

# Survival and Predictors of Mortality Among HIV Positive Adult Patients on Highly Active Antiretroviral Therapy in Public Hospitals of Kambata Tambaro Zone, Southern Ethiopia: A Retrospective Cohort Study

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Wondimu Abuto<sup>1</sup>  
Admas Abera<sup>2</sup>  
Tesfaye Gobena<sup>2</sup>  
Tariku Dingeta<sup>2</sup>  
Melese Markos<sup>3</sup>

<sup>1</sup>Public Health Emergency Management, Kambata Tembaro Zonal Health Department, Durame, Southern Nations Nationalities Peoples' Region, Ethiopia; <sup>2</sup>School of Public Health, Haramaya University, Harar, Ethiopia; <sup>3</sup>Department of Public Health, Dire Dawa University, Dire Dawa, Ethiopia

**Background:** Human Immune Deficiency Virus (HIV) infection remains the leading cause of morbidity and mortality. In Ethiopia, despite test and treat all HIV positives are adopted, a significant number of people eligible for Anti-Retroviral Therapy (ART) show up with advanced disease and at lower CD4 count. There is currently paucity of studies conducted that investigate predictors of mortality among adults on ART in the study area.

**Objective:** To explore Survival and predictors of mortality among adult HIV-positive patients on ART in Kambata Tambaro Zone, Ethiopia, from August 2013 to February 2019.

**Methods:** A health facility-based retrospective cohort study was conducted among records of 467 adult HIV-positive patients on ART selected using simple random sampling. Data were collected using standardized abstraction tool. Kaplan-Meier, Log rank tests and Cox regression model was applied to estimate survival status and identify predictors of mortality, respectively.

**Results:** Of the total 467 study subjects, 59 (12.63%) of them died in the study period. The median follow-up time of the cohort was 40.1 (IQR=13.6–59.0) months. The mortality rate of the cohort was 4.1 per 100 PYO. The overall survival probability of the cohort was 84.38% (95% CI=80.08–87.82) at 66 months. Bedridden function AHR=3.0 (95% CI, 1.44–6.64), Fair-adherence AHR=3.3 (95% CI, 1.50–7.07), Poor-adherence AHR=3.8 (95% CI, 1.88–7.96), presence of OIs AHR=4.2 (95% CI, 1.98–8.50), Late diagnosis (CD4 count  $\geq$  350) AHR=3.0 (95% CI, 1.91–6.42) and Immunologic failure AHR=3.5 (95% CI, 1.41–6.29) were independent predictors of time to death in Cox-Regression.

**Conclusion:** Late Diagnosis, poor adherence, being bedridden, having OI and Immunologic failure were independently associated with time to death. Early diagnosis to start treatment and emphasizing on close follow-up care to improve treatment adherence should be given special emphasis.

**Keywords:** survival, mortality, predictors, HIV/AIDS, HAART, low-resource setting

## Introduction

Human Immune Deficiency Virus (HIV) infection remains the leading cause of morbidity and mortality, and continues to be a challenge to major global public health issues. Globally, 78 million people have been infected with this virus and 35 million people have died from AIDS-related illnesses since the start of the epidemic.<sup>1</sup>

Correspondence: Admas Abera  
Haramaya University, P.O. Box 235, Harar,  
Ethiopia  
Email admasabera10@gmail.com

In 2016, 36.9 million people were living with HIV of which 35.1 million were adults and 21.7 million People Living With HIV (PLWHIV) were accessing ART and about 1 million AIDS-related deaths have been registered.<sup>2,3</sup> AIDS-related death remained static, 1 million deaths each year since 2015.<sup>1,7</sup> Although Sub Saharan Africa (SSA) is home to only 12% of the global population, it accounts for 75% AIDS-related deaths in which 81% of deaths in SSA were recorded only in 10 countries including Ethiopia.<sup>3</sup> Moreover, higher rate of mortality as high as 26% has been recorded particularly at early year of ART initiation in SSA including Ethiopia.<sup>8,9</sup>

Antiretroviral therapy (ART) is a great public health success that led to improved survival among HIV-infected people. The reduction in AIDS-related death is attributed to global scale-up of ART.<sup>4</sup> By the end of 2016, 70% of PLWHIV were diagnosed, 77% of those who knew their HIV status received ART and 82% of those on treatment were virally suppressed.<sup>5</sup> Even though none of the countries achieve the so-called 90–90–90 targets, the lowest achievement rates were in low and middle-income countries (LMICs).<sup>6</sup>

