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The potential effects of widespread community transmission of SARS-CoV-2 infection in the World Health Organization African Region: a predictive model

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ABSTRACT

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Dr Humphrey Cyprian Karamagi; karamagih@gmail.com The spread of severe acute respiratory syndrome coronavirus-2 (SARS-CoV-2) has been unprecedented in its speed and effects. Interruption of its transmission to prevent widespread community transmission is critical because its effects go beyond the number of COVID-19 cases and deaths and affect the health system capacity to provide other essential services. Highlighting the implications of such a situation, the predictions presented here are derived using a Markov chain model, with the transition states and country specific probabilities derived based on currently available knowledge. A risk of exposure, and vulnerability index are used to make the probabilities country specific. The results predict a high risk of exposure in states of small size, together with Algeria, South Africa and Cameroon. Nigeria will have the largest number of infections, followed by Algeria and South Africa, Mauritania would have the fewest cases, followed by Seychelles and Eritrea. Per capita, Mauritius, Seychelles and Equatorial Guinea would have the highest proportion of their population affected, while Niger, Mauritania and Chad would have the lowest. Of the World Health Organization's 1 billion population in Africa, 22% (16%–26%) will be infected in the first year, with 37 (29 - 44) million symptomatic cases and 150 078 (82 735-189 579) deaths. There will be an estimated 4.6 (3.6-5.5) million COVID-19 hospitalisations, of which 139 521 (81 876-167 044) would be severe cases requiring oxygen, and 89 043 (52 253-106 599) critical cases requiring breathing support. The needed mitigation measures would significantly strain health system capacities, particularly for secondary and tertiary services, while many cases may pass undetected in primary care facilities due to weak diagnostic capacity and non-specific symptoms. The effect of avoiding widespread and sustained community transmission of SARS-CoV-2 is significant, and most likely outweighs any costs of preventing such a scenario. Effective containment measures should be promoted in all countries to best manage the COVID-19 pandemic.

INTRODUCTION

The spread of the severe acute respiratory syndrome coronavirus-2 (SARS-CoV-2) and its resulting coronavirus disease of 2019 (COVID-19) has been unprecedented in speed and impact, contributing to widespread socioeconomic disruption. It has spread to all regions of the world, with 3018681 cases and 207973 deaths reported in the World Health Organization (WHO)'s 100th situation report by 29 April 2020.¹ The report showed that 45 of the 47 countries in the WHO African Region had cases, although fewer than expected. The WHO African Region is comprised of 47 member states from the African continent. These do not include Djibouti, Egypt, Libya, Morocco, Somalia, Sudan and Tunisia. The region had 0.77% of the worldwide cases even though it represents 13.7% of the global population, and 0.43% of deaths, with a case fatality rate of 3.88%, compared with 9.19% in the European region and 6.89% globally.

Containment and mitigation

As with other infectious diseases, transmission of SARS-CoV-2 is a function of three attributes: sources of infection, routes of transmission and susceptibility of individuals to infection.² Most current infection is from other humans, and the greater the exposure the higher the probability of infection. There are four distinct phases of infection intensity for SARS-CoV-2, of increasing possibility of exposure of a susceptible person: (1) sporadic cases (isolated cases primarily from persons known to have recently travelled, with the population

Summary box

What is already known?

- The SARS-CoV-2 pandemic is worldwide, affecting all countries and with everyone at risk of infection and death. However, countries have had different infection rates, and outcomes have suggested a main role for socioecological factors in its transmission, and personal vulnerabilities in determining its outcomes.
- Non-pharmaceutical interventions present the best approach to contain the outbreak. Failure of containment leads to a significant increase in cases and deaths, with mitigation measures overwhelming health system capacities.

What are the new findings?

- The countries in the World Health Organization (WHO) African Region have a lower risk of exposure—varied for each country—than other parts of the world. This suggests a lower rate of transmission in the region.
- Only 22% of the population would be infected in the first year, with widespread community transmission, with fewer numbers of severe cases and deaths compared with that seen in other countries due to different personal vulnerabilities in the countries. However, the increase in hospitalisations and care needs and impact on morbidity and mortality of other conditions would have significant effects due to limited capacity to mitigate against the effects of the disease.

What do the new findings imply?

- The success of the containment measures is critical for the region as their health systems are not designed to mitigate against the implications of widespread community transmission of SARS-COV-2.
- Countries of the WHO African Region need to expand capacity, particularly of their primary hospitals, to mitigate the implications of widespread community spread of SARS-CoV-2. Basic emergency care needs to be included in primary care systems.

at risk largely limited to contacts with the cases); (2) local transmission (additional cases from persons who can be traced to known cases, with the population at risk being the networks to which the cases belong); (3) clustered community transmission (additional cases are from persons who cannot be traced to known cases, but are within a definable cluster, such as a city, slum or

large event, with possible exponential increases in cases within the clusters; the population at risk is the persons in the cluster to which the cases belong); and (4) widespread sustained community transmission (additional cases are from multiple, unconnected clusters and selfpropagating with exponential increases of cases across these many clusters, with the population at risk being most of the population of the country or territory of interest) (figure 1).

