



OPEN HIV-1 prevalence, drug resistance, and associated factors in the urban Ethiopian population

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Despite progress in HIV control, gaps persist in understanding the prevalence, drug resistance, and associated factors in urban Ethiopia. This cross-sectional study analyzed data from 19,136 consenting participants (aged 15–64 years) for interview and blood testing in the 2017–2018 Population-based HIV Impact Assessment (PHIA) survey to assess HIV-1 prevalence, drug resistance mutations, and associated factors. Participants confirmed to be living with HIV (PLHIV) provided additional samples for CD4 + T-cell counts, viral load testing, and drug resistance analysis. Forty-three samples were analyzed for viral subtyping and drug resistance profiling, of which 42 samples successfully amplified. Data were analyzed using descriptive statistics and logistic regression. The overall HIV-1 prevalence was 3%. Among genotyped samples, 79.1% harbored drug resistance mutations, with high rates of resistance to Non-Nucleoside Reverse Transcriptase Inhibitors (NNRTIs) (78.8%) and Nucleoside Reverse Transcriptase Inhibitors (NRTIs) (66.7%). Regarding associated factors, females had 1.7 times higher odds of HIV infection than males (AOR = 1.7). Participants aged 35–44 and 45–54 had significantly higher odds of infection (AOR = 6.7 for both) compared to those aged 15–24. Lower educational attainment (primary school: AOR = 2.5; secondary school: AOR = 2.2) was associated with increased HIV prevalence compared to post-secondary education. Previously married individuals had higher infection odds than never-married participants (AOR = 2.7). Geographically, residents of Gambella showed significantly higher odds (AOR = 2.8) compared to those in Tigray. This study identified a high HIV-1 prevalence in Ethiopia, along with elevated virological failure rates, a significant proportion of PLHIV unaware of their status, and widespread drug resistance. Sex, age, education level, marital status, and region were the identified factors associated with HIV-1 prevalence. To improve treatment outcomes and curb transmission, Ethiopia should strengthen HIV prevention strategies focused on vulnerable populations, enhance testing and linkage to care, and implement routine drug resistance monitoring.

Keywords HIV-1 prevalence, Drug resistance, Associated factors, Urban Ethiopia, EPHIA

Background

The Human Immunodeficiency Virus (HIV) infections remain a major public health issue in Ethiopia, particularly in urban areas^{1,2}. It can be classified as either recently acquired (within 6–12 months), characterized by high viral loads (≥ 1000 copies/mL) and distinct antibody maturation markers, or long-term infection (persisting beyond this period)^{3,4}. Across Africa, HIV prevalence varies widely, ranging from 0.54% in Niger to 23.98% in Lesotho⁵. In Ethiopia, HIV-1 is the predominant strain, accounting for over 95% of HIV cases⁶, with its prevalence showing significant regional variation, reaching 3.4% in Addis Ababa city and 4.8% in the Gambella region⁷.

The emergence of drug resistance mutations (DRMs) is a growing concern, significantly compromising the efficacy of antiretroviral therapy (ART)^{8–10}. Mutations such as M184V and K103N^{11,12} are increasingly reported, with cross-class resistance¹³. Previous studies highlighted a high resistance to protease inhibitors,

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including atazanavir/ritonavir (28.6%) and lopinavir/ritonavir (14.3%)¹⁴, and 9.8% for Non-Nucleoside Reverse Transcriptase Inhibitor (NNRTI) mutations in the ART-naïve population¹⁵. For patients on ART, the existence of dual resistance to Nucleoside/Nucleotide Reverse Transcriptase Inhibitors (NRTIs) and NNRTIs is alarmingly prevalent (65.3%)¹⁶, with mutations including K65R and M184V having been frequently observed in first-line ART failures^{17,18}.

As previous studies showed, HIV transmission in Ethiopia is driven by a combination of behavioral, socioeconomic, and structural factors^{19,20}. The key behavioral factors include inconsistent condom use^{1,21–23}, multiple sexual partners^{1,22–25}, transactional sex²³, early sexual debut²⁴, and substance use^{21,25,26}. Socioeconomic disparities further exacerbate the epidemic, including poverty^{1,22,27–29}, low education levels^{1,22,30}, limited healthcare access^{27,30}, urban residence^{28,29,31}, and the lack of media exposure²⁸.

While previous studies have provided valuable insights, their small-scale designs limit generalizability to Ethiopia's urban population. By analyzing the PHIA dataset, this study generated evidence on HIV-1 prevalence, drug resistance, and associated factors in urban HIV dynamics that can inform targeted interventions, optimize ART regimens, and guide national HIV policies to curb the epidemic effectively.

