

# Child mortality associated with maternal HIV status: a retrospective analysis in Rwanda, 2005-2015

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## ABSTRACT

**Introduction** Child mortality remains highest in regions of the world most affected by HIV/AIDS. The aim of this study was to assess child mortality rates in relation to maternal HIV status from 2005 to 2015, the period of rapid HIV treatment scale-up in Rwanda.

**Methods** We used data from the 2005, 2010 and 2015 Rwanda Demographic Health Surveys to derive under-2 mortality rates by survey year and mother's HIV status and to build a multivariable logistic regression model to establish the association of independent predictors of under-2 mortality stratified by mother's HIV status.

**Results** In total, 12 010 live births were reported by mothers in the study period. Our findings show a higher mortality among children born to mothers with HIV compared with HIV negative mothers in 2005 (216.9 vs 100.7 per 1000 live births) and a significant reduction in mortality for both groups in 2015 (72.0 and 42.4 per 1000 live births, respectively). In the pooled reduced multivariable model, the odds of child mortality was higher among children born to mothers with HIV, (adjusted OR, AOR 2.09; 95% CI 1.57 to 2.78). The odds of child mortality were reduced in 2010 (AOR 0.69; 95% CI 0.59 to 0.81) and 2015 (AOR 0.35; 95% CI 0.28 to 0.44) compared with 2005. Other independent predictors of under-2 mortality included living in smaller families of 1–2 members (AOR 5.25; 95% CI 3.59 to 7.68), being twin (AOR 4.93; 95% CI 3.51 to 6.92) and being offspring from mothers not using contraceptives at the time of the survey (AOR 1.6; 95% CI 1.38 to 1.99). Higher education of mothers (completed primary school: (AOR 0.74; 95% CI 0.64 to 0.87) and secondary or higher education: (AOR 0.53; 95% CI 0.38 to 0.74)) was also associated with reduced child mortality.

**Conclusions** This study shows an important decline in under-2 child mortality among children born to both mothers with and without HIV in Rwanda over a 10-year span.

## INTRODUCTION

Of the approximately 37.9 million individuals living with HIV in 2018, the majority resided in sub-Saharan Africa including approximately 1.7 million children under 15 years of

## Key questions

### What is already known?

- ▶ Previously, children's mortality was assessed regardless of the mothers' HIV status.
- ▶ The survival of children born to mothers with HIV was typically worse than those children born to mothers without HIV
- ▶ The antiretroviral treatment scale-up among pregnant women has considerably reduced the vertical transmission of HIV.

### What are the new findings?

- ▶ This study determines the impact of treatment scale-up and the option B+ programme on child mortality, which has not yet been evaluated in Rwanda.
- ▶ This manuscript investigates the association between maternal HIV status and childhood mortality over time as HIV treatment scale-up was occurring in the country.
- ▶ Using Demographic Health Surveys from three time points ranging from just before rapid antiretroviral therapy scale-up to after implementation of 'option B+', this analysis compares the decline in under-2 mortality among children born of both mothers with and without HIV.

### What do the new findings imply?

- ▶ Further biomedical interventions are needed to achieve the Sustainable Development Goals.
- ▶ More studies are recommended to monitor the encouraging decline in child mortality and test emerging interventions.

age.<sup>1,2</sup> HIV/AIDS is a major cause of mortality among children under 5 years of age.<sup>3</sup> Globally, remarkable gains in reducing child mortality in the previous decades have decreased mortality from 93 to 39 deaths per 1000 live births between 2000 and 2018.<sup>4</sup> However, important regional differences contributed to these gains. Sub-Saharan Africa (57% decline), for instance, had a slower decline compared with Northern Africa and Western

Asia (65% reduction), Central and Southern Asia (68% reduction), Eastern and South-Eastern Asia (74% reduction), Latin America and the Caribbean (70% reduction) and Europe and Northern America (60% reduction).<sup>4</sup> As of 2018, sub-Saharan Africa continues to rank as the region of the world with the highest child mortality of 78 deaths per 1000 live births.<sup>5</sup> While the United Nation's Sustainable Development Goals target to reduce under-5 mortality to less than 25 per 1000 live births by 2030 in all countries, a global review of national mortality trends showed that efficient control of HIV/AIDS would decrease the under-five mortality rate by 14% per decade from 2010 to 2030.<sup>6</sup>