The modes of transmission for SARS-CoV-2 are varied. Person-to-person transmission is documented, through respiratory droplets, contact and fomites,³ in addition to the possibility of zoonotic, nosocomial, faecal-oral and aerosol mechanisms.^{4–6} These routes of transmission are thought to be influenced by the socioecological context—the bio-geo-physical variables influencing spread of disease, such as population density, weather, population mobility, social behaviour and others—and build a complex picture of transmission influences.^{7 8} Finally, looking at susceptibility to SARS-CoV-2, older age, male sex and the presence of chronic comorbidities are associated with more severe disease and mortality.^{9–12}

Control therefore is premised on addressing these attributes of transmission through containment measures aimed at avoiding widespread and sustained community transmission. Where these fail, costly mitigation measures are needed to reduce resultant morbidity and mortality. As seen in Europe and the USA, these measures place significant strain on health system capacities due to the inability to cope with the increased demand, with negative effects on other health services due to the diversion of essential supplies, staff and financial resources to the mitigation measures. In contrast with Europe and the USA, many countries in the WHO African Region have implemented containment measures early, primarily focused on physical distancing and improved hygiene practices. Physical distancing measures have been introduced through lockdown of schools, markets, places of worship and other locations to limit clustering mostly to the home. On the other hand, hygiene practices have



Figure 1 Phases for SARS-COV-1 transmission in a country/territory. Source: produced by the authors for this publication.

focused on behavioural practices for improved hand hygiene, face contact, regular disinfection and other mechanisms aimed at reducing person-to-person spread. These containment measures have been introduced and enforced differently depending on perceptions of risk, balanced against enforcement capacity and socioeconomic implications.

Given the fewer events relating to SARS-CoV-2 in Africa, there have been efforts to predict how the pandemic would evolve, leading to varied estimates of impact and projected mortality.^{13–18} There are two reasons for this: knowledge of the virus transmission characteristics is still evolving and the prediction models are not adequately reflecting the socioecological factors. The unique context of a country or territory influences disease transmission.⁷⁸ This may contribute to the different transmission dynamics seen in countries that have more mature SARS-CoV-2 outbreaks, with cases and deaths guite varied. These socioecological factors may play a role in the early pattern of SARS-CoV-2 transmission in many African countries-with slower transmission, fewer cases, less severe infections and fewer deaths than other parts of the world. Underreporting of cases is another possible contributor, but that alone would not explain the unique transmission dynamics in the region as some countries, such as South Africa, with better reporting systems would have outbreaks similar to those seen in the badly affected countries, which they do not, and conversely, we would not see relatively many cases in countries such as Cameroon and Burkina Faso with similar detection capacities to their peers.¹

We therefore designed a model that uses both virus transmission characteristics and country specific socioecological factors to predict the most likely outcome of widespread and sustained community transmission of SARS-CoV-2.

What current epidemiological and statistical evidence tells us

Seven coronaviruses are now known to be associated with disease in humans, and present with clinical similarities to infections by other pathogens, including rhinovirus and influenza A.¹⁹ As symptoms tend to be mild in most people, they are prone to spreading extensively, eventually afflicting persons who may be susceptible to severe disease and leading to their known mortality effects.²⁰

Genome sequencing has shown that SARS coronaviruses merit classification as a new group within the family of *Coronaviridae*.^{20 21} The spike protein of SARS-CoV-2 has undergone several mutations that heighten the probability of human cell infection.²² Human transmission is further amplified by asymptomatic carriers, enabling further undetected spread.^{20 23}

Predictive models exist to capture transmission dynamics^{24–28} focused on different elements, such as the effects of lockdown measures,²⁸ specialised equipment needs^{25 26} and projections on the speed of spread.^{24 27} The differential spread and impact show that influences beyond the characteristics of the virus are significant, in both determining transmission and its effects in terms of disease severity. The virus is currently understood to be highly transmissible, with various estimates for the basic reproductive number (R_0) ranging from 0.41 to 2.39,²⁹ but some estimates are as high as 6.49.³⁰ This is higher than other coronaviruses, justifying strict containment measures.^{31–33}

Estimates of attack rate vary in different studies, depending on the numerator and denominator used. According to the official WHO China Joint Mission Report on SARS-CoV-2, an attack rate of 3-10% was calculated, based on infection rate within a household as a cluster of exposed persons.²⁰ Another analysis suggested an attack rate of 11.2-14.9% within households, but a rate of 6.6-9.7% across the population at risk during the infection.³⁴ These are based on comparing reported cases against clusters of at risk persons. Lower attack rates have been estimated when the whole population in a country is considered susceptible as opposed to a specific cluster, with this ranging from 0.0899% (Canada) to 0.288% (Italy) of the population.³⁵

Multiple pharmaceutical and non-pharmaceutical interventions (NPIs) have been used to slow the rate of infection, with varying degrees of success.^{36 37} Lockdown measures and effective hygiene practices have shown the most promise but are more difficult to enforce the more advanced a country's disease transmission phase is. They are however still effective in reducing new infections, even in the event of widespread sustained community transmission.^{28 31 38 39}

Current evidence classifies infected persons as being asymptomatic, or symptomatic with mild, moderate, severe or critical disease.⁴⁴⁰ Mild disease is associated with non-debilitating symptoms and no radiology features; moderate disease with fever, respiratory symptoms and radiological features; severe disease with either tachypnoea or oxygen saturation <93% or PaO₂/FiO₂ >300 mg; and critical disease with respiratory failure, septic shock or multiorgan failure.⁴⁴¹