In Ethiopia, HIV infection continues to be a major public health problem and AIDS has claimed the lives of 1.3 million since the start of the epidemic.<sup>10</sup> The prevalence of HIV among adults was 1.16%, with substantial prevalence variation between regions. An estimated number of PLWHIV in Ethiopia were 738,976 in which 426,000 were on ART.<sup>2,10–13</sup>

Ethiopia launched implementing test and treat all policy in testing and preventing target populations for strengthening prevention, monitoring and response to HIV by sustaining HIV treatment scale-up. Despite this, there are still people who present to healthcare facilities with advanced HIV disease. Nationally in 2018, 67% of PLHIV know their status, 88% of them were on ART and 86% of people on ART have viral suppression.<sup>10</sup> Insufficient detection, late diagnosis, inadequate viral suppression; and inadequate support to sustained HIV care and treatment increase AIDS-related mortality.<sup>11</sup>

Survival is reduced among PLWHIV after 2011, even though eligibility has progressively increased and patients are starting care earlier.<sup>10,20</sup> One systematic review estimates the mortality rate to be 40.8% with the highest mortality at early months of follow-up within 3 to 12 months of ART initiation.<sup>19</sup> Different predictors were

independently associated with mortality in different parts of Ethiopia.<sup>14–18</sup>

Understanding context-specific predictors of survival in PLWHIV are essential to monitor and improve program effectiveness which in turn improves the clinical outcome of patients on ART. However, little is known about the survival status and its predictors after ART initiation in the study area. Moreover, studies showed variations and inconsistencies from region to region<sup>21–24</sup> which indicates predictors of time to death have not yet been well understood in resource-limited settings like Ethiopia including Kambata Tambaro Zone. Therefore, this study was aimed to assess Survival and predictors of mortality among adult HIV-positive patients on ART in Kambata Tambaro Zone, Ethiopia.

## Methods and Materials

### Study Design and Area

Institution-based retrospective longitudinal study was used for 5 years in public hospitals of Kambata Tambaro Zone, Southern Ethiopia from March 10–31, 2019. Kambata Tambaro zone is one of the central zone of SNNPR and has 3 administrative cities and 8 woredas. There are 7 health centers and 4 public Hospitals that provide ART services. The study was conducted in Durame General Hospital, Shinshicho Primary Hospital, Doyogena Primary Hospital and Mudula Primary Hospital. There were 1174 patients ever start ART and currently 786 patients were receiving ART during the study period.

### Study Participants and Follow-Up Time

The study participants were adults aged 15 or older who started ART between August 2013 and August 2018. Additional 6 months follow-up was added to account for those patients entering the cohort late in the study period and, to account for those patients in the cohort who had to wait until their CD4 count falls below 350 which was the guideline used before 2017. Patients who started ART outside the study facilities and have incomplete baseline data (transferred in), and patients with incomplete baseline parameters (no recorded CD4 count, no recorded WHO stage and no recorded specific ART regimen type) were excluded from the study. A total of 110 records were excluded (43 of them had no recorded CD4 count and 67 were transferred in at initiation from other facilities).

The sample size was calculated based on the assumption that type I error of 5%, 80% power and considering the

proportion of death among exposed patients (using CD4 cell count <350) was 11.25% and AHR= 2.34<sup>25</sup> and the proportion of lost to follow-up was anticipated to be 15%. Based on the above assumptions the final estimated sample size of the study was 467. These participants were selected using simple random sampling from 1174 patient records obtained from the ART clinics of the health institutions.

## Measurements

The dependent variable is time from ART initiation to an event, categorized into Censored (Alive, Transferred out and LTFU) and Death. The date of censoring for alive patient was January 31, 2019; for LTFU was the last visit date and for transferred out was date of transfer. The main predictor variable taken to identify the implicit group of the study was CD4 count at ART initiation. CD4 count measures the immune status of patients, clinical, risk of OI and support HIV diagnosis decision.<sup>26</sup> CD4 count below 350 cells/mm<sup>3</sup> suggested a late diagnosis and CD4 count  $\geq$  350 cells/mm<sup>3</sup> was considered as early diagnosis.<sup>7,18,47</sup> Socio-demographic predictors (Age, sex, residence, marital status, occupational status and educational status); Baseline Clinical parameters (ART start period, baseline WHO clinical stages, functional status, TB treatment, ART regimen type, ART regimen change, CPT use, IPT use, ART Adherence status, OI other than TB, disclosure status, BMI) and Baseline Laboratory tests (viral load test, hemoglobin) were the other covariates. We have also included a time-dependent variable, immunologic failure, which is defined as patients' CD4 count declines from the baseline, or CD4 levels are persistently below 100 cells/mm<sup>3</sup> (more than two times after baseline).<sup>27</sup>