A number of factors have been identified as associated with higher child mortality in sub-Saharan Africa. These include low coverage of vaccination, poor hygiene and sanitation, inadequate nutrition and maternal socioeconomic and HIV status.<sup>5 7–10</sup> Many countries in the region, including Eritrea, Ethiopia, Liberia, Malawi, Mozambique, Niger, Rwanda, Uganda and Tanzania observed a reduction in the under-five mortality rate between 1990 and 2018 by up to two-thirds. Substantial gains in reducing child mortality have been made in Rwanda since the scale-up of HIV and other primary health services over the last decade, resulting in the country being one of the few to meet Millennium Development Goal (MDG)–4.<sup>5</sup>

In Rwanda, many efforts have been made over the past two decades to reduce mother-to-child HIV transmission as part of the country's efforts to reduce child mortality.<sup>11</sup> In 2012, all pregnant women with HIV were enrolled into the 'option B+ programme' that provides lifelong antiretroviral therapy (ART), irrespective of a mother's CD4-cell count,<sup>12</sup> as well as a close follow-up of mothers and their children until 24 months postdelivery. By June 2019, 98% of health facilities countrywide provided Prevention of Mother to Child Transmission (PMTCT) services. This resulted in 99% of pregnant women with HIV attending at least one antenatal care visit, 86% being accompanied by their partners for HIV testing and 91% delivering in a health facility with the assistance of a skilled healthcare provider.<sup>13</sup> The survival of children of mothers with HIV has been investigated in Rwanda using cohort studies for shorter durations of follow-up,<sup>11 14 15</sup> yet these analyses were not based on data available from nationally representative population surveys. Furthermore, the impact of nationwide HIV treatment scale-up on child mortality has not yet been evaluated in Rwanda. In this study, we assess child mortality among children less than 2 years of age born to mothers with and without HIV using population-based survey data (Rwanda Demographic and Health Survey, RDHS) from 2005 to 2015, corresponding to the timeframe of massive HIV treatment scale-up in Rwanda.

## METHODS

### Study design

The RDHS is a standardised cross-sectional household survey conducted every 5 years. RDHS uses a two-stage

cluster sampling procedure that aims to guarantee a representative sample of the target population. The sampling frame was generated first by a random sample of villages, followed by a second stage of random sampling of households within each of these villages. Sampling of each year's RDHS is done independently from each other and there is little chance that a household is part of two or more RDHS, given the low selection probability of a cluster. The survey collects data about adult males 15–54 and females 15–49 years of age.<sup>16</sup>

### Public involvement

The RDHS is implemented by the National Institute of Statistics of Rwanda and the Ministry of Health of Rwanda under the technical assistance of Inner City Fund (ICF) International through the DHS Programme. Together with other stakeholders, the Ministry of Local Government participates in the design, implementation and dissemination of the study. Specifically, local authorities play a key role during the data collection to ensure the smooth implementation in their respective areas.

### Data

We used data from RDHS from 2005, 2010 and 2014/2015 on children born in the 5 years prior respective surveys administration from all mothers tested for HIV. During each survey, every woman sampled was interviewed about her maternal history and the vital status of her children at the time of the survey. RDHS obtained informed consent for interviews from all participants and for blood draws for HIV testing of all mothers. RDHS sampled 50% of interviewed mothers for HIV testing. The response rate was 98.1% in 2005, 99.1% in 2010 and 99.5% in 2015 and the HIV test consent rate was 97.0% in 2005, 98.9% in 2010 and 99.5% in 2015. We included all children from sampled mothers tested for HIV under the age of 2 years born 24–84 months preceding the survey, excluding the month of the interview. Therefore, a cohort of children who were deemed to reach the age of 2 years in the 5 years prior to the survey was considered for analysis of under-2 mortality, defined as death between birth and 23 months of age. Deaths were solely based on the report of the mothers.

In total 92 769 children were born to sampled mothers (30 072 in 2005, 32 639 in 2010 and 30 058 in 2015, respectively) and 23 587 were born within an interval of 24 and 84 months prior to each survey (RDHS 2005, 2010 and 2014/2015). Of them, 11 816 children were born to mothers, who were offered an HIV test. Our study population was constituted with 3627 mothers from the RDHS 2005, 4381 mothers from the RDHS 2010, and 3808 mothers from the RDHS 2015 (figure 1).