On the other hand, the number of asymptomatic infections is still not well estimated. In the early stages of an outbreak, it is difficult to determine this as detection is biased towards clinically severe disease.⁴² The number of asymptomatic cases is best determined from population based seroepidemiology data that provide more accurate numbers of infected persons.^{43 44} The current testing in many countries that is focused on symptomatic suspects and their contacts represents this bias, and implies asymptomatic cases will be underrepresented among cases. This experience was also documented with the recent MERS-CoV outbreak that showed increasing identification of asymptomatic case numbers over time as surveillance and testing strategies expanded.⁴⁵ Initial evidence at the beginning of the pandemic suggested that only 1–5% of cases were asymptomatic.²⁰ An analysis of early data suggested 86.1% of cases are undocumented, and given the bias towards clinical cases, suggested a potentially large number of asymptomatic cases.²³ As testing

is increased to wider populations, we see an increase in asymptomatic SARS-CoV-2 infections being detected, such as in South Korea^{46 47} and Germany.⁴⁸ Testing strategies not focused on symptomatic patients or contacts also show high levels of asymptomatic cases, such as the findings from testing of flight returnees to China, reflecting up to 80% asymptomatic cases.⁴⁹ Studies to ascertain the number of COVID-19 cases being identified suggest the SARS-CoV-2 ascertainment rate as 2.22-35.58%, indicating that many non-severe cases are not reflected in reported cases.³⁷ However, the actual rate of asymptomatic cases can only be determined from seroepidemiology tests to find actual numbers of infected persons. Early results from these studies carried out in multiple places point to the total number of infections being up to 90 times the number of reported cases. We see infection in 30% of the population in Chelsea, Massachusetts, 14% in Heinsberg, Germany, 11% in Stockholm, Sweden, 21% in New York City, USA and 2-4% in Santa Clara County, California, USA, all of which point to asymptomatic cases of 50–95% of total reported ones. 5051 As of 20 April 2020, the WHO estimated that 2-3% of the global population are infected (150-230 million persons) compared with 2314621 reported cases, according to its 91st situation report, translating to only 1.5-2% of estimated cases reported.52 53

Evidence shows that among symptomatic cases, the probability of having mild, moderate, severe or critical disease is approximately 40%, 40%, 15% and 5%, respectively.⁵⁹ Hospitalisation rates vary depending on hospitalisation policy and capacity, but it is estimated that 30% of symptomatic patients need hospitalisation, with case fatality highest for critical cases (up to 89% without intervention) and 49% for severely ill patients. The infection mortality rate, however, is estimated at <0.1%.^{42 54}

These different states and probabilities allow us to build a model for infection in the WHO African Region, with appropriate assumptions tested through sensitivity analyses.

METHODS

Infectious diseases are typically modelled using the conceptual model of a flow of population between four states: susceptible (S), exposed (E), infectious (I) and removed (R). We based our model on this concept, parameterising it to SARS-CoV-2 to incorporate the socioecological parameters that influence its transmission. We considered susceptible persons to be the whole population as there is no evidence of inherent immunity. We however do not expect every person to be susceptible at the same time, with various country specific factors influencing individual susceptibility and leading to the differential numbers of cases. We consolidated these into a 'risk of exposure (RoE)' that allows us to identify a subset of the susceptible persons who will be exposed to infection at a given time. This subset that we classified as the exposed persons differs from the SEIR model, where the exposed

are defined as pre-symptomatic persons. Our model defines I as infected persons among the exposed population, as opposed to infectious. The attack rate represents the probability of an exposed person becoming infected. We considered 'infected' to be 'infectious', but incorporated a pre-symptomatic latent phase within the model to cater for the pre-symptomatic period for all infected persons, irrespective of severity.²⁰ These infected persons were captured as the source of new transmissions in the model, thereby closing the loop. The removed population are the absorbing states of either recovery or death.

Based on markovian assumptions,^{55,56} we modelled the evolution of the disease using transition states and fitted a stochastic Markov chain model representing where any member of a population could exist at a given time. Each of these Markov states, denoted S_i with i = 1, 2, ..., 10, is associated with probabilities of transitioning to the next allowable state:

$$\Pr\left(X_{n+1} \in A | X_0, \ldots, X_n\right) = K(X_n, A).$$

where $A \subset S$ denotes a subset of states, and X_n refers to a sequence of random variables indexed by a variable $n \in \{0, 1, 2, ...\}$ which forms a discrete time random process, where $X_n = \{X_n: n = 0, 1, 2, ...\}$. K refers to the invariant distribution for the Markov chain.

The distribution of the population, X, is represented across the Markov states. X_0 gives the initial distribution of the Markov chain; the subsequent distribution of X_n would be dependent on the distribution of X_0 . This conditional distribution of X_n given X_0 is described by:

$$Pr(X_0 \in A|X_0) = K^n(X_0, A)$$

where K^n is utilised to denote the *nth* application of *K*, an invariant distribution $\Upsilon(x)$ for the Markov chain is a density satisfying the equation:

$$\gamma(A) = \int K(x, A) \gamma(x) dx$$

where $\Upsilon(x)$ refers to both the distribution and density of the random variable.

