coverage and 28 due to sequencing problems.				
Replication	Reproducibility were performed for maximum likelihood and bayesian MCMC phylogenetic tree reconstruction. We computed MCMC (Markov chain Monte Carlo) triplicate runs of 100 million states each, sampling every 10.000 steps for each data set.				
Randomization	Samples for South Africa were randomly selected in the most sampled province. This mean that every week before the peak of infection, we would receive 50 samples for sequencing that were randomly selected by the national health laboratory service. During the peak of infections, we received around 150 samples per week for sequencing.				
Blinding	Geographical blinding of data was not necessary for the study as it involves phylogeographical analysis. Data identification from the samples were anonymized as this was not necessary for the analysis.				
Reportin	g for specific n	naterials, systems and methods			
'	**	of materials, experimental systems and methods used in many studies. Here, indicate whether each material, are not sure if a list item applies to your research, read the appropriate section before selecting a response.			
Materials & ex	perimental systems	Methods			
n/a Involved in the study		n/a Involved in the study			
Antibodies		ChIP-seq			
Eukaryotic cell lines		Flow cytometry			
Palaeontology and archaeology		MRI-based neuroimaging			
Animals and other organisms					
Human research participants					
∐ Dual use r	Dual use research of concern				

Human research participants

Policy information about studies involving human research participants

Population characteristics

We obtained deidentified remnant nasopharyngeal and oropharyngeal swab samples from patients testing positive for SARS-CoV-2 by RT-qPCR from public health and private medical diagnostics laboratories.

Recruitment

The patients were mostly selected randomly (>90%), however, four outbreak investigations (3 hospitals) and 1 shopping facility (total of 120 sequences) were nor randomly selected as these were individuals in the outbreak.

Ethics oversight

The project was approved by University of KwaZulu-Natal Biomedical Research Ethics Committee. Protocol reference number: BREC/00001195/2020. Project title: COVID-19 transmission and natural history in KwaZulu-Natal, South Africa: Epidemiological Investigation to Guide Prevention and Clinical Care. This project was also approved by University of the Witwatersrand Human Research Ethics Committee. Clearance certificate number: M180832. Project title: Surveillance for outpatient influenza-like illness and asymptomatic virus colonization in South Africa. Sequence data from the Western Cape was approved by the Stellenbosch University HREC Reference No: N20/04/008_COVID-19. Project Title: COVID-19: sequencing the virus from South African patients.

Note that full information on the approval of the study protocol must also be provided in the manuscript.