### Under-2 mortality predictors

Following a literature review and the development of a conceptual framework we identified the following available variables as potential predictors of under-2 mortality: the mother's HIV and marital status at the time of the survey, the mother's education level (no education, primary,



**Figure 1** Recruitment flow.

secondary/higher) and age at childbirth (<25, 25–34, 35+ years old), residence type (urban/rural), mother’s occupation (not working; skilled work (professional, clerical, sales, services, skilled manual)); unskilled work: (agriculture, domestic, unskilled manual), size of the family (1–2, 3–4, 5+ members), use of modern contraceptives (contraceptive pill, intrauterine device, injectable, male condom and implants, female sterilisation) at the time of the survey, preceding birth spacing (first born, less than 24 months, 25–36 months, 37–48 months and 49 plus months), twin birth (yes/no), source of drinking water (categorised as ‘improved’ if the source is piped water into dwelling/yard/plot, public tap/standpipe, borehole, protected dug well, protected spring, rainwater, bottled water) as a socioeconomic proxy, and the availability of treated bed nets in the household (yes/no).

### Statistical analysis

Mortality rates were calculated for each RDHS period using a self-modified version of the ‘chmort’ function from the R statistical package ‘DHS.rates’, used to calculate under five mortality, to allow calculation of under-2 child mortality.<sup>17</sup>

Multivariable logistic regression models were constructed for each RDHS period with the aforementioned variables identified as potential predictors of under-2 mortality. Variables that were significantly associated ( $p < 0.05$ ) were considered as important predictors and were carried forward to develop a final reduced multivariable logistic model on the pooled data over the three RDHS periods in order to quantify the effect of the calendar period on the risk of under-2 childhood mortality. Logistic regression models were constructed with Stata V.15 software<sup>18</sup> using ‘svy’ commands accounting for sample weights.

RDHS data were used for this study.

### RESULTS

In total, the sample consisted of 12 010 mothers tested for HIV during the three RDHS in 2005, 2010 and 2015. Of these, 450 (3.7%) were HIV positive. Most of the tested mothers were in marital unions at the time of the surveys (85.3%), had primary or higher education (78.5%), lived in a rural area (86.2%), lived in households with an improved source of drinking water (73.8%) and did not use contraceptives at the time of the surveys (63.5%). Roughly, half of them gave birth between ages 25 and 34 years (48.8%), lived in families of more than five members (47.7%) and were poor (43.4%). With the exception of primary or higher education which increased from 70.3% in 2005 to 82.8% in 2015 and availability of bed nets at household, which increased from 17.4% in 2005 to 72.3% in 2015 the characteristics of mothers who tested positive for HIV were stable across the RDHS periods (table 1).

Under-2 mortality per 1000 live births in 2005 was higher among children born to mothers with HIV compared with HIV negative mothers (216.9; 95% CI 144.5 to 289.3 versus 100.7; 95% CI 88.8 to 112.7). In 2010, the respective mortality rates were 131.4 (95% CI 79.8 to 182.9) among children born to mothers with HIV versus 66.7 (95% CI 58.7 to 74.7) per 1000 live births among children born to HIV negative mothers. In 2015, mortality rates were 72.0 (95% CI 31.2 to 111.9) versus 42.4 (95% CI 35.5 to 49.4) per 1000 live births born to mothers with HIV and HIV negative mothers, respectively (table 2).

Results from the logistic regression models are presented in table 3. The odds of dying for children born to mothers with HIV increased from 1.93; 95% CI 1.16 to 3.22 in 2005 to 2.66; 95% CI 1.72 to 4.14 in 2010. The multivariable model shows that children born from mothers with HIV were at higher risk of dying before age 2 years compared with children born from HIV negative mothers (adjusted OR (AOR) 2.09; 95% CI 1.57 to 2.78). The risk of dying before age 2 was reduced by 31% in 2010 compared with 2005 (AOR 0.69; 95% CI 0.59 to 0.81) and by 65% in 2015 compared with 2005 (AOR 0.35; 95% CI 0.28 to 0.44). Other independent predictors of under-2 mortality included living in smaller families of 1–2 members (AOR 5.25; 95% CI 3.59 to 7.68), being a twin (AOR 4.93; 95% CI 3.51 to 6.92), living in a rural area (AOR 1.47; 95% CI 1.18 to 1.84) and being an offspring from mothers not using contraceptives at the time of the survey (AOR 1.66; 95% CI 1.38 to 1.99). Being a child from mothers with completed primary school was preventative (AOR 0.74; 95% CI 0.64 to 0.87) and secondary or higher education (AOR 0.54; 95% CI 0.38 to 0.75) as well as increased preceding birth spacing, compared with children who were born less than 24 months after their siblings, of 25–36 months (AOR 0.55; 95% CI 0.45 to 0.66), 37–48 months (AOR 0.50; 95% CI 0.39 to 0.64), above 49 months (AOR