Considering the still evolving evidence on the probabilities of transitioning between states, sensitivity analysis was conducted based on the range for each probability representing best or worst case, and a 10% deviation where there was no range from the literature.

Transition states

Ten states were defined. In a scenario of widespread and sustained community transmission, a country's population is deemed susceptible (S_1 , the initial Markov state), while the exposed population (those at risk of getting infected at any given time) are the next Markov state, S_2 . Those who get infected are the next state (S_3). If not infected, they will return to the previous Markov state (S_2) to face a continued risk of exposure.

What happens to an infected person is represented by five mutually exclusive Markov states: asymptomatic (S_4) , mild symptoms (S_5) , moderate symptoms (S_6) , severe symptoms (S_7) or critical symptoms (S_8) . These represent



Figure 2 Transition states and probabilities for SARS-CoV-2.

the final states for a case, and are not point-in-time descriptions. Infected persons are the source of transmission for the susceptible population.

The final absorbing states are death (S_{0}) or recovery (S_{10}) , with distribution based on probabilities adjusted for existing vulnerabilities.957

The model recognises that the states are time dependent. The infection has an incubation period of 1-14 days, with an average of 7 days.²⁰ We therefore aligned the model cycle length to 7 days to reflect this time dependency. The simulation was repeated every 7 days for 52 weeks to produce the results. To account for the fact that persons entering each Markov state may not enter at the same time during the cycle, we applied a half cycle correction.⁵⁶ The different transition states and probabilities are summarised in figure 2.

Transition probabilities

We defined transition probabilities for movement across the different Markov states, to obtain the actual numbers of persons in each state at a given time. These are based on current knowledge of SARS-CoV-2 infection. The prevalence of infection is taken to be the same in countries given the scenario is one of widespread sustained community transmission. This implies transmission is occurring across multiple clusters and is self-propagating and characterised by exponential growth in cases from multiple unconnected clusters. This is similar to the transmission that was occurring in Northern Italy, the USA and the UK before they instituted lockdown measures.

To better explore the role of contextual factors, we derived a probability to reflect the given population that is at risk of getting infected at a given time-RoE. This is different from the risk of infection as it identified the subset of the susceptible population that is at risk of infection (but not infected) at a given time, based on contextual specificities in a country. We used four contributory factors based on the knowledge of socioecological effects on disease transmission⁷ and coronavirus transmission modalities.

1. Gathering factor (gf). The tendency for people to cluster together, creating opportunities for transmission. The higher the 'gathering' tendency in a country, the higher the factor and its contribution to the RoE. People cluster during their activities of daily living, in the home, work/school or social environments. We used the following indicators based on data availability to reflect these: household size; number of children in school per capita; per cent of the population living in slums; and per cent of the population living in urban areas. We considered but did not include alcohol consumption per capita (as a proxy for social gathering) as most countries do not have recent or available data on this indicator. The normalised values for the applied indicators were averaged to produce a country specific gf.

Gathering factor $(gf) = \frac{\sum_{i=1}^{n} P_i}{n}$, where P_i refers to the *i*th gathering factor and *n* refers to the total number of factors.

- 2. Weather factor (wf). Although the influence of seasonality on SARS COV-2 is not clear,⁵⁸ weather is a recognised socioecological factor affecting transmission of disease caused by other coronaviruses.⁵⁹ Early studies also suggest that humidity and temperature could potentially affect the activity and transmissibility of SARS-CoV-2.^{60–64} We therefore included precipitation as a contributor to the RoE, being the only indicator relating to these with data across the countries in the region.
- 3. Distribution factor (df). Countries differ in the ease of movement of an exposed person, thus varying the extent to which an infected person can spread the virus from one location to another. We used permanent road network per square kilometre and vehicle density as proxy indicators for distribution factor. Both were applied as a country may have a limited road network (due, for example, to low population) but with heavy utilisation, or vice versa.
- 4. Sanitation and hygiene practices (sh). The mainstay of prevention of SARS-COV-2 is hygiene practices. It was therefore deemed that the effectiveness of a population's hygiene practices would influence the RoE. The percentage of people using at least basic sanitation services-that is, improved sanitation facilities that are not shared with other households-was used as a proxy for hygiene practices.

We recognise that these factors are distributed unevenly in each country, and so the results should only be used at the national level, because extrapolation to the subnational level would depend on the specific values for each of the sub-national units.

Examining the relationship across these factors, the gf and df were assumed to behave in an interactive way, as clustering and distribution are related, while the wf and sh were assumed to be additive. Hence the product of gf and df was then averaged with the wf (precipitation) and sh to derive the estimated RoE for each of the countries.

[(Gathering factor \times Distribution factor)+

Pick of antosura -	Weather factor)]	× hypigma factor
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We considered applying more weight to the gathering and hygiene factors, and less to the weather factor due to the uncertainty of the influence of weather factors on the speed of transmission. However, there was no evidence to guide weighting criteria. We therefore maintained an equal weighting to the contribution of the four factors.

Other factors are being explored to explain the differential transmission, such as possible cross immunity with other coronaviruses or BCG vaccine,⁶⁵ and population behavioural characteristics. The model allows for the RoE to be updated when evidence on these becomes conclusive.²³

The remaining transition probabilities relate to the probability of an exposed person becoming infected, the attack rate, an infected person becoming either asymptomatic or falling into one of the defined symptomatic states, and the probability of recovery or death.