## Operational Definitions

**Good adherence:** If the percentage of missed dose is >95% (<2 doses of 30 doses or <3 doses of 60 doses) as documented by ART physicians.<sup>7</sup>

**Fair adherence:** If the percentage of missed dose is between 85–94% (3–5 doses of 30 doses or 3–9 doses of 60 doses) as documented by ART physician.<sup>7</sup>

**Poor adherence:** If the percentage of missed dose is <85% (>6 doses of 30 doses or >9 doses of 60 doses) as documented by ART physician.<sup>7</sup>

**Late diagnosis:** HIV positive patients (who present with CD4 cell count <350 cells/mm<sup>3</sup>)<sup>5,18,47</sup>

**Early diagnosis:** HIV positive patients (who present with CD4 cell count  $\geq$ 350 cells/mm<sup>3</sup>)<sup>5,18,47</sup>

**Functional Status:**

**Ambulatory:** able to perform routine activities of daily living.<sup>7</sup>

**Bedridden:** A patient unable to perform routine activities of daily living.<sup>7</sup>

**Working:** A patient able to perform his/her usual work.<sup>7</sup>

**Lost to follow up:** A patient not seen for 1–3 months after the last appointment.<sup>7</sup>

**Immunologic Failure:** patients' CD4 count declines from the baseline, or CD4 levels are persistently below 100 cells/mm<sup>3</sup> (more than two times after baseline).<sup>27</sup>

## Data Extraction and Data Quality Control

A pre-prepared data retrieval (abstraction tool) form was developed from records of the Federal Ministry of Health, ART entry and follow-up forms, patient records, laboratory requests and computer data. Death information was confirmed from reviewing the death summary, medical registration in ART clinic, or registration by ART adherence supporter through calling using the registered phone number or neighborhoods of the patients. The recent data before ART initiation was taken as baseline and data before one month of ART initiation was used for patients who started ART immediately after diagnosis. Data was extracted after giving 2 days training for data collectors and supervised by the Principal investigator. The extracted data were checked for the completeness, accuracy and clarity.

## Ethical Considerations

Ethical approval of the study was obtained from the Haramaya University, College of Health and Medical Sciences, Institutional Health Research Ethics Review Committee (IHRERC) with Reference Number IHRERC/039/2019. Letter of permission was obtained from Kambata Tambaro Zonal Health Department and agreement consent was obtained from each Public Hospital managers prior to the study. Due to difficulty to reach patients to obtain informed consent, data were extracted anonymously ensuring patient data confidentiality, and all data collection was conducted in compliance with the declaration of Helsinki.

## Data Processing and Analysis

Data were entered into Epi-Data version 4.1 and exported to STATA version 14.1 for data processing and analysis. Data were coded, labeled and categorized for

analysis. The time that each participant contributed to the study was calculated by months from date of ART start to the date of censored or death. The time that the patient discontinued ART was reduced for re-entered patients after attrition. Kaplan–Meier was used to estimate the survival probability of the cohort and category of each predictor. Log rank test was employed to compare (test) equality of surviving, ie, checking whether survival probabilities are statistically significant. Semi-parametric (Cox regression) Model was applied to identify the predictors of time-to-death and to calculate the hazard ratios (HR).

Bivariate Cox-Regression analysis was performed, forward variable selection using AIC (Akaike Information Criteria) was used with a liberal p-value of 0.1 to ensure essential predictor variables are retained. The proportionality hazard assumption test was checked using Schoenfeld residuals on functions of time.

## Results

A total of 467 study participants' records were reviewed, among them 263 (56.3%) were females and the overall median age was 30 years (IQR=26-38) and 213 (54.4%) participants live in rural areas (Table 1).

## Clinical and Laboratory Characteristics

From the total 437 records reviewed, 173 (37.0%) patients had BMI <18.5 Kg/m<sup>2</sup>, 107 (22.9%) patients were in Ambulatory functional status and 60 (13%) were Bedridden. About 171 (36.6%) had OI other than TB, 183 (39.2%) were in WHO stage III or IV and 57 (12.2%) patients had not received CPT. Further, about 189 (40%) had CD4 <350 cells/mm<sup>3</sup>, 31 (11.3%) had Viral load >1000 copies, and 116 (30.8%) had Hemoglobin level below 10 (Table 2).