**Table 1** Characteristics of mothers tested for HIV in 2005, 2010 and 2015

	2005		2010		2015		Total	
	n (weighted)	Per cent						
Overall	3659		4443		3908		12010	
<b>Mother's HIV test results</b>								
HIV negative	3517	96.1	4283	96.4	3759	96.2	11560	96.3
HIV positive	141	3.9	160	3.6	148	3.8	450	3.7
<b>Current marital status</b>								
Not in union	516	14.1	592	13.3	654	16.7	1762	14.7
In union	3143	85.9	3851	86.7	3254	83.3	10248	85.3
<b>Highest educational level</b>								
No education	1086	29.7	938	21.1	672	17.2	2696	22.5
Primary	2279	62.3	3127	70.4	2837	72.6	8242	68.6
Secondary/higher	294	8.0	378	8.5	399	10.2	1071	8.9
<b>Mother age at birth</b>								
Below 24 years	1116	30.5	1388	31.2	1201	30.7	3705	30.8
25–34 years	1706	46.6	2144	48.3	2013	51.5	5863	48.8
35+ years	836	22.9	911	20.5	695	17.8	2442	20.3
<b>Source of drinking water</b>								
Not improved	725	19.9	1193	27.2	1200	31.2	3119	26.2
Improved	2913	80.1	3199	72.8	2652	68.8	8764	73.8
<b>No of household members</b>								
1–2 members	69	1.9	95	2.1	91	2.3	255	2.1
3–5 members	1641	44.9	2157	48.5	2231	57.1	6028	50.2
5+ members	1949	53.3	2191	49.3	1586	40.6	5726	47.7
<b>Child is twin</b>								
No	3544	96.9	4307	96.9	3796	97.1	11647	97.0
Yes	115	3.1	136	3.1	112	2.9	363	3.0
<b>Type of place of residence</b>								
Urban	509	13.9	522	11.8	625	16.0	1657	13.8
Rural	3150	86.1	3921	88.2	3283	84.0	10353	86.2
<b>Wealth index</b>								
Poor/poorer	1574	43.0	1871	42.1	1767	45.2	5211	43.4
Middle	708	19.3	912	20.5	811	20.8	2431	20.2
Rich/richer	1377	37.6	1660	37.4	1330	34.0	4367	36.4
<b>Contraceptive use</b>								
Modern	434	11.9	2124	47.8	1828	46.8	4386	36.5
Non-modern	3225	88.1	2319	52.2	2079	53.2	7623	63.5
<b>Mothers occupation</b>								
Not working	810	22.2	356	8.0	211	5.4	1376	11.5
Skilled work	227	6.2	407	9.2	534	13.7	1169	9.7
Unskilled work	2612	71.6	3674	82.8	3163	80.9	9448	80.9
<b>Available of bed net at household</b>								
No	3021	82.6	1059	23.8	1084	27.7	5163	43.0
Yes	638	17.4	3384	76.2	2824	72.3	6846	57.0
<b>Preceding birth interval</b>								
9–24 months	879	24.1	972	22.0	542	14.0	2393	20.0

Continued

Table 1 Continued

	2005		2010		2015		Total	
	n (weighted)	Per cent						
25–36 months	1133	31.0	1330	30.1	888	22.9	3351	28.0
37–48 months	553	15.1	638	14.4	548	14.1	1739	14.5
49+ months	457	12.5	501	11.3	772	19.9	1731	14.5
First born	628	17.2	983	22.2	1133	29.2	2745	23.0

0.43; 95% CI 0.33 to 0.55) and the first born (AOR 0.73; 95% CI 0.59 to 0.91) (table 3).

## DISCUSSION

In this comparative study of nationwide representative health survey data from 2005 (the first DHS to collect HIV biomarker) to 2015, we demonstrated a marked decline in under-2 child mortality among children born of both mothers with and without HIV. The decrease in the under-2 childhood mortality was more pronounced in children born from mothers with HIV, and children born to mothers with and without HIV had a similar mortality by 2015. Despite the widespread scale up of HIV services starting in 2004, data show a higher odds of dying before the age of two among children born from mothers with HIV compared to those with no HIV in 2010. This near normalisation of childhood mortality for children born to mothers with HIV compared with those without HIV seen in 2015 likely reflects the vast gains in HIV care and mother-to-child HIV transmission experienced in Rwanda over the last decade. Children born in small households, of one to two members (most probably from single mothers), as well as from mothers with lower education, in rural areas, and those born twins were predictive of a higher risk of mortality. The family size includes the nuclear family members and dependents residing in the same household. Other studies in resource-limited settings have revealed that single motherhood, unwanted pregnancy and lack of spousal support are a higher risk of child mortality. However, this risk may also be attributed to emotional violence and the lack of economical support.<sup>19 20</sup>