The attack rate represents the transition probability from S_2 to S_3 states. We used the average value of 6.5%, based on the 6.6% overall rate found in the community transmission,³⁴ with 3–15% representing the two extreme values from the literature for best and worst case scenarios, respectively.

The model incorporates the NPIs relating to the distancing and hygiene measures a country has implemented to reduce the attack rate. Ten NPIs are included: seven distancing ones (closure of schools, formal workplaces, markets and social entertainment places, together with restrictions on inter city travel, social gatherings such as burials and public transportation) and three hygiene practices (enforcement of regular handwashing and safe hygiene practices). The attack rate can then be customised for each country based on which NPIs had been introduced, and how much they have been implemented.

The more the NPIs and higher their implementation, the greater the effect of reducing he country's attack rate.

From S_a, the probabilities of transitioning to one of the five states are based on the literature. We applied an estimate of 80% of cases (allowing 10% variation for best and worst case scenarios) being asymptomatic based on the emerging literature of asymptomatic proportions. We applied sensitivity analysis to explore how this varies, reflecting the still emerging evidence. For symptomatic cases, we applied the proportions of 40%, 40%, 15% and 5% representing the mild, moderate, severe and critical forms of the disease, respectively. This categorisation translates to 88% of infected persons not realising they are infected with COVID-19 (have mild/ no symptoms), while 4% will have severe/critical disease requiring specialist intervention. The remaining 8% will have moderate but non-life threatening symptoms. For recovery following infection, we applied 100% of asymptomatic, mild and moderate cases, 50% of severe cases and 12% of critical cases.²

Regarding hospitalisation, assuming 30% of symptomatic cases require hospitalisation,⁶⁶ we considered all severe and critical cases would need hospitalisation, and applied the remainder to moderate cases who would be hospitalised (25%). The initial probabilities associated with each transition are shown in table 1. The values in the model differ for each country, based on its context specific information.

Factoring in vulnerabilities

The final application of socioecological factors was with biological aspects influencing disease progression. Current evidence suggests that COVID-19 is more severe among older people, men, and people with underlying chronic health problems.⁹⁵⁷ Given the difference in these parameters in the WHO African countries compared with Europe and China where most probability assumptions have been developed, we applied an adjustment factor

Table 1 Para	ameter values for transmission dynamics of SARS-CoV-2 ir	n the World Health O	rganization Africa	In region
Parameter	Probability description	Value	Best case	Worst case
P ₁	Risk of exposure (S_1 to S_2)	Country specific	10% lower	10% higher
P ₂	Attack rate (S_2 to S_3)	0.065	0.03	0.10
P ₃	Asymptomatic infection (S_3 to S_4)	0.8	0.88	0.72
P ₄	Mild infection (S_3 to S_5)	0.08	0.088	0.072
P ₅	Moderate infection (S_3 to S_6)	0.08	0.088	0.072
P ₆	Severe infection (S_3 to S_7)	0.03	0.027	0.033
P ₇	Critical infection (S_3 to S_8)	0.01	0.009	0.011
P ₈	Recovery from severe state (S_7 to S_{10})	0.5	0.45	0.55
P ₉	Recovery from critical state (S_8 to S_{10})	0.12	0.108	0.132
P ₁₀	Death from severe state (S_7 to S_9)	0.5	0.45	0.55
P ₁₁	Death from critical state (S_8 to S_9)	0.88	0.868	0.892
P ₁₂	Recovery from asymptomatic, mild and moderate state	1.00	1.00	1.00

based on the relative difference in selected vulnerabilities between a country and Europe (it was not possible to get reference values for Wuhan, China). In deriving this vulnerability adjustment factor, data on tuberculosis prevalence, diabetes prevalence, HIV prevalence, total alcohol consumption per capita, smoking prevalence and protein energy malnutrition were considered as most likely to cause differences in infection outcomes. However, of these, HIV and diabetes prevalence were the only ones with an interpretable pattern on severe and critical cases. This is possibly because some of these estimates are also modelled. Tuberculosis and HIV prevalence are highly correlated across countries due to the comorbid nature of the two diseases; therefore, the tuberculosis effect is not lost, particularly in countries with a high prevalence of HIV. As a result, we factored in three vulnerabilities: per cent population >65 years, prevalence of HIV as a proxy for chronic communicable conditions and prevalence of diabetes as a proxy for chronic non-communicable chronic conditions. We did not consider sex, as we also assumed this was double counting of the vulnerabilities (men are more prone to these risk factors). These proxy indicators for vulnerability were assumed to contribute to the mortality pattern in a proportional manner, based on their respective burdens in each country. The indicator values per country were normalised between 0 and 1 and then averaged to yield a vulnerability adjustment factor.

Vulnerability adjustment factor =
$$\begin{pmatrix} \sum_{i=1}^{n} p_i \\ \frac{1}{n} \end{pmatrix}$$

where p_i refers to the normalised specific vulnerability factor. On average, the age factor reduces, HIV prevalence increases, and diabetes prevalence is largely similar to numbers with severe/critical disease; however, these trends vary by country.