## Survival Status of the Cohort

In this cohort, out of 467 study subjects on ART, 408 (87.4%) patients were censored and 59 (12.6%) patients died. From 408 censored subjects, 308 (75.5%) were Alive and 69 (16.9%) were LTFU and 31 (7.6%) were transferred out (Figure 1).

The overall follow-up of the cohort was 1412.7 person-years of observation (PYO). The median follow-up of the overall cohort was 40.1 (IQR=13.63–59.0) months. The total mortality rate of the cohort over 66 months of follow-up was 4.1 per 100 PYO. The survival probability of the cohort at 3 months, 6 months, 1 year, 2 years, 3 years, 4

**Table 1** Patients Baseline Sociodemographic Characteristics, Kambata Tambaro Zone, Ethiopia, August 2013 to February 2019

Socio-Demographic Characteristics	Frequency	Percentage
<b>Sex (467)</b>		
Female	263	56.3
Male	204	43.7
<b>Age group (in years)</b>		
15–24	58	12.4
25–34	229	49.0
35–44	121	26
45+	59	12.6
<b>Marital status (422)</b>		
Never Married	72	16.9
Married	252	59.3
Divorced	58	13.6
Widowed/Separated	43	10.1
<b>Educational status (436)</b>		
No formal education	58	13.3
Primary	231	53
Secondary	112	25.7
Tertiary	35	8
<b>Residence (467)</b>		
Urban	254	54.4
Rural	213	45.6
<b>Occupational status (429)</b>		
Government employee	52	12.4
Farmer	102	24.2
House wife	51	12.1
Merchant	94	22.3
Self-employee	94	22.3
Others	36	6.6
<b>ART start period (467)</b>		
Before 2017	334	71.5
After 2017	133	28.5

years and 5 and half years was 96.9%, 93.8%, 91.9%, 88.5%, 85.9%, 85.5% and 84.4%, respectively. The overall failure probability of the cohort was 15.6% at 66 months (Figure 2).

The survival probability descends sharply in the first 6 months, steadily descends until 36 months and descends slowly until it reaches its minimum (84.23%) at 60 months (Figure 3A). On the other hand, the probability of survival among patients diagnosed late and early was 78.2% and 93.2%, respectively (Figure 3B). The survival rate of patients diagnosed late declines steadily after ART start in the first month until 30 months reaching the minimum

**Table 2** Clinical and Laboratory Characteristics of Patients in Kambata Tambaro Zone, Ethiopia, August 2013 to February 2019

Clinical Characteristics	Frequency (n=467)	Percentage
<b>BMI (kg/m<sup>2</sup>)</b>		
Below 18.5	173	37.0
18.5 and Above	294	63.0
<b>Functional status</b>		
Working	300	64.2
Ambulatory	107	22.9
Bedridden	60	12.9
<b>WHO clinical stages</b>		
I and II	284	60.8
III and IV	183	39.2
<b>Disclosure status</b>		
Yes	294	63.0
No	173	37.0
<b>ART adherence</b>		
Good	334	71.5
Fair	74	15.9
Poor	59	12.6
<b>ART side effect</b>		
Yes	31	6.6
No	436	93.4
<b>CPT</b>		
Yes	410	87.8
No	57	12.2
<b>IPT</b>		
Yes	382	81.8
No	85	18.2
<b>TB co-infection</b>		
Yes	111	23.8
No	356	76.2
<b>CD4 count</b>		
Below 350 cells/mm <sup>3</sup>	278	59.5
350 and Above cells/mm <sup>3</sup>	189	40.5
<b>OIs other than TB</b>		
Yes	71	36.6
No	296	63.4
<b>Viral load</b>		
Below 1000	252	88.7
1000 and Above	31	11.3
<b>Immunologic failure</b>		
Yes	66	15.0
No	375	85.0
<b>Hemoglobin</b>		
Below 10	116	30.8
Above 10	261	69.2

**Abbreviations:** BMI, body mass index; ART, anti-retroviral therapy; CPT, cotrimoxazole preventive therapy; IPT, isoniazid preventive therapy; OIs, opportunistic infections; TB, tuberculosis.

of 78.2%. Fourteen (23.7%) of deaths were recorded in the first 3 months and another 14 (23.7%) deaths between 3 and 6 months of follow-up and 28 (47.4%) death happened between 6 and 36 months (Table 3).