Several factors contributed to the decline of childhood mortality under 5 years of age in Rwanda as outlined in a recent review.<sup>21</sup> These factors include the establishment of community health workers, a premium-based

community health insurance (Mutuelle de Santé), the scale up of contraceptive use, the decentralisation and integration of health services, increased domestic funding for health and the scale up of a national HIV PMTCT programme.<sup>22 23</sup> By 2011, 22.1% of general government expenditures were allocated to the public health sector. As a result, rates of child and maternal mortality, and deaths due to tuberculosis and malaria have fallen alongside the burden of HIV, and Rwanda is now on track for each of the health-related MDGs.<sup>21</sup> Other factors contributing to these trends are the country's engagement in initiatives to obtain gender parity and equal access to education.<sup>24</sup>

In addition to interventions that reduce mortality among all children, interventions to reduce disparities among children born of HIV positive mothers, compared with negative mothers have also been scaled up in Rwanda. Most notably, in 2005, only 5.5% of persons living with HIV were taking HIV treatment compared with 81.2% in 2015.<sup>25</sup> During the same period, access to PMTCT services more than doubled; from 41.3% in 2005 to 85.1% in 2015.<sup>25</sup> Similarly, infant delivery at a health facility has steadily increased from 28% in 2005 to 91% in 2015.<sup>26</sup> The difference in childhood mortality in 2010 and 2005 concurs with findings of a study that was conducted in sub-Saharan Africa when PMTCT and ART programme was not yet scaled up.<sup>27</sup> In addition, a study conducted in South Africa has reported a decline in under-2 mortality as a result of HIV prevention and treatment,<sup>28</sup> and different studies conducted in Africa highlighted equal chances of child survival as result of maternal HIV treatment.<sup>29</sup> Higher education and access to modern contraceptive methods was also found in many other studies to be associated with reduced child mortality.<sup>10 30–34</sup>

Our study has several limitations: First, the analysis is based on cross-sectional survey data and, the HIV status of the mother and the child was not determined at the time of delivery or during the follow-up because data were collected during surveys and time of infection could not be determined. Second, the design of the study also precludes the determination of the HIV status of children, and hence whether or not the child's death was directly attributable to HIV remained undetermined. The latter may be a source of bias, however, as no available data on a child's HIV status, we had to make the best use of available data. Third, mothers with

Table 2 Under-2 mortality rates from 2005 to 2015 in Rwanda, by HIV mother status

Survey period	Under-2 mortality (95% CI) (per 1000 births)	
	HIV negative mothers	HIV positive mothers
2005	100.7 (88.8 to 112.7)	216.9 (144.5 to 289.3)
2010	66.7 (58.7 to 74.7)	131.4 (79.8 to 182.9)
2015	42.4 (35.5 to 49.4)	72.0 (31.2 to 111.9)

