

Forecasting the effect of HIV-targeted interventions on the age distribution of people with HIV in Kenya

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Objectives: To provide accurate forecasts of the age distribution of people with HIV (PWH) in Kenya from 2025 to 2040.

Design: Development of a compartmental model of HIV in Kenya, calibrated to historical estimates of HIV epidemiology.

Methods: We forecasted changes in population size and age distribution of new HIV infections and PWH under the status quo and under scale-up of HIV services.

Results: Without scale-up, new HIV infections were forecasted to fall from 34 000 (28 000–41 000) in 2,025 to 29 000 (15 000–57 000) in 2,040; the percentage of new infections occurring among persons over 30 increased from 33% (20–50%) to 40% (24–62%). The median age of PWH increased from 39 years (38–40) in 2025 to 43 years (39–46) in 2040, and the percentage of PWH over age 50 increased from 26% (23–29%) to 34% (26–43%). Under the full intervention scenario, new infections were forecasted to fall to 6,000 (3,000–12 000) in 2,040. The percentage of new infections occurring in people over age 30 increased to 52% (34–71%) in 2,040, and there was an additional shift in the age structure of PWH [forecasted median age of 46 (43–48) and 40% (33–47%) over age 50].

Conclusion: PWH in Kenya are forecasted to age over the next 15 years; improvements to the HIV care continuum are expected to contribute to the growing proportion of older PWH.

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AIDS 2024, 38:1375–1385

Keywords: Africa, aging, mathematical models

Introduction

As a result of advances in the effectiveness and coverage of antiretroviral therapy (ART), the global population of people with HIV (PWH) is aging. A PWH who initiates ART today has the same life expectancy as an HIV-negative individual of the same age [1]. This longer lifespan has shifted the global HIV age structure: one in five adults living with HIV today is over age 50; this proportion has doubled from one in ten adults in 2005 [2]. Modeling studies suggest that the number of PWH over the age of 50 will continue to increase over time [3].

As this population ages, countries must consider how to provide comprehensive care to older individuals living

with HIV. These PWH are at greater risk for many age-associated comorbidities, including cardiovascular, metabolic, pulmonary, and renal conditions; as well as non-AIDS-defining cancers [4–12]. Such multimorbidity leads to increasingly complex care management and increased likelihood of polypharmacy, or the simultaneous use of multiple medications [13,14]. Although research on age-related comorbidities and the age distribution of PWH is gaining traction, studies often focus on high-income settings [15–17].

Kenya has one of the largest HIV epidemics globally, with an estimated 1.4 million PWH in 2021 [2]. Although HIV care efforts have been largely successful (over 75% of Kenyan PWH are virally suppressed [2]), management of

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Received: 2 October 2023; revised: 27 February 2024; accepted: 21 March 2024.

other comorbidities remains inadequate. In Kenya, an estimated 62% of PWH were living with at least one noncommunicable disease (NCD) in 2020 [18]. However, NCD awareness is low: in 2015, 88% of Kenyans had never been measured for raised blood sugar and 56% had never been measured for hypertension [19]. Therefore, a key component of investing in new care models for HIV is estimating the population size and age distribution of PWH that will require these NCD services in the future.

Therefore, we sought to forecast the effect of demographic shifts and HIV-targeted interventions on the population size and age distribution of PWH in Kenya through 2040. Specifically, we examined demographic changes under a status quo scenario representing current trends in HIV care, and a ‘full intervention’ scenario representing broad scale-up of HIV care continuum interventions.

Methods

We developed a dynamic, compartmental model of HIV transmission among the general population in Kenya (Fig. 1). A compartmental model is a type of mathematical model that simplifies the population into homogenous compartments and models the average behavior within each. Compartments were defined to

characterize the HIV continuum of care: uninfected; infected but undiagnosed; diagnosed but unengaged in care; engaged in care but unsuppressed; and engaged in care and virally suppressed. We defined engagement in care as being on ART, based on the UNAIDS definition of receiving an ART prescription in the past 28 days [2]. We stratified the population according to age (5-year bands from 0 to 80 and an 80+ group) and sex (male/female). The full set of differential equations governing the model can be found in Appendix Text S1, <http://links.lww.com/QAD/D158>.

Model calibration

The model was initiated in 1970 and calibrated through 2021. Calibration is defined as the process of finding a set of input parameter values that results in a simulated HIV epidemic closely representing the observed epidemic. Parameters are the numerical variables that define a model and govern its function. Two general data types informed the model: input data for prior distributions of model parameters (e.g. the rate at which individuals disengage from HIV care) and calibration target data (e.g. the number of new HIV infections reported in Kenya annually by age and sex).