In the current study, sex, marital status, education level, residence, occupational status, ART start year, care category, eligibility criteria, disclosure status, ART regimen type, ART side effect, ART regimen change and Viral load (<1000 Vs ≥1000) were not significantly associated with survival of patients.

In bivariate cox regression model, variables that were significantly associated with survival of participants include; Age at ART initiation, BMI, functional status, WHO clinical stages, ART adherence, tuberculosis, IPT use, CPT use, OI other than Tb, CD4 count at ART initiation (Late diagnosis, ie, CD4 Count <350 cell/μL Vs Early Diagnosis, ie, CD4 Count ≥350cell/μL), Immunologic failure and hemoglobin level (Table 4).

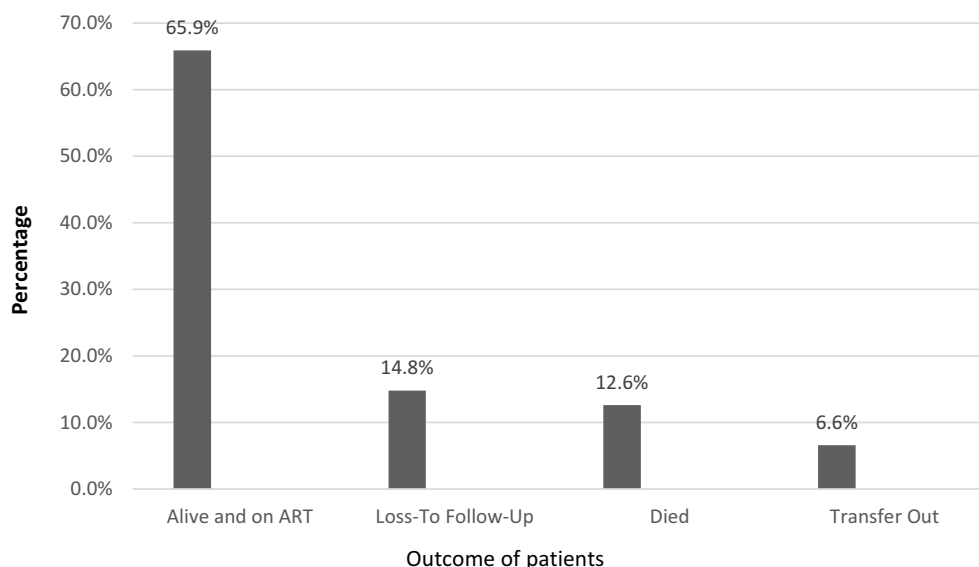
After controlling for confounding in the multivariable analysis of Cox Regression Model, variables that are independent predictors of time to death include: being bedridden patient (AHR = 3.0, 95% CI = 1.44–6.64), Fair adherence to ART (AHR=3.26, 95% CI=1.50–7.05), Poor adherence to ART (AHR=3.87, 95% CI=1.88–7.95), presence of OIs other than TB (AHR= 4.09, 95% CI= 1.98–8.50), Late diagnosis measured by CD4 count (CD4 count ≥350) (AHR=2.99, 95% CI=1.41–6.30), and Immunologic failure (AHR=3.50, 95% CI, 1.90–6.41) (Table 4).

## Discussion

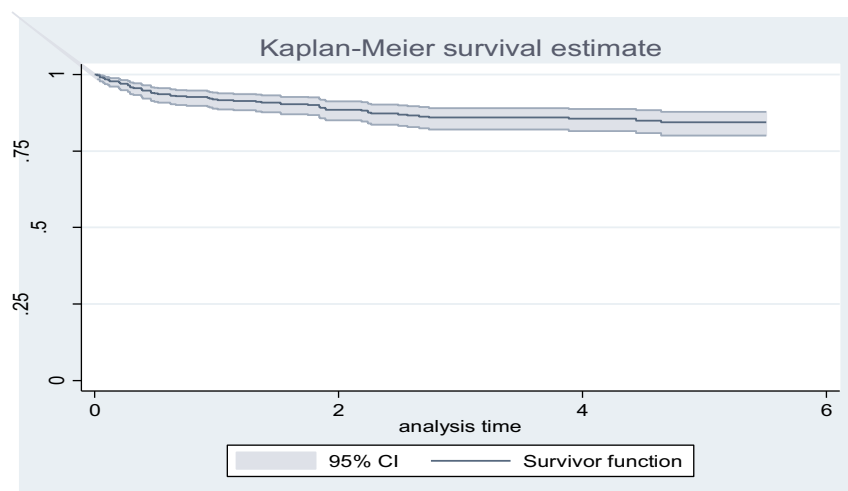
This retrospective cohort study shows early diagnosis before starting ART treatment regardless of CD4 count and WHO staging improves survival of patients on ART. Further, CD4 count <350 (Late diagnosis), poor adherence, being bedridden functional status, having OI and immunologic failure on ART follow-up patients have higher risk of mortality among adults on ART.