In populating the model, data were triangulated from multiple sources. The population parameters were obtained from the United Nations Population Division database.⁶⁷ Data on school populations (tertiary, secondary and primary), prevalence of HIV, prevalence of non-communicable diseases, percentage of the population older than 65 years, average annual precipitation, road network and the proportion of population using at least basic sanitation were obtained from the World Development Indicators database.⁶⁸ The results are specific for each country, with the model run for a 52 week period. Please see the online supplementary material for further information. Neither patients nor the public were directly involved in the design of the study as it was primarily analytical.

RESULTS

Risk of exposure in countries in the WHO African Region

The RoE ranged from 0.002 to 0.461 for the Africa Region, with the relative values shown in figure 3. The



Figure 3 Risk of exposure of the population in countries in the World Health Organization African region.

RoE was higher in small states, whether island or mainland Africa, with these being seven of the top 10. These countries face the highest risk of transmission as their exposed population is high. Mauritius had the highest RoE. South Africa, Cameroon and Algeria were the only large countries among the top 10 highest RoE. The lowest RoE was mainly observed in sparsely populated countries, with Niger, Mauritania and Chad having the lowest RoE. Eritrea, Ethiopia, South Sudan and Madagascar were among the lowest 10 RoE countries that are relatively more densely populated than the other low RoE countries.

Expected impact of COVID-19 in the WHO African Region

We present the 1year prediction of the implications arising from widespread community transmission of SARS-CoV-2 in each WHO African country, which would result from failure of the containment measures (table 2). From the model, the region would have 223 281 401 (166 016 889–275 695 204) persons infected, representing 22% (16–26%) of the population. There would be 36967 532 (28 953 415–44 071 349) symptomatic cases, with 150078 deaths (82 735–189 579). The estimated infection mortality rate would be 0.06% (0.05–0.07%). A total of 4637 240 additional hospitalisations (3 592 443–5 529 368) would occur, of which 139 521 would be severe cases, requiring oxygen (81 876–167 044), and 89 043 critical cases, requiring ventilation support (52 253–106)

Table 2 Predicte	ed implications ov	er 52 weeks (of widespread cc	mmunity tra	Insmission of S/	RS-CoV-2 by	country in the	e World Heal	thOrganizat	tion African	Region
		Risk of			Distribution by	ype of infectic	u				
Country	Total population estimate (2019)	exposure(u - Iowest; 1 - highest)	Total estimated infections	Infections per capita	Non- symptomatic	Mild infections	Moderate infections	Severe infections	Critical infections	Deaths	NO OI hospital admissions
Algeria	42 228 429	0.216	25601319	0.26	21336366	2 036 229	2 036 229	26646	17389	29354	532730
Angola	30 809 762	0.085	9374921	0.30	7 811 646	744 559	744559	3762	2462	4066	184918
Benin	11 485 048	0:030	1 360 309	0.12	1 132 376	107 960	107960	725	482	792	27117
Botswana	2254126	0.024	213522	0.09	177688	16945	16945	139	92	152	4298
Burkina Faso	19751535	0.023	1811521	0.09	1 507 818	143757	143757	1154	476	1007	36 131
Burundi	11 175 378	0.136	4 921 938	0.44	4 103 321	391 135	391135	2215	1378	2336	97 466
Cabo Verde	543767	0.081	158736	0.29	132170	12606	12 606	118	77	129	3220
Cameroon	25216237	0.099	8 650 261	0.34	7 206 950	687117	687117	4791	2990	5074	172690
Central African Republic	4666377	0.052	924272	0.20	769514	73375	73375	560	369	611	18539
Chad	15477751	0.003	220640	0.01	183645	17505	17505	95	60	101	4356
Comoros	832 322	0.066	204 291	0.25	170159	16221	16221	97	65	106	4055
Congo, Dem Rep	84 068 091	0.051	16467004	0.20	13712570	1 307 261	1 307 261	8239	5428	8960	327410
Congo, Rep	5244363	0.053	1 053 489	0.20	877 184	83635	83 635	588	391	644	21 051
Cote d'Ivoire	25 069 229	0.080	7 192 921	0.29	5 992 069	571224	571224	3475	2304	3791	142872
Equatorial Guinea	1 308974	0.227	819289	0.63	683646	65171	65171	391	255	423	16286
Eritrea	4475000	0.006	118425	0.03	98560	9396	9396	59	37	62	2351
Eswatini	1 136 191	0.093	371606	0.33	309 591	29516	29516	197	128	213	7409
Ethiopia	109224559	0.009	4 254 002	0.04	3 539 877	337 525	337525	2497	1636	2713	85 139
Gabon	2119275	0.136	935 148	0.44	779146	74314	74314	200	455	762	18 990
Gambia, The	2280102	0.059	503 998	0.22	419826	40015	40015	199	129	213	9931
Ghana	29767108	0.041	4 783 076	0.16	3 981 908	379 663	379663	2789	1767	2978	95675
Guinea	12414318	0.059	2 764 527	0.22	2 302 216	219489	219489	1470	946	1581	55 094
Guinea-Bissau	1874309	0.055	388 820	0.21	323 799	30868	30868	193	127	210	7729
Kenya	51 393 010	0.030	6157172	0.12	5 125 709	488 663	488663	3445	1805	3340	122529
Lesotho	2108132	0.051	410895	0.19	341 999	32620	32 620	304	202	335	8335
Liberia	4818977	0.065	1 155 575	0.24	962 431	91753	91753	585	386	637	22 991
Madagascar	26262368	0.026	2 729 032	0.10	2271625	216577	216577	1562	872	1563	54413
Malawi	18143315	0.050	3 463 021	0.19	2 883 220	274913	274913	2019	1343	2213	69 341
Mali	19077690	0.015	1 154 252	0.06	960 677	91588	91588	602	398	656	22 981
Mauritania	4 403319	0.003	60309	0.01	50053	4785	4785	114	72	126	1335
Mauritius	1 265 303	0.461	1 097 013	0.87	916588	87424	87 424	1005	358	837	22 345
											Continued