We assessed model fit via a defined likelihood function quantifying how well each simulation reproduced the calibration targets. Using an Adaptive Metropolis sampler [a Markov-chain Monte-Carlo (MCMC) approach [20,21]], the calibration ran simulations with different

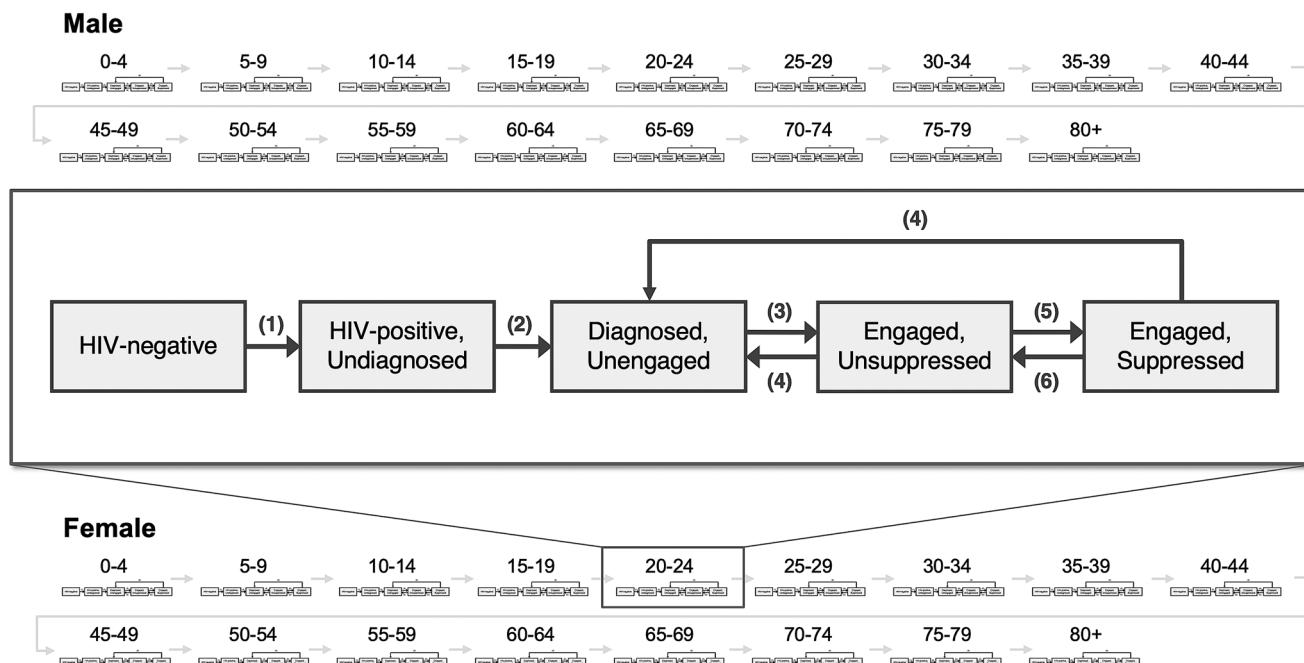


Fig. 1. Model structure. Compartmental model representing the general HIV epidemic in Kenya by sex (male, female), 5-year age groups, and relevant HIV states: HIV-negative; HIV-positive but undiagnosed; diagnosed but unengaged in care; engaged in care but unsuppressed, and engaged in care and virally suppressed. Transitions along the continuum of care include (labeled in figure): (1) HIV incidence, (2) HIV diagnosis, (3) engagement in care, (4) disengagement from care (with separate rates of disengagement from the suppressed versus unsuppressed compartments), (5) gain of viral suppression, and (6) loss of viral suppression.

sampled values of each parameter and weighted these simulations according to the likelihood function. We ran the MCMC simultaneously across four chains, with a burn-in period of 50 000 iterations followed by 50 000 iterations for analysis in each chain. The resulting 200 000 simulations (across four chains) were thinned to a final set of 1000 simulations by selecting every 200th simulation.

Input parameters and prior distributions

Model input parameter values came from available literature and survey data (see Table 1; a complete list of parameters is provided in Table S1, <http://links.lww.com/QAD/D160>). We parameterized the model using the simplest set of parameters that could still replicate the observed epidemic and continuum of care by age and sex over the calibration period.

Parameters governing continuum of care transitions (i.e. the probabilities of receiving an HIV test [22–24], starting or stopping ART [2,25], becoming virally suppressed [26], or losing viral suppression [27]) came from Kenya-specific sources, such as the Kenya Demographic and Health Surveys (KDHS), or cohort studies conducted among Kenyan populations. For testing, we fit logistic models to KDHS data on the probability of receiving an HIV test, with separate models by sex and by age for those under versus over 17. We included

covariates for calendar year and each year of age over 17; the resulting odds ratios are in Table 1. For engagement, we fit a logistic model to UNAIDS data on the number of individuals starting ART, with separate slopes for the time periods of preuniversal ART, rollout of universal ART, and postuniversal ART. These slopes are represented as odds ratios in Table 1. Annual probabilities were transformed to per capita rates by assuming the probabilities were derived from exposure to a constant hazard, that is, $P = 1 - S(t) = 1 - e^{-(\text{rate} * \text{time})}$, where $S(t)$ is the survival function at time t .