The probability of survival for 66 months was 84.4%, which is consistent with the study done in Nepal,<sup>32</sup> India<sup>33</sup> and elsewhere in Ethiopia.<sup>14,15,25</sup> This finding, however, is higher than the study done in Cameroon<sup>28</sup> where 1167 aged 15 years and above patients were followed from 2001 to 2006, and in Ethiopia,<sup>24,29</sup> where records of 930 adults were studied between 2005 and 2010. The discrepancy can be attributed to the difference in study time. The finding of the present study was lower than study done in Southern





**Figure 1** Survival status of HIV positive patients on ART in Kambata Tambaro Zone, Ethiopia from August 2013 to February 2019.



**Figure 2** Kaplan-Meier survival rate estimate of HIV positive patients on ART in Kambata Tambaro Zone, Ethiopia from August 2013 to February 2019.

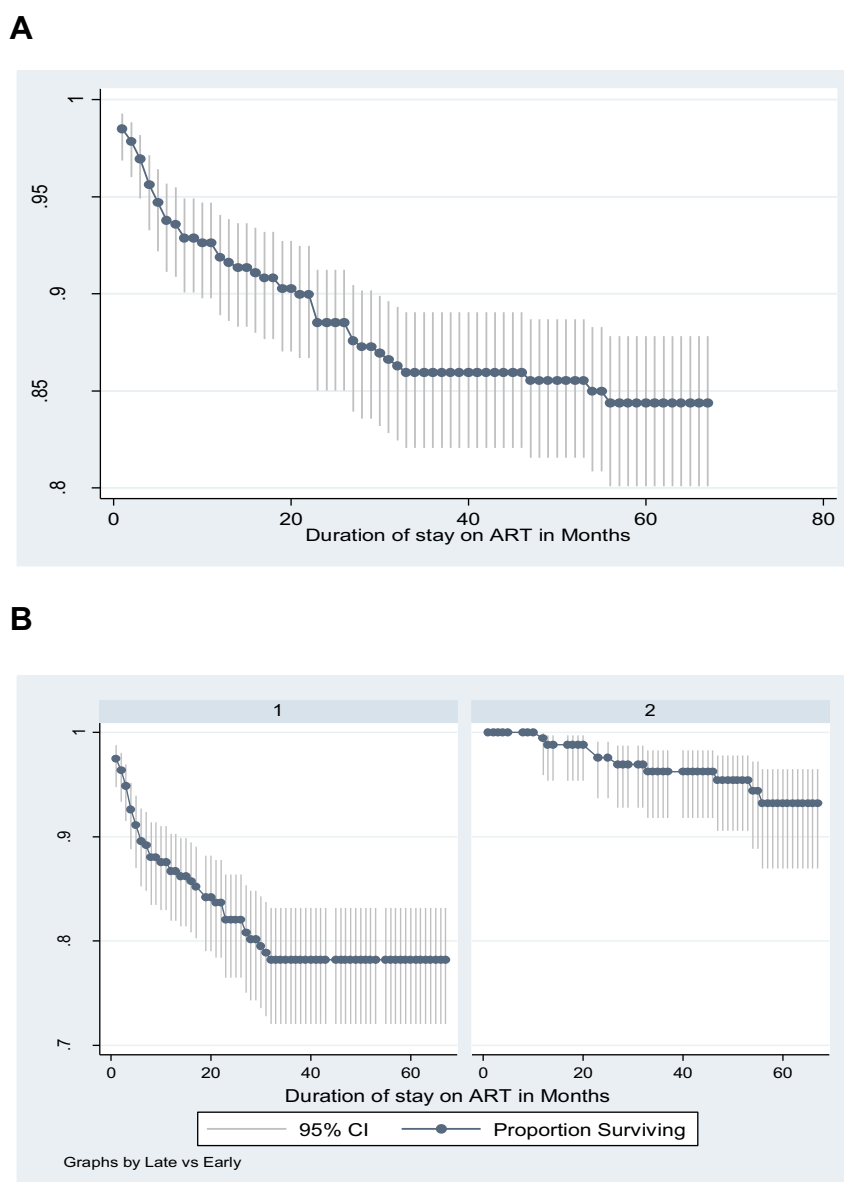
Ethiopia,<sup>30</sup> Aksum,<sup>31</sup> Nekemte<sup>17</sup> in which the studies recruited relatively higher number of participants. These differences can be attributed to the difference in study time and sampling as well as the difference in length of follow-up.