**Table 3** Predictors of under 2 years childhood mortality in 2005, 2010 and 2015 in Rwanda

	DHS 2005			DHS 2010			DHS 2015			Pooled database			Pooled reduced model		
	AOR	95% CI		AOR	95% CI		AOR	95% CI		AOR	95% CI		AOR	95% CI	
<b>Mother HIV status</b>															
HIV negative (ref)	1.00			1.00			1.00			1.00			1.00		
HIV positive	1.93*	(1.16 to 3.22)		2.66*	(1.72 to 4.14)		1.64*	(0.8 to 3.34)		2.16*	(1.61 to 2.89)		2.09*	(1.57 to 2.78)	
<b>DHS year</b>															
2005 (ref)															
2010							0.69*	(0.57 to 0.83)		0.69*	(0.57 to 0.83)		0.69*	(0.59 to 0.81)	
2015							0.35*	(0.27 to 0.44)		0.35*	(0.27 to 0.44)		0.35*	(0.28 to 0.44)	
<b>Mother's education level</b>															
No education (ref)	1.00			1.00			1.00			1.00			1.00		
Primary	0.73*	(0.57 to 0.93)		0.71*	(0.55 to 0.93)		0.91*	(0.6 to 1.39)		0.75*	(0.64 to 0.88)		0.74*	(0.64 to 0.87)	
Secondary/higher	0.49*	(0.26 to 0.91)		0.53*	(0.31 to 0.9)		0.77*	(0.34 to 1.72)		0.54*	(0.38 to 0.77)		0.53*	(0.38 to 0.74)	
<b>Marital status</b>															
Not in union (ref)	1.00			1.00			1.00			1.00			1.00		
In union	1.01	(0.71 to 1.44)		0.93	(0.64 to 1.35)		1.29	(0.79 to 2.11)		1.03	(0.82 to 1.29)				
<b>Mother age at child birth</b>															
Below 24 (ref)	1.00			1.00			1.00			1.00			1.00		
25–34	1.29	(0.98 to 1.71)		1.00	(0.74 to 1.35)		0.97	(0.66 to 1.42)		1.09	(0.91 to 1.3)		1.04	(0.87 to 1.26)	
35+	1.36	(0.96 to 1.93)		1.25	(0.83 to 1.88)		1.87	(0.99 to 3.54)		1.34	(1.05 to 1.72)		1.24	(0.97 to 1.60)	
<b>Mothers occupation</b>															
Not working (ref)	1.00			1.00			1.00			1.00			1.00		
Skilled work	0.79	(0.47 to 1.31)		0.89	(0.52 to 1.52)		0.59	(0.21 to 1.66)		0.77	(0.56 to 1.06)				
None skilled work	0.92	(0.72 to 1.18)		0.80	(0.52 to 1.25)		1.62	(0.69 to 3.81)		0.93	(0.75 to 1.14)				
<b>Preceding birth interval</b>															
Below 24 months (ref)	1.00			1.00			1.00			1.00			1.00		
25–36 months	0.63*	(0.48 to 0.84)		0.44*	(0.32 to 0.59)		0.63	(0.36 to 1.11)		0.55*	(0.46 to 0.66)		0.55*	(0.45 to 0.66)	
37–48 months	0.59*	(0.41 to 0.86)		0.41*	(0.27 to 0.62)		0.54	(0.28 to 1.04)		0.50*	(0.39 to 0.64)		0.50*	(0.39 to 0.64)	
49+ months	0.47*	(0.31 to 0.69)		0.30*	(0.19 to 0.49)		0.58	(0.32 to 1.04)		0.42*	(0.32 to 0.56)		0.43*	(0.33 to 0.55)	
First born	0.93	(0.65 to 1.33)		0.57*	(0.41 to 0.78)		0.87	(0.48 to 1.56)		0.73*	(0.590.91)		0.73*	(0.59 to 0.91)	
<b>Child is a twin</b>															
No (Ref)	1.00			1.00			1.00			1.00			1.00		
Yes	5.04*	(2.99 to 8.49)		4.77*	(2.78 to 8.20)		6.22*	(2.79 to 13.89)		5.04*	(3.58 to 7.09)		4.93*	(3.51 to 6.92)	

Continued

Table 3 Continued

	DHS 2005			DHS 2010			DHS 2015			Pooled database			Pooled reduced model		
	AOR	95% CI		AOR	95% CI		AOR	95% CI		AOR	95% CI		AOR	95% CI	
<b>No of family members</b>															
1–2 members	8.71*	(4.57 to 16.61)	4.41*	(2.37 to 8.21)	4.49*	(1.87 to 10.78)	5.70*	(3.81 to 8.52)	5.25*	(3.59 to 7.68)					
3–5 members	3.04*	(2.37 to 3.89)	1.78*	(1.36 to 2.33)	2.00*	(1.28 to 3.14)	2.30*	(1.94 to 2.73)	2.15*	(1.81 to 2.54)					
5+ (ref)	1.00		1.00		1.00		1.00		1.00						
<b>Modern contraceptive use at time of survey</b>															
Yes (ref)	1.00		1.00		1.00		1.00		1.00						
No	1.54*	(1.03 to 2.32)	1.37*	(1.08 to 1.75)	2.31*	(1.58 to 3.39)	1.61*	(1.34 to 1.94)	1.66*	(1.38 to 1.99)					
<b>Residence</b>															
Urban (ref)	1.00		1.00		1.00		1.00		1.00						
Rural	1.54*	(1.09 to 2.18)	1.35	(0.89 to 2.03)	0.99	(0.55 to 1.81)	1.39*	(1.09 to 1.78)	1.47*	(1.18 to 1.84)					
<b>Source of drinking water</b>															
Nor improved (ref)	1.00		1.00		1.00		1.00		1.00						
Improved	0.92	(0.70 to 1.22)	0.92	(0.72 to 1.18)	1.09	(0.75 to 1.60)	0.95	(0.80 to 1.13)							
<b>Available treated bed net at household</b>															
No (ref)	1.00		1.00		1.00		1.00		1.00						
Yes	1.05	(0.78 to 1.43)	1.06	(0.80 to 1.40)	0.91	(0.63 to 1.32)	1.01	(0.85 to 1.21)							

\*P&lt;0.05.