Methods

Survey setting and population characteristics

The survey was conducted in Ethiopia, a country divided into nine regional states and two city administrations. The study targeted urban areas across Ethiopia, focusing on participants aged 15 to 64 years. Data collection took place from October 2017 to April 2018 as part of the EPHA survey. The survey was designed to generate comprehensive national and subnational estimates of HIV-1 prevalence, viral load suppression, and other key indicators among urban residents.

Sampling strategy and study population

The survey utilized a multi-stage stratified sampling design. In the first stage, 393 enumeration areas (EAs) were selected. From each EA, an average of 30 households were randomly chosen, resulting in a total of 11,810 households. Overall, 19,136 study participants aged 15 to 64 consented to participate in the interview and HIV-1 testing (Fig. 1).

Data collection procedures

The data collection process was conducted through face-to-face interviews using tablet-based data capture systems. Trained interviewers gathered information at the household level across all urban administrative regions of the country. The survey collected data on demographic characteristics, socioeconomic factors, HIV testing history, knowledge about HIV/AIDS, sexual behavior and practices, HIV prevention methods, access to healthcare services, ART usage among people living with HIV (PLWH), and blood samples for biomarker analysis.

HIV screening, viral subtyping analysis, and drug resistance

Blood samples from consenting participants underwent on-site HIV-1 screening using the national testing algorithm (Wantai HIV 1/2 test, Uni-Gold HIV 1/2™ confirmatory test, and Vikia HIV 1/2 tie-breaker test). Additional samples from PLWH were collected for CD4 + T-cell counts, viral load testing, drug resistance analysis, and recent infection assessment. CD4 + T-cell counts were measured in the field for all PLWH participants and a 2% random sample of HIV-negative individuals using a Pima™ Analyzer and Cartridge by trained staff. Viral load testing was performed on plasma (Roche COBAS®) and dried blood spots (DBS; Abbott m2000), both employing automated RNA extraction, purification, amplification, and detection. Roche's 1 mL plasma protocol and Abbott's open-mode DBS protocol (for low-volume samples) were applied³². ART coverage was evaluated by analyzing samples from all PLWH participants for ARVs (efavirenz, lopinavir, and tenofovir) using LC-MS/MS. These drugs were chosen based on their common use in treatment regimens and extended half-lives, though broader ARV profiling was limited by cost³². Drug resistance mutations in the protease and reverse transcriptase genes were analyzed using Stanford University's Genotypic Resistance Interpretation Algorithm or Calibrated Population Resistance Tool³³. Transmitted resistance was assessed by comparing recently infected PLWH with long-term infection cases, with mutations detected in the pol gene via TaqMan® SNP Genotyping Assay (Applied Biosystems)³². Viral RNA extraction was performed using NucliSens easyMAG (bioMérieux), followed by sequencing of ~1.1-kilobase amplicons on an Applied Biosystems Genetic Analyzer³³.

The study analyzed 43 samples (all 5 confirmed recent infections and 38 from long-term infections) for drug resistance and viral subtyping, with successful amplification achieved in 42 samples³². The HIV pol gene was amplified by a one-step reverse transcription polymerase chain reaction (RT-PCR), followed by nested PCR. Each sample was subtyped using the REGA HIV-1 & 2 Automated Subtyping Tool^{34,35}.

Recent and long-term infections classification

Recent and long-term HIV infections were classified using two algorithms combining HIV-1 LAg avidity, viral load (VL), and antiretroviral (ARV) detection. Recent infections were defined by a median normalized optical density (ODn) ≤ 1.5 on the LAg avidity test (confirmed through triplicate testing) coupled with VL ≥ 1,000 copies/mL; cases meeting these criteria but with no detected ARVs were also classified as recent, reflecting active viral replication in untreated individuals. Long-term infections were identified through an initial LAg avidity ODn > 2.0; a median ODn > 1.5 upon repeat testing if the initial result was ≤ 2.0; a median ODn ≤ 1.5 with VL < 1,000 copies/mL (indicating viral suppression); or cases with median ODn ≤ 1.5 and VL ≥ 1,000 copies/mL but with detected ARVs (suggesting treatment failure or non-adherence rather than recent infection). This