We assumed sexual partnerships formed with a probability related to the age difference between two partners, based on a phylogenetic analysis of sexual partnerships in South Africa [28]. We allowed the relative risk of HIV transmission to vary by age as a function of the proportion in each age group reporting multiple partners or high-risk sex practices (i.e. condomless sex) [22–24,29]. Maternal–fetal transmission risk came from Kenya's Framework for Elimination of Mother-To-Child Transmission of HIV [30] for the modern era and a study of maternal–fetal risk for the pre-ART era [31].

During calibration, parameter values were sampled from prior distributions with corresponding uncertainty ranges; in most cases, this was the 95% confidence

Table 1. Prior (pre-calibration) values of key parameters^a.

Parameter	Value (median prior, with sampling range)	Reference
Transmission ^b		
Transmission rates by age/sex/time	(Calibrated)	(Calibrated)
Testing (probability of receiving an HIV test within the past year)		
Female, age 35–39, 2015	0.52 (0.24–0.73)	KDHS 2003 [22], 2008 [23], 2014 [24]
Female, age 17 and younger, 2015	0.35 (0.13–0.63)	
Yearly change in odds of receiving a test, for individuals over 17 (odds ratio)	1.26 (0.32–5.04)	
Yearly change in odds of receiving a test, for individuals age 17 and under (odds ratio)	1.23 (0.31–4.90)	
Change in odds of receiving a test for every one-year increase in age, for individuals over 17 (odds ratio)	0.96 (0.24–3.86)	
Ratio of testing rates among male versus female	1 (0.25–4)	(Calibrated)
Engagement (annual probability of starting ART)		
Female, 2015	0.32 (0.08–1)	AIDSinfo [2]
Yearly change in odds of starting ART prior to universal ART, 2016 (odds ratio)	1.34 (0.34–5.36)	
Yearly change in odds of starting ART during rollout of universal ART, 2016–2017 (odds ratio)	1.68 (0.42–6.70)	
Yearly change in odds of starting ART after initial rollout of universal ART, 2017 (odds ratio)	1.12 (0.28–4.49)	
Ratio of engagement rates among male versus female	1 (0.25–4)	(Calibrated)
Disengagement (annual probability of stopping ART) ^c		
Among suppressed	0.098 (0.026–0.329)	Lee et al. [25]
Among unsuppressed	0.13 (0.035–0.418)	Lee et al. [25]
Suppression (average time to suppression, years) ^d		
In 1993	1.49 (0.38–5.78)	Njuguna et al. [26]
In/after 2003	1.49 (0.38–5.78)	Njuguna et al. [26]
Ratio of suppression rates among male versus female	1 (0.25–4)	(Calibrated)
Viral rebound (average time to loss of suppression, years)		
All years	5.07 (1.3–19.7)	Maina et al. [27]

ART, antiretroviral therapy; KDHS, Kenya Demographic and Health Survey.

^aSelect parameters only; see for full parameterization of all parameter values, <http://links.lww.com/QAD/D160>.

^bA composite of number of sexual encounters and rate of transmission per encounter.

^cSame for all ages and sexes; constant over time

^dSame for all ages.

interval of a lognormal distribution. We used lognormal distributions to reflect the intuition that rates function on a multiplicative scale; for example, simulating a rate of starting ART twice what is estimated from the literature would be as likely as a rate half of the literature estimate.

Calibration targets

The model was calibrated to HIV incidence (1990–2021), prevalence (1990–2021), mortality (1990–2021), status awareness (2015–2021), care engagement (2015–2021), and viral suppression (2019–2020) as reported by UNAIDS [2]. Incidence and prevalence were reported as totals, by age, and by age*sex; awareness, engagement, and suppression were reported as totals and by age*sex; and HIV mortality was reported as totals and by age.

General population size was calibrated to UN World Population Prospects data from 1950 to 2021, reported as totals and by age, sex, and age*sex [32]. See Appendix Text S2, <http://links.lww.com/QAD/D159> and Table S2, <http://links.lww.com/QAD/D161> for detailed reporting of data and likelihood methodology.