AOR, adjusted OR; DHS, Demographic and Health Survey.

HIV who had died between the time of the health survey but had given birth within the retrospective time window for assessing childhood mortality are not represented in this analysis and the life status of their children is unknown; however, we estimate that this might be lower given the high ART coverage and increase in life expectancy among PLHIV on ART in Rwanda.<sup>35</sup> Fourth, infant death was self-reported and not able to be verified by national mortality statistics. However, it is very unlikely that reporting of mothers might be differential to HIV status of the mother, thereby limiting the impact of any potential recall bias. Fifth, the cause of deaths for child mortality and impact PMTCT programmes cannot be directly assessed from this data. Finally, there are factors other than HIV treatment scale-up, which likely contributed to the reduction in child mortality, such as other improvements in maternal–child health interventions or general socioeconomic growth in the country.

## CONCLUSION

This study shows a large reduction of childhood mortality between 2005 and 2015 in Rwanda. By 2015 the former considerable discrepancy in mortality of children less than 2 years of age and born to mothers with HIV has disappeared and now resembles rates of progeny from HIV negative mothers. Substantial gains have been made in Rwanda to scale-up health services and, maternal and child health programmes that include family planning and vaccination, ART and to improve the health system. However, more biomedical interventions related to empowering women, improving their education and economic status, and increasing the uptake of family planning services are needed to achieve the strategic development goal of reducing under-five mortality to less than 25 per 1000 live birth. As the health system is maturing in Rwanda, more studies are recommended to monitor these encouraging decline in child mortality and test emerging interventions.

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## REFERENCES

- UNAIDS. UNAIDS. Global HIV & AIDS statistics fact sheet, 2019. Available: <https://www.unaids.org/en/resources/fact-sheet>
- Kharsany ABM, Karim QA. HIV infection and AIDS in sub-Saharan Africa: current status, challenges and opportunities. *Open AIDS J* 2016;10:34–48.
- Lane CE, Bobrow EA, Ndatimana D, *et al*. Determinants of growth in HIV-exposed and HIV-uninfected infants in the Kabeho study. *Matern Child Nutr* 2019;15:e12776.
- UNICEF: WHO: World Bank: UN DESA. *Levels & Trends in Child Mortality 2019*, 2019: 52.
- Amoroso CL, Nisingizwe MP, Rouleau D, *et al*. Next wave of interventions to reduce under-five mortality in Rwanda: a cross-sectional analysis of demographic and health survey data. *BMC Pediatr* 2018;18:27.
- Norheim OF, Jha P, Admasu K, *et al*. Avoiding 40% of the premature deaths in each country, 2010–30: review of national mortality trends to help quantify the UN sustainable development goal for health. *Lancet* 2015;385:239–52.
- Chihana ML, Price A, Floyd S, *et al*. Maternal HIV status associated with under-five mortality in rural Northern Malawi: a prospective cohort study. *J Acquir Immune Defic Syndr* 2015;68:81–90.
- Yaya S, Bishwajit G, Okonofua F, *et al*. Under five mortality patterns and associated maternal risk factors in sub-Saharan Africa: a multi-country analysis. *PLoS One* 2018;13:e0205977.