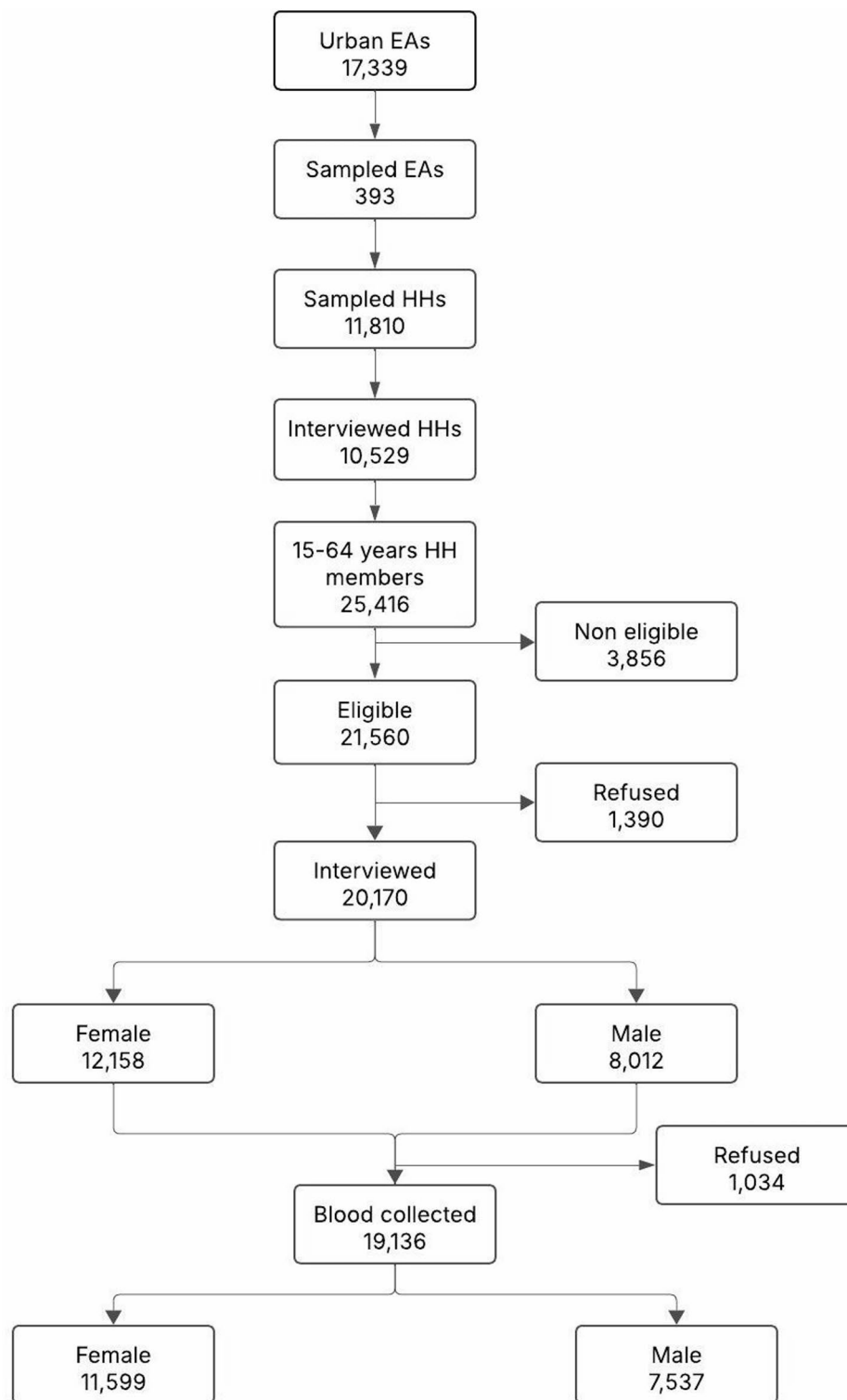


Fig. 1. Flow diagram showing participant selection in the PHIA survey.

multi-parameter approach enhances diagnostic accuracy by distinguishing true recent infections from long-standing ones with similar biomarker profiles due to treatment effects or natural viral control.

Variable selection and category

The primary outcome variables in this study were HIV-1 prevalence and the presence of drug-resistant mutations. Independent variables, selected based on a review of existing literature, included demographic factors (age, sex, education level, marital status, region, and household wealth index), behavioral factors (alcohol

and non-prescribed drug use, sexual history including age at first sex, condom use, number of sex partners in the last 12 months, relationship with the most recent partner, and condom use consistency in non-marital or non-cohabiting relationships; HIV testing history such as ever being tested and self-reported HIV status; and STI history), and laboratory variables (HIV-1 infection duration, viral load suppression, CD4 count, ART status [self-reported and ARV testing], viral load, ARV detection, duration of ART, time since ART initiation, and drug-resistant mutations). These variables were used to evaluate the health status of PLWH and/or ART effectiveness.