Modeled interventions

Calibrated models were followed to 2040 under two scenarios: a ‘status quo’ scenario assuming no additional scale-up of HIV interventions, but allowing current trends in transmission rates, HIV testing, and care engagement to continue, and a ‘full intervention’ scenario with the following improvements to the HIV care continuum:

1. HIV testing: 75% of the population tested for HIV annually [average preintervention value: 32% (95% credible interval: 27–37%)].
2. Engagement in HIV care: 90% of diagnosed, unengaged PWH engage in care annually [preintervention: 40% (23–60%)].
3. Viral suppression gain: 90% of engaged, unsuppressed PWH gain suppression annually [preintervention: 88% (71–98%)].
4. Maintain viral suppression: 90% of engaged, suppressed PWH maintain suppression annually [preintervention: >90% (83–97%)].
5. Retention: 90% of all engaged PWH (regardless of suppression status) remain engaged annually [preintervention: >90% (91–98%)].

Preintervention values of each intervention are population-weighted averages; however, model values varied within strata of age and sex. For some age-specific and sex-specific strata of certain interventions (especially maintenance of suppression and retention), the preintervention value was greater than the intervention target; in these cases, the preintervention value was carried forward in the intervention scenario as well. All interventions began in 2025, scaled up linearly to reach

full effect by 2030, and remained at the target level through 2040. We chose a linear scale-up for the simplicity of representing a phased roll-out, as opposed to assuming immediate implementation at full scale.

Outcomes

Our primary outcomes were the median forecasted age of all PWH and PWH engaged in HIV care in 2040 under both the ‘status quo’ and ‘full intervention’ scenarios (overall and by sex). We also reported the median age of people with incident infections, the median age of individuals entering care, the number of new HIV infections, and total prevalence in 2040. For each outcome, we calculated the median value and the 95% credible interval (2.5th and 97.5th quantiles) from the final set of 1000 simulations.

Sensitivity analyses

We conducted sensitivity analyses to identify parameters with the strongest influence on the percentage of PWH over age 50 in 2040. We assessed the monotonicity of these parameter–outcome relationships visually using scatterplots and calculated partial rank correlation coefficients (PRCCs, a measure of the correlation between each parameter and the outcome). To complement the PRCCs, we ranked simulations based on each parameter value and compared the percentage of PWH over age 50 from the 25% of simulations with the lowest versus highest values of each parameter. This method of comparing outcomes helps depict how much results might change if key parameter values are changed, and has been described elsewhere [33].

Results

New HIV infections and total prevalence

Assuming continuation of status quo, annual new HIV infections in Kenya were forecasted to fall from 34 000 [95% credible interval (CrI): 28 000–41 000] in 2025 to 29 000 (15 000–57 000) in 2040 (Fig. 2, red line). Assuming implementation of the full intervention, new HIV infections fell to 6000 (3000–12 000) by 2040, representing 78% (54–90%) fewer incident infections than in the status quo scenario. Under the status quo, HIV prevalence fell from 1.44 million (1.39–1.50 million) in 2025 to 1.38 million (1.27–1.65 million) in 2040. Under the intervention scenario, prevalence fell to 1.11 million (1.01–1.25 million) by 2040, representing 20% (10–30%) fewer prevalent infections than in the status quo scenario. (See Figures S1–2, <http://links.lww.com/QAD/D163>, <http://links.lww.com/QAD/D164> for full calibration period 1990–2020 and additional stratifications.)

Age structure of people with HIV

The median forecasted age of PWH (both diagnosed and undiagnosed) in 2025 was 39 (38–40), and 376 000