The mortality rate of the cohort in the current study was 4.1 per 100 PYO. This finding is lower than the study done in Cameroon,<sup>34</sup> India,<sup>33</sup> Nepal<sup>32</sup> and Ethiopia,<sup>30</sup> but higher than the study done in Jinka,<sup>29</sup> Debre Markose,<sup>24</sup> Harar.<sup>25</sup> This might be probably due to the difference in diagnosis and ART initiation time and the difference in proportion of advanced WHO stages in the studies.

Late HIV diagnosis was a strong predictor of mortality in this study. This was in line with the study done in

Tigray.<sup>18</sup> Other studies reported in agreement with this study, thus an increase in baseline CD4 count reduce AIDS-related mortality<sup>35</sup> and the lower CD4 count at ART initiation have higher risk of mortality, Hind,<sup>33</sup> Uganda,<sup>36</sup> Kenya,<sup>37</sup> Ethiopia.<sup>15,23,25,38–40</sup> This is because high CD4 count reduces the occurrence of opportunistic infection and reduces AIDS-related mortality.<sup>7</sup>

This study revealed that the presence of opportunistic infections is a strong predictor of time to death. Accordingly, patients with OI are four folds at increased risk of mortality compared to patients without OI. The finding is supported by various other studies elsewhere.<sup>35,41,42</sup> The occurrence of Opportunistic infection determines the level of immunity



**Figure 3** Kaplan–Meier survival function estimate of all HIV positive patients on ART (A) and patients diagnosed late (left) and early (right) diagnosed (B), in Kambata Tambaro Zone, Ethiopia, August 2013 to February 2019.

and is the predominant cause of morbidity and mortality among HIV-infected people.<sup>7,47</sup>

This study also revealed that adherence level to ART is a strong predictor of survival. Patients with poor adherence are almost four times at increased risk of mortality compared to patients with good adherence. Different studies in India<sup>43</sup> and Ethiopia<sup>16,24,41</sup> reported in agreement with the present study. This might be due to the fact that low adherence for treatment decreases the effectiveness and benefits of the ART treatment.<sup>48</sup>

Functional status was the indicator for poor performance scale for capability of self-care and able to perform routine

daily activities by themselves to improve quality of life and survival. This study revealed that bedridden functional status was a significant predictor of reducing survival of patients on ART. This is supported by a study done in India,<sup>33</sup> in Kenya<sup>37</sup> and elsewhere in Ethiopia.<sup>14,29,38,39,44</sup>

Moreover, the study revealed that immunologic failure was strong predictors of reducing survival in patients on ART which is in line with previous studies.<sup>21,45</sup> A successful ART regimen offers the best opportunity for effective immune recovery and prevent OIs. Continuous monitoring of CD4 count alerts to start and continue prophylaxis for OI to recover immunologic

**Table 3** Life Table Showing Pattern of Death with Time Intervals Among HIV Positive Patients on ART in Kambata Tambaro Zone, Ethiopia, August 2013 to February 2019

Time Interval in Months	Failure/Death	Survival Probability	95% CI
0 to 3 months	14	0.9695	(0.9490, 0.9818)
3 to 6 months	14	0.9379	(0.9113, 0.9567)
6 to 12 months	8	0.9185	(0.8888, 0.9406)
12 to 24 months	12	0.8852	(0.8502, 0.9124)
24 to 36 months	8	0.8592	(0.8202, 0.8902)
36 to 48 months	1	0.8551	(0.8154, 0.8869)
48 to 66 months	2	0.8423	(0.7984, 0.8773)

**Table 4** Predictors of Survival Rate in Cox Regression Model Analysis Among Adult HIV Positive Patients, Kambata Tambaro Zone, from August 2013 to February 2019