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		Risk of			Distribution by t	type of infectic	u				
Country	Total population estimate (2019)	exposure(0 - lowest; 1 - highest)	Total estimated infections	Infections per capita	Non- symptomatic	Mild infections	Moderate infections	Severe infections	Critical infections	Deaths	No of hospital admissions
Mozambique	29 495 962	0.048	5 380 072	0.18	4 479 773	427 085	427085	2773	1843	3031	107116
Namibia	2448255	0.030	289700	0.12	240624	22992	22 992	475	315	536	6308
Niger	22 442 948	0.002	166248	0.01	138369	13189	13 189	71	46	77	3283
Nigeria	195874740	0.081	56941648	0.29	47438888	4 522 073	4522073	25771	17103	28107	1128172
Rwanda	12 301 939	0.128	5197581	0.42	4 332 265	413 005	413005	2599	1689	2809	103409
Sao Tome and Principe	211028	0.227	132 079	0.63	110211	10506	10506	64	42	69	2627
Senegal	15854360	0.067	3 937 580	0.25	3279437	312 654	312654	2136	1293	2225	78467
Seychelles	96762	0.449	83212	0.89	69382	6631	6631	164	78	159	1834
Sierra Leone	7650154	0.064	1810218	0.24	1 507 815	143729	143729	843	501	868	35 839
South Africa	57 779 622	0.126	24023691	0.42	20007604	1 908 888	1 908 888	21 692	14028	23661	493853
South Sudan	10975920	0.016	725250	0.07	603 553	57549	57549	436	272	462	14 520
Tanzania	56318348	0.050	10663921	0.19	8 8 7 9 2 9 9	846553	846553	5885	3618	6180	212676
Togo	7 889 094	0.033	1 038 493	0.13	864616	82423	82 423	506	322	539	20609
Uganda	42 723 139	0.037	6213392	0.15	5174085	493170	493170	2724	1470	2670	122555
Zambia	17 351 822	0.042	2841111	0.16	2 365 797	225520	225520	1390	769	1381	56284
Zimbabwe	14 439 018	0.045	2515899	0.17	2 094 954	199713	199713	1257	775	1319	49 963
Africa region	1 064 747 476	0.081	231 281 401	0.22	192 651 016	18369484	18369484	139521	89043	150078	4637240
Best case scenario	1 064 747 476	0.073	166 016 889	0.16	134 792 528	14409643	14 409 643	81 876	52253	82 735	3592443
Worst case scenario	1 064 747 476	0.089	275 695 204	0.26	260 049 209	21898853	21 898 853	167 044	106 599	189579	5529368

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Figure 4 Impact of different variables, constituting the gathering factor, on overall deaths.

599). The model strength is in provision of country specific values based on their attributes of disease transmission.

Sensitivity analysis

Given the high uncertainty with several probabilities, it is important to see how the predictions change when varied. We specifically explored variations in four probabilities: RoE, specifically the gathering factor, as this had the least empirical evidence supporting its inclusion; the attack rate value; the effect of NPIs; and the proportion of asymptomatic cases. We used the number of deaths as the proxy for results (150078 (82 735–196 645)). We tested the effect on the overall number of deaths when each of the indicators constituting the gathering factor was removed (figure 4).

None of the indicators constituting the gathering factor significantly changed the results when eliminated, and the total number of deaths remained within the projected range. Children in primary school had the largest positive change (2.3%, 3758 deaths), while the proportion of the population living in slums had the largest negative change (3.2%, 4778 deaths). Similarly, changing the proportional implementation of the NPIs did not change the results significantly. Reducing the implementation effectiveness of NPIs to 75%, 50%, 25% and 0% changed the number of deaths by 3.8%, 7.5%, 11.1% and 14.7%, respectively. Implementing lockdown interventions only, without improved hygiene practices, increased deaths by 6.4%, while only hygiene practices without any lockdown interventions increased them by 8.5%.

Furthermore, changing the attack rate did not have an effect on the results, and raising it did not increase the results beyond the reported best and worst case ranges. An increase to 60% on the attack rate augmented the number of deaths to 166 148, which lies within the range of the best and worst cases of the results.

Finally, the proportion of asymptomatic cases was varied to see what proportion it needs to be for the study results to change. The model is based on 80% of infected persons being asymptomatic, or 88% if asymptomatic and mild disease patients are combined. To remain within the range of possible results predicted by the model, the

proportion of asymptomatic cases ranges between 74% and 89% (or 84.4–93.4% of persons having asymptomatic/mild disease).