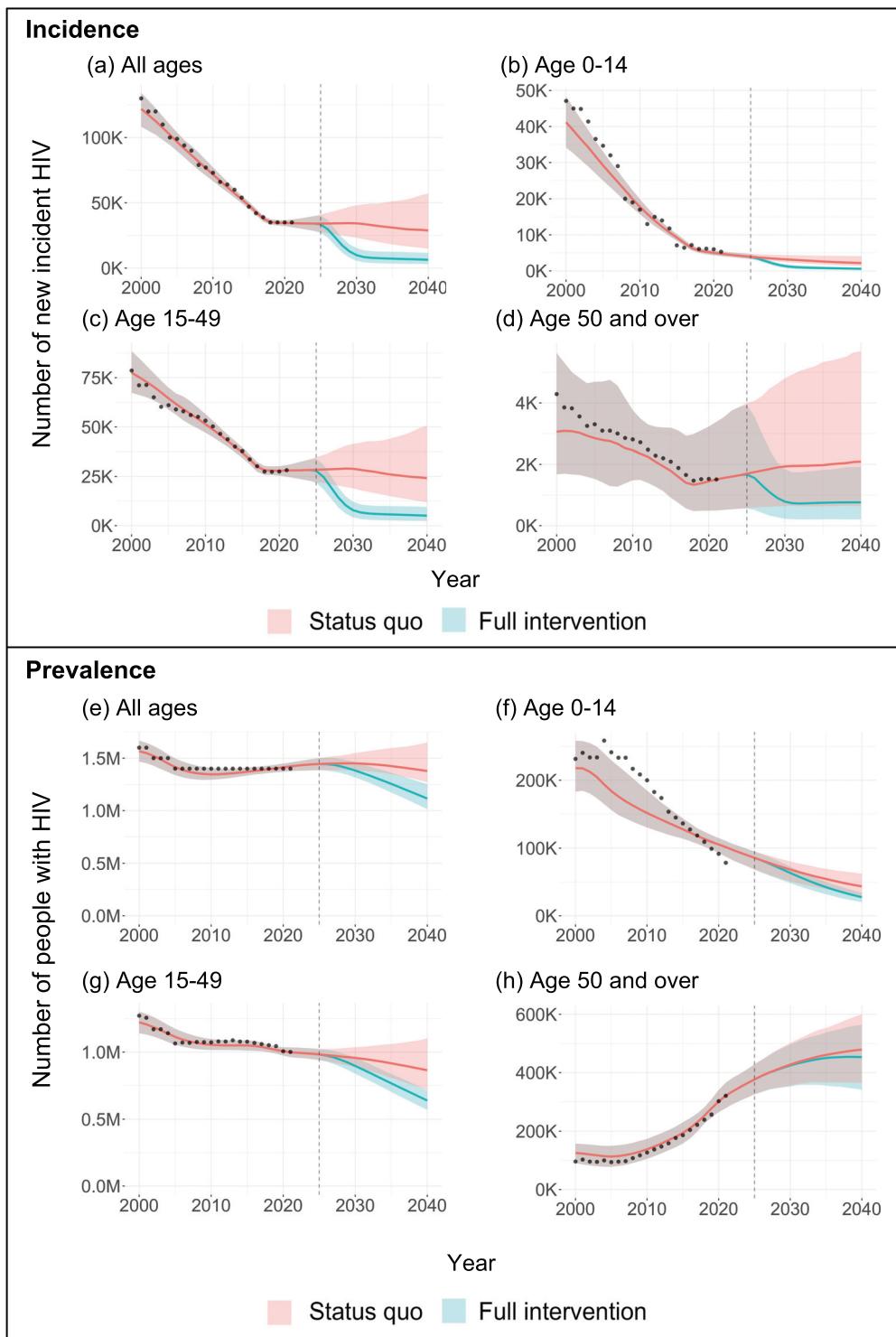


Fig. 2. Forecasted HIV incidence (top panel) and HIV prevalence (bottom panel) under status quo scenario versus full combined interventions, overall and by age, overlaid with calibration data. The median value across 1000 model simulations is shown as a red line for the 'status quo' scenario and green line for the 'full intervention' scenario, with 95% credible intervals shown as the shaded ribbons. Black dots indicate calibration target data. The vertical dashed line indicates the intervention start year in 2025, at which point the two scenarios diverge.

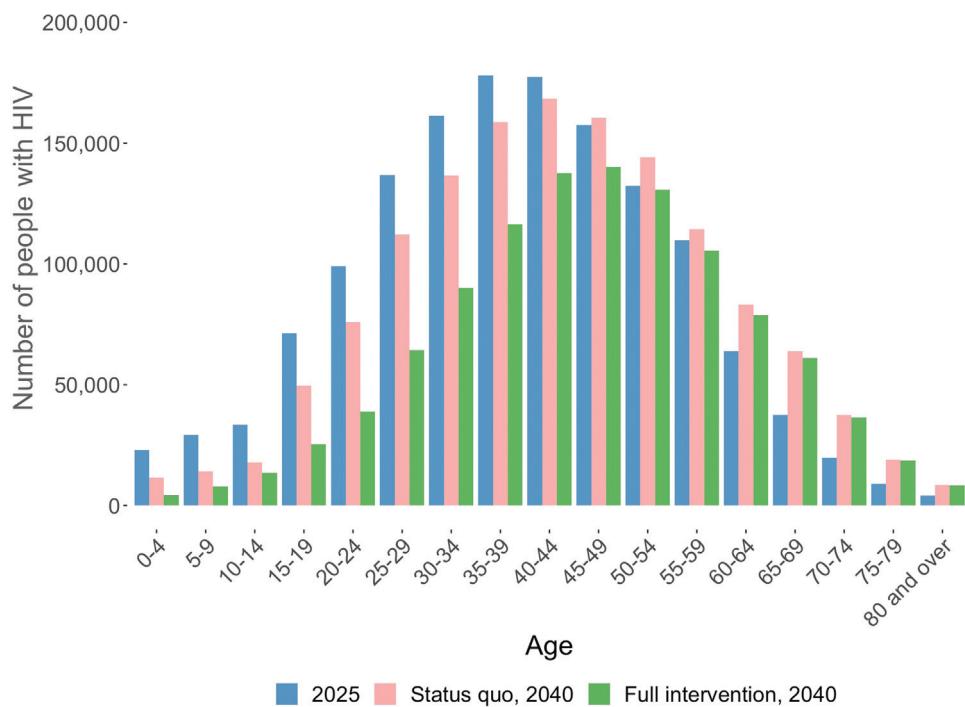


Fig. 3. Age structure of people with HIV in Kenya. Age structure of people with HIV in Kenya in 2025, prior to intervention (blue, left bars), in 2040 under the status quo scenario (pink, center), and in 2040 under the full intervention scenario (green, right).