Variables	Crude HR (95%, CI)	P value	Adjusted HR (95%, CI)	P value
<b>Age</b>				
15–24	1			
25–34	1.71 (0.60–4.88)	0.318	1.32 (0.38–4.6)	0.481
35–44	1.66 (0.54–5.04)	0.371	1.06 (0.29–2.96)	0.677
45+	3.31 (1.09–10.07)	0.035	2.82 (0.75–10.70)	0.097
<b>BMI (kg/m<sup>2</sup>)</b>				
Below 18.5	3.01 (1.78–5.08)	<0.001	1.38 (0.63–3.00)	0.344
18.5 and Above	1			
<b>Functional status</b>				
Working	1			
Ambulatory	1.92 (31.02–3.708)	0.043	1.44 (0.72–2.85)	0.299
Bedridden	5.65 (3.11–10.25)	<0.001	<b>3.09 (1.44–6.64) *</b>	0.004
<b>WHO clinical stages</b>				
I and II	1			
III and IV	1.70 (1.02–2.84)	0.041	1.50 (0.67–3.30)	0.210
<b>ART adherence</b>				
Good	1			
Fair	2.65 (1.35–5.20)	<0.001	<b>3.26 (1.50–7.05)*</b>	0.003
Poor	6.37 (3.57–11.38)	<0.001	<b>3.87 (1.88–7.95)*</b>	<0.001
<b>CPT</b>				
Yes	1			
No	2.22 (1.15–4.29)	0.017	2.31 (0.67–8.01)	0.246
<b>IPT</b>				
Yes	1			
No	3.38 (1.96–5.80)	<0.001	1.47 (0.60–3.61)	0.296
<b>TB co-infection</b>				
No	1			
Yes	1.77 (1.04–3.01)	0.036	1.41 (0.66–3.01)	0.377

(Continued)



Table 4 (Continued).

Variables	Crude HR (95%, CI)	P value	Adjusted HR (95%, CI)	P value
<b>CD4 count</b>				
Below 350	6.14 (3.10–12.16)	<0.001	<b>2.99 (1.41–6.30)*</b>	0.004
350 and Above	I			
<b>OIs other than TB</b>				
Yes	4.86 (2.70–8.73)	<0.001	<b>4.09 (1.97–8.47)*</b>	<0.001
No	I			
<b>Immunologic fail</b>				
Yes	7.32 (4.09–13.10)	<0.001	<b>3.50 (1.90–6.41)*</b>	<0.001
No	I			
<b>Hemoglobin</b>				
Below 10	2.80 (1.51–5.20)	0.001	1.90 (0.92–3.90)	0.077
Above 10	I			

**Note:** \*Statistically significant association. The bold texts represent statistically significant values.

**Abbreviations:** HR, hazard ratio; CI, confidence interval; BMI, body mass index; WHO, World Health Organization; Art, anti-retroviral therapy; Cpt, cotrimoxazole preventive therapy; IPT, isoniazid preventive therapy; Ois, opportunistic infections; Tb, tuberculosis.

failure. Immune suppression may persist in patients who do not experience a significant increase in CD4 count, failure to achieve CD4 recovery and presence of CD4 decline that reduce length of survival.

WHO stage was not a predictor of time to death in the current study, while the WHO Stage was a significant predictor of mortality elsewhere.<sup>24,25</sup> This is indeed a surprising finding that the advanced stage of HIV has no association with mortality. This might be due to compared to the proportion of people in WHO stage I and II, the low proportion of patients in stages III and IV in our study sample. Further, the presence of TB was not independently associated with mortality in this study whereas other studies indicate that TB was a significant predictor of mortality.<sup>25,46</sup> This can be attributed to the early initiation of ART after Tuberculosis infection with high CD4 count does not increase the risk of death.

## Limitations of the Study

Selection bias might have been introduced since records with incomplete information or charts that were lost for some patients were excluded. All-cause death might overestimate the mortality since the cause of death in this study was not identified. On the other hand, participants who were lost to follow up and those who do not have phone numbers on their folder were not traced back which might underestimate the mortality rate of the cohort.

## Conclusion

Our study found that the proportion of death late diagnosis independently predicts time to death among people with HIV/AIDS on ART. The present study also revealed that bedridden patients, poor adherence to ART, presence of OIs and immunologic failure were positively associated with increased risk of death. The findings indicate that local health bureaus in collaboration with NGOs and other stakeholders need to continually work to raise awareness that HIV/AIDS is still a major public health issue so as to diagnose patients early and start them on ART. In longitudinal HIV care, patients with OI and patients with poor adherence to ART ought to be highlighted for close follow-up care. Last but not least, health institutions need to strengthen their data recording practices including baseline information (phone number, address of patients and family) which are essential in tracking of LTFU patients.

## Data Sharing Statement

All data are available upon request of the authors of the study.

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## Author Contributions

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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The authors declare that they have no conflicts of interests for this work.

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