DISCUSSION

Comparability with current knowledge

The results suggest a slower rate of infection compared with other areas of the world, together with lower mortality rates among the infected. The calculated reproduction rate, R_0 , of 1.7 overall (range 1.5–1.8 depending on country), is within the known range for COVID-19 of 0.41–2.39,²⁹ and consistent with that from an earlier model.²⁸ This suggests that while the rate of infection is slower, the outbreak would continue until R_0 drops to <1.

The probabilities used were corroborated in the sensitivity analysis, where the ones with the most uncertainty of evidence were shown to still predict the study results, even when extensively varied. The range of asymptomatic cases for which the results are sensitive (74–89%, or 84.4– 93.4% of asymptomatic/mild persons) is plausible given the data emerging from the seroepidemiological studies. This could also contribute (together with possible low reporting) to the very low numbers of cases seen within the WHO African Region. At the time of writing, 25 of the 47 countries in the region had fewer than 100 cases.

The RoE is also aligned with what is seen with SARS-CoV-2 in the region. The top 10 countries with the highest RoE are seven small states, where transmission is easier due to size, together with South Africa, Algeria and Cameroon. As of 30 April, the top three countries with the most cases were South Africa (5350), Algeria (3848) and Cameroon (1806).

The model predicts fewer deaths primarily because of the effects of the RoE, reducing overall cases in the first year. The correction made for regional vulnerabilities also further reduces the proportion of severe/critical disease observed in the African Region. The effect of the vulnerability correction is attributable to a younger population and the region's similarity in non-communicable disease burden with other affected countries. High access and use of interventions targeted at chronic communicable diseases also reduce the potential impact of these factors. The region will have fewer deaths, but occurring more in relatively younger age groups, among people previously considered healthy, due to undiagnosed noncommunicable diseases. We already see this picture emerging based on the characteristics of persons affected in the African Region compared with other regions.

The case fatality for COVID-19 is still difficult to estimate globally, as countries are using different testing strategies for identification of cases and have different RoE. However, the infection mortality rate estimate of 0.06% is consistent with the current expectations of it being <0.1%.⁴² We expect the regional infection mortality rate to be lower because of the vulnerability index. However, the case fatality rate, particularly when based on deaths among known cases with moderate to critical symptoms, may be high due to weaker mitigation measures, especially pertaining to care of severe and critically ill persons.

Impact on health system capacity

Deaths due to COVID-19 will not significantly change crude death rates; the regional crude death rate is 8.9/1000 persons.⁶⁹ It will, however, lead to more deaths from other conditions due to changing health seeking behaviours among the population, who may perceive health service delivery areas as hotspots of infection, diversion of time, money and other health resources and/or overwhelmed health services unable to cope.^{70–72} Predicted hospitalisations would lead to an additional 69550600 bed days assuming 15 day admissions, translating to 60 bed days per 1000 persons, ranging from 284 in the Seychelles to 2 in Niger. These hospitalisations would be widespread, with some occurring in areas where access to services is poor and inaccessible to the most disadvantaged.

The creation of special COVID-19 service areas to reduce nosocomial infections would be difficult to establish and sustain in the African Region. Facilities would need to be effectively prepared with diagnostic and functional triage systems, many of which are currently ineffective. Tertiary care capacity is also limited, due to an historical focus on provision of a limited set of primary care mother and child interventions, at the expense of basic emergency interventions.⁷³ Given the low coverage of service provision units and death reporting, coupled with diagnostic challenges, many cases may also go undocumented.

These system capacity challenges highlight the need to ensure the success of the containment measures to avoid the need for mitigation measures that, despite the relatively fewer cases expected in the region, will be difficult to institute. These containment measures, specifically physical distancing, should be implemented in the context of Africa's unique socioeconomic challenges and limited economic safety nets.⁷⁴ They nonetheless are already appearing to be effective, with cases stagnating/ reducing in some countries, such as South Africa, despite its high RoE, at the time of writing.

Challenges in building the model

Effective predictive modelling relies on the development of assumptions and availability of quality data. Both presented challenges for the development of a model useful for countries in the region. Innovative ways of overcoming these challenges therefore had to be used and have been presented through this paper. The region lacks data in many areas, relying on values modelled with different assumptions that make it difficult to use for this purpose. As a result, selection of indicators to be used was not only based on the indicators' appropriateness, but also the availability of reliable data across the countries. Different indicators were tested, especially in the generation of the RoE and the regional vulnerability factor, to arrive at those used. A focus on capacity for locally generated data is urgently needed for countries in the region.

CONCLUSION

The picture we have modelled is based on a scenario of widespread and sustained community transmission of SARS-CoV-2 in each country of the WHO African Region. It introduces unique ways of overcoming traditional challenges with modelling in the region, specifically in improving accuracy in the number of infections and capturing the effect of a different disease burden. The model predicts that the region's unique socioecological context is contributing to reductions in the number of cases, with infections spread over a longer time period. Its specific vulnerabilities are also changing the numbers, age focus and nature of events arising from infection in most countries. Moving forward, more analysis and research, particularly around refinement of the RoE and vulnerability adjustments within each country, would allow for further improvement in estimates.

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agencies depending on the indicator. The tool consolidating and analysing these data is publicly available with all countries of the World Health Organization African Region and upon request.

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