(325 000–429 000) PWH were over age 50, representing 26% (23–29%) of all PWH. Without additional intervention, the median age increased to 43 (39–46) in 2040, with 479 000 (364 000–600 000) over age 50 (or 34% (26–43%) of all PWH, Fig. 3; Table S3, <http://links.lww.com/QAD/D162>). Under the full intervention scenario, the age structure of PWH was forecasted to shift even older, reaching a median age of 46 (43–48) by 2040, with 453 000 (341 000–564 000) over age 50, or 40% (33–47%) of all PWH.

We also found a shift in the age distribution of new HIV infections: in 2025, only 33% (20–50%) of new infections were forecasted to occur among individuals over age 30. Without intervention, this percentage increased to 40% (24–62%) by 2040, and further to 52% (34–71%) under the full intervention scenario (Figure S4, <http://links.lww.com/QAD/D166>).

Relative to the population of PWH, the forecasted age distribution of the Kenya's general population was much younger, with a median age of 22 (21–22) in 2025 and 25 (24–26) in 2040 (Figure S3, <http://links.lww.com/QAD/D165>). Additionally, the percentage of the general population over age 50 was considerably lower in comparison [10% (9–11%) in 2025 and 13% (11–15%) in 2040].

Age structure of people engaged in HIV care

As most PWH in Kenya are engaged in care, the age structure of people engaged in HIV care was similar to

that of all PWH: the median forecasted age shifted from 40 (39–42) in 2025 to 44 (41–47) in 2040 without any intervention and 46 (43–49) with full intervention (Figure S5, <http://links.lww.com/QAD/D167>). Of those engaged in care, 28% (25–31%) were over 50 in 2025, increasing to 36% (28–44%) in 2040 without intervention and 41% (34–48%) with full intervention.

In 2025, the median age of individuals entering into care (either for the first time or re-entering care after disengagement) was 35 (32–38), with 63% (57–69%) over age 30. In the full intervention scenario, the median age of those entering care was 44 (41–47) by 2040, with 81% (77–85%) over age 30.

Differences in age structure by sex

Despite a similar median age of women and men living with HIV in 2025 (38 [37–40] for women, 41 [39–42] for men), the percentage of HIV-positive men over age 50 was forecasted to grow more rapidly in the next 15 years, from 28% (24–31%) in 2025 to 40% (32–48%) in 2040 [versus 25% (22–28%) to 31% (22–39%) among women, Figures S6–S7, <http://links.lww.com/QAD/D168>, <http://links.lww.com/QAD/D169>]. This growth is primarily due to the older age at HIV infection among men (29 [26–34]) versus women (22 [19–24]).

Sensitivity analysis

Under the intervention scenario, the parameters most strongly associated with the percentage of PWH over age 50 in 2040 were related to general (i.e. non-HIV-specific)