- 9 Ezeh OK, Agho KE, Dibley MJ, *et al*. Risk factors for postneonatal, infant, child and under-5 mortality in Nigeria: a pooled cross-sectional analysis. *BMJ Open* 2015;5:e006779.
- 10 Ayele DG, Zewotir TT. Comparison of under-five mortality for 2000, 2005 and 2011 surveys in Ethiopia. *BMC Public Health* 2016;16:930.
- 11 Gill MM, Hoffman HJ, Ndatimana D, *et al*. 24-Month HIV-free survival among infants born to HIV-positive women enrolled in option B+ program in Kigali, Rwanda: the Kabeho study. *Medicine* 2017;96:e9445.
- 12 Ministry of Health (Rwanda). *National guidelines for prevention and management of HIV and STIs*. Kigali, Rwanda, 2016.
- 13 Rwanda Ministry of Health. *National HIV and viral hepatitis annual report*, 2018.
- 14 Mugwaneza P, Lyambabaje A, Umubyeyi A, *et al*. Impact of maternal art on mother-to-child transmission (MTCT) of HIV at six weeks postpartum in Rwanda. *BMC Public Health* 2018;18:1248.
- 15 Peltier CA, Ndayisaba GF, Lepage P, *et al*. Breastfeeding with maternal antiretroviral therapy or formula feeding to prevent HIV postnatal mother-to-child transmission in Rwanda. *AIDS* 2009;23:2415–23.
- 16 ICF International. *Demographic and health survey sampling and household listing manual*. ICF Int, 2012: 1–98.
- 17 Elkasabi M. Calculating fertility and childhood mortality rates from survey data using the DHS.rates R package. *PLoS One* 2019;14:e0216403.
- 18 College Station TSL. *StataCorp 15*. Release: Stata Statistical Software, 2017.
- 19 Clark S, Hamplová D. Single Motherhood and Child Mortality in Sub-Saharan Africa : A Life Course Perspective 2013:1521–49.
- 20 Bawah AA, Asuming PO, Debpuur C, *et al*. Child wanted and when? fertility intentions, Wantedness, and child survival in rural Northern Ghana. *Stud Fam Plann* 2016;47:252–63.
- 21 Binagwaho A, Farmer PE, Nsanzimana S, *et al*. Rwanda 20 years on: investing in life. *Lancet* 2014;384:371–5.
- 22 Schwandt HM, Feinberg S, Akotiah A, *et al*. "Family planning in Rwanda is not seen as population control, but rather as a way to empower the people": examining Rwanda's success in family planning from the perspective of public and private stakeholders. *Contracept Reprod Med* 2018;3:18.
- 23 Abimpaye M, Kirk CM, Iyer HS, *et al*. The impact of "Option B" on HIV transmission from mother to child in Rwanda: An interrupted time series analysis. *PLoS One* 2018;13:e0192910.
- 24 USAID/Rwanda. Gender analysis for Rwanda, 2014. Available: <https://www.usaid.gov/sites/default/files/documents>
- 25 Rwanda Biomedical Centre. *National HIV and viral hepatitis annual report 2014/15*, 2015.
- 26 Ministry of Health Rwanda, National Institute of Statistics of Rwanda I macro. *Rwanda demographic health survey 2014-15*.
- 27 Mugwaneza P, Umutohi NWS, Ruton H, *et al*. Under-two child mortality according to maternal HIV status in Rwanda: assessing outcomes within the National PMTCT program. *Pan Afr Med J* 2011;9:37.
- 28 Ndirangu J, Newell M-L, Tanser F, *et al*. Decline in early life mortality in a high HIV prevalence rural area of South Africa: evidence of HIV prevention or treatment impact? *AIDS* 2010;24:593–602.
- 29 Ndirangu J, Newell M-L, Thorne C, *et al*. Treating HIV-infected mothers reduces under 5 years of age mortality rates to levels seen in children of HIV-uninfected mothers in rural South Africa. *Antivir Ther* 2011;17:81–90.
- 30 Andriano L, Monden CWS. The causal effect of maternal education on child mortality: evidence from a Quasi-Experiment in Malawi and Uganda. *Demography* 2019;56:1765–90.
- 31 Maniruzzaman M, Suri HS, Kumar N, *et al*. Risk factors of neonatal mortality and child mortality in Bangladesh. *J Glob Health* 2018;8:10417.
- 32 Maïga A, Hounton S, Amouzou A, *et al*. Trends and patterns of modern contraceptive use and relationships with high-risk births and child mortality in Burkina Faso. *Glob Health Action* 2015;8:29736.
- 33 Chola L, McGee S, Tugendhaft A, *et al*. Scaling up family planning to reduce maternal and child mortality: the potential costs and benefits of modern contraceptive use in South Africa. *PLoS One* 2015;10:e0130077.
- 34 Bado AR, Sathya Susuman A. Women's education and health inequalities in Under-Five mortality in selected sub-Saharan African countries, 1990–2015. *PLoS One* 2016;11:e0159186.
- 35 Nsanzimana S, Remera E, Kanters S, *et al*. Life expectancy among HIV-positive patients in Rwanda: a retrospective observational cohort study. *Lancet Glob Health* 2015;3:e169–77.