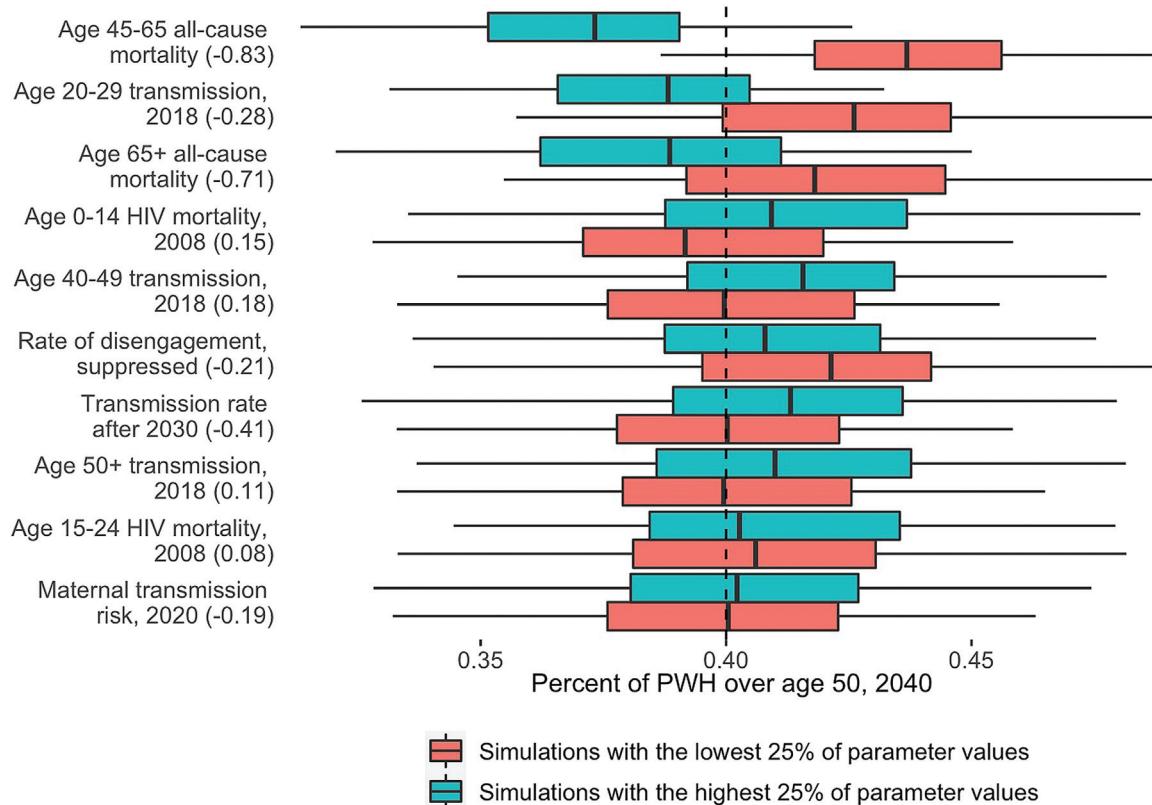


Fig. 4. Sensitivity analysis for parameters most strongly associated with the percentage of people with HIV (PWH) over age 50. The red (lower) bars represent the estimate of the change in this percentage for the 25% of simulations with the lowest values of each parameter in 2040; the blue (upper) bars represent the estimate for the 25% of simulations with the highest values. The solid vertical line of each bar represents the median, the box spans the interquartile range, and the whiskers span the 95% credible interval. Parameters are ordered by the univariate difference between the lowest and highest estimates, that is, in one-way sensitivity analyses, age 20–29 transmission was the second most influential parameter. The vertical dashed line represents the median value from the main analysis. Parameters are labeled with their partial rank correlation coefficient in parentheses, a multivariate estimate of the correlation between each parameter and the outcome. Values closer to 1 or -1 indicate stronger correlation.

mortality by age (PRCCs: -0.83, mortality age 45–65 years old; -0.71, mortality age 65 and older, Fig. 4). In the 25% of simulations with the lowest general mortality age 45–65, the percentage of PWH over age 50 in 2040 was 44% (39–49%). Conversely, in the 25% of simulations with the highest general mortality age 45–65, the percentage of PWH over age 50 was 38% (32–43%). We conducted an additional sensitivity analysis with select model outputs; these results are shown in Figure S8, <http://links.lww.com/QAD/D170>.

Discussion

This analysis illustrates the effect of demographic shifts and HIV-specific interventions on the age distribution of PWH in Kenya over the next 15 years. Assuming no additional scale-up of HIV interventions, new infections are forecasted to fall by 2040, a product of historically strong HIV programming and a hopeful sign for future

health system needs. This reduction in new HIV infections – typically clustered among younger ages – combined with longer survival among persons receiving ART will lead to an aging HIV population: our results suggest that by 2040, the population of PWH over age 50 will grow by nearly one-third, from 376 000 to 479 000. Moreover, this population over 50 will constitute a greater percentage of all PWH, increasing from 26 to 34%.

The addition of HIV-specific interventions was forecasted to further reduce new infections and prevalence of HIV. Because younger PWH tend to have greater room for improvement along the care continuum, interventions that brought all ages up to a common level (as implemented in this model) led to an even older population living with HIV by 2040.

Men with HIV were forecasted to have an older median age than women with HIV in 2040, both with and without intervention. The pattern of a higher age at

infection among men versus women has been seen in other countries across eastern and southern Africa [34] and is important for both HIV-specific programming and NCD screening, given the elevated risk of cardiovascular disease in men [35].

This analysis adds to the literature on the aging HIV population and, importantly, focuses on a lower middle-income country. Research on healthy aging among PWH is growing; however, studies in African populations remain sparse [17]. Although other models have similarly evaluated HIV outcomes by age, many models in Africa have not included finer age strata above 50 [36,37]. Those studies that have done so have found both increases in prevalence as well as greater progress towards reaching care continuum targets among those over 50 [38,39] – both results that align with our findings.

Additional research in this setting will hopefully lead to a reevaluation of the care needs of African PWH. For instance, recent studies of NCDs among PWH in Africa have found high rates of elevated blood pressure, dysglycemia, renal insufficiency, and obesity [40–44]. Combined with factors such as increased inflammation from viral replication, PWH may have increased risk for cardiovascular events such as myocardial infarction and stroke [5,6,8,45]. Therefore, interventions mitigating noncommunicable risk factors among PWH will likely rise in importance over time. Our sensitivity analysis found that the parameters most strongly influencing the percentage of PWH over 50 were those related to all-cause mortality, suggesting that future challenges of HIV care may focus more on managing other comorbidities driving general mortality than managing HIV itself.

Lessons learned from other settings with aging PWH suggest prioritizing overall health-related quality of life by integrating HIV and NCD screening and treatment [46–65]. Different service delivery modalities have proven effective in different settings: in western Kenya, the Academic Model Providing Access to Healthcare (AMPATH) program has leveraged experience in managing HIV to address NCDs by training nurses and clinical officers in diabetes and hypertension care [50]. The Sustainable East Africa Research in Community Health (SEARCH) program in rural Uganda and Kenya has improved outcomes through mobile, multi-disease community health campaigns [66–68]. Therefore, our analysis can be used to compare the potential impact of such integration options at a national level in Kenya.

Our analysis is subject to limitations. Although the UNAIDS data are the official estimates for Kenya and we believe they represent the best estimates of HIV epidemiology in the country, these estimates of incidence, prevalence, and mortality are in fact themselves generated from the Spectrum/AIM model [69,70]. Using model-generated projections as calibration data is a limitation;

however, Spectrum has been validated across settings. Furthermore, a comparison of incidence estimation approaches in Kenya found that Spectrum estimates compared reasonably well with survey-derived and assay-derived methodologies [71]. We have used Spectrum in previous modeling analyses in Kenya [72,73], but developed this new model to allow more flexibility and control over model assumptions, parameters, and calibration.

We did not conduct model validation against external datasets in Kenya, as we aimed to include as much data as possible in the calibration targets themselves. However, our results can be compared with future estimates of Kenya's HIV age distribution – particularly among finer age strata over 50 – as they become available. As our calibration targets are all Kenya-specific, the generalizability of our results to other settings may require model recalibration.

We forecast 15-year trends in the HIV epidemic using data from the recent past; findings should be interpreted with consideration for the challenges of forecasting HIV incidence beyond the immediate term. In the United States, for instance, incidence patterns have changed in the past decade, owing to a concentration and persistence of transmission in key populations, such as men who have sex with men and persons who inject drugs [74]. Although we model HIV in the general population, effective HIV programs must reach key populations (which vary geographically, including female sex workers in Kenya, e.g.) who may benefit the most.

Kenya's programs will have to maintain annual rates of engagement, viral suppression, and retention as proposed in this article in order to reach the UNAIDS 95–95–95 targets [75]. However, though we described the potential impacts of meeting the targets, this analysis did not cover the specific strategies that Kenya may use to reach them. Literature on best practices – such as CDC's compendium of evidence-based interventions or UNAIDS' checklist for developing a national strategic plan – will be crucial to achieving these targets [76,77]. Improving testing rates from 32 to 75%, for instance, will require differentiated and innovative modalities to reach those who are missing, such as those focused on men, adolescents, or key populations. Our decision to linearly scale up interventions to full effect by 2030 may also not reflect implementation realities, with harder-to-reach risk groups often requiring more effort.

As our model collapses rural and urban settings, we are modeling the average behavior across all settings in Kenya. Our predictions might overstate or underestimate the effectiveness of interventions depending on how future trends in HIV epidemiology deviate from past patterns and how interventions are differentially implemented in these settings. With devolution of power from the

national government, Kenyan counties are responsible for intervention implementation, likely resulting in heterogeneous rollout. Reaching care continuum targets may be significantly harder in areas where transmission persists. For instance, while 15 counties saw new HIV infections reduce by 75% or more from 2013 to 2021, five counties – including four rural counties in the Western region and Nairobi – all had an increase in new infections [78].

Our findings suggest that Kenya will face an aging HIV population over the next 15 years, owing to demographic shifts and declining HIV infections. Successful HIV programming in the coming years may further accelerate these trends. Although HIV policy in Kenya has historically focused on preventing new infections and providing HIV treatment to those infected, new policy must also consider the needs of an aging HIV population as a whole – including healthy aging and the treatment of age-associated comorbidities.

Acknowledgements

Authors' contributions: conception and design: M.S., P.K., D.D., A.F. Model development: M.S., P.K., A.F. Analysis: M.S., P.K., A.F. Interpretation and important intellectual input: all authors. First draft of manuscript: M.S., P.K., A.F. Manuscript review and revision: all authors.

This research was funded by the National Institutes of Health, including grants T32AI102623 (M.C.S.); T32HL007024 (M.C.S.); K01AI138853 (P.K.); K08MH118094 (A.T.F.); R01MD018539 (A.T.F.).

Conflicts of interest

There are no conflicts of interest.

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