Operational definitions

Virological failure A confirmed HIV viral load of $\geq 1,000$ copies/mL in PLWH who are on ART³².

Viral load suppression An HIV viral load of $< 1,000$ copies/mL in PLWH who are on ART, indicating effective ART response³².

Early sexual initiation (sexual debut) The initiation of sexual activity before the age of 18 years. This threshold was selected based on established literature linking early sexual initiation to increased HIV risk and its relevance to adolescent health policies in Ethiopia^{36–38}.

Data management and analysis

All potential variables were selected based on existing literature. Data were recoded and analyzed using STATA version 17.0 (StataCorp, College Station, TX, USA). A complete case analysis approach was employed, excluding participants with missing values for any variables in the multivariable logistic regression model to ensure the reliability of estimates and the validity of statistical inferences. The statistical analysis also accounted for the complex survey design by incorporating weighting adjustments to address the sampling strategy and non-response. Descriptive statistics were used to present means, frequencies, and proportions. The bivariable logistic regression analyses were performed between each independent variable and the primary outcome (HIV-1 prevalence) to identify potential candidate variables for the multivariable model. Variables showing a suggestive association with HIV-1 prevalence ($p < 0.25$) in the bivariable analyses were considered for inclusion in the multivariable analysis to avoid prematurely excluding potentially important variables. In the final multivariable model, variables with $p < 0.05$ were considered statistically significant independent predictors of HIV-1 prevalence.

Ethics approval

The survey protocol was approved by the Institutional Review Boards (IRBs) of the Ethiopian Public Health Institute (EPHI), the Centers for Disease Control and Prevention (CDC), and Columbia University. Written informed consent was obtained from emancipated minors (aged 15–17) and adult participants (aged 18–64) for both interview and biomarker components. The IRBs waived the requirement for parental/guardian consent for emancipated minors. For non-emancipated minors (aged 15–17), written consent was obtained from a parent or guardian, accompanied by the participant's assent. All methods were performed in accordance with the relevant guidelines and regulations.

Results

Demographic, socioeconomic, and behavioral characteristics of study participants

The study analyzed data from 19,136 participants, the majority of whom were female (60.6%). The mean participant age was 29.95 ± 11.92 years (mean \pm standard deviation). The age distribution indicated a predominantly young adult population, with 39.4% aged 15–24 and 29.6% aged 25–34 years. Regarding educational attainment, 35.7% of participants had completed primary school, while 28.8% had completed secondary school. In terms of sexual behavior, most participants (65.7%) initiated sexual activity at age 18 or older. Furthermore, 16.2% reported engaging in non-marital or non-cohabiting sexual relationships in the 12 months preceding the survey. Among this subgroup, only 34.9% reported using condoms (Table 1).

HIV-1 prevalence and distribution

The overall HIV-1 prevalence was 3.0% (614/18,522; 95% CI: 2.6–3.5). Among PLWH, 15.3% (88/614; 95% CI: 12.2–18.9) reported being HIV-negative, while 10.5% (59/614; 95% CI: 7.9–13.7) were unaware of their status. Prevalence was higher among females (4.1%) than males (2.0%). Age-stratified analysis revealed an inverted U-shaped trend in HIV-1 prevalence, with rates increasing with age, peaking at 6.2% among individuals aged 35–44 years, and subsequently declining to 3.4% in the 55–64 age group. The prevalence decreased with higher educational levels, ranging from 5.2% (no formal education) to 1.0% (post-secondary education). Similarly, HIV-1 prevalence was higher among those who initiated sex before age 18 (5.1%) compared to those who began at 18 or older (3.2%). Additionally, participants reporting consistent condom use or abstinence in the 12 months before the survey had a higher prevalence (4.9%), while those with inconsistent use (2.0%) or no condom use (2.2%) showed lower rates.

HIV-1 infection duration, ART adherence, and treatment outcomes

Among the study participants, 0.5% (95% CI: 0.2–1.5) had acquired HIV-1 recent infections. Of those receiving ART, 12.4% (95% CI: 9.4–16.3) showed virological failure. Regarding immune status, 35.8% (221/614; 95% CI: 31.7–40.1) had CD4 T-cell counts below 350 cells/mm³. Approximately three-fourths (76.7%; 95% CI: 72.3–80.7) of participants were confirmed to be on ART, based on a combination of blood testing and self-reported data. Blood tests for specific ARVs revealed that 46.7% (95% CI: 41.8–51.6) tested positive for Efavirenz, 0.2% (95% CI: 0.0–1.6) for Lopinavir, and 26.0% (95% CI: 21.7–30.7) for Nevirapine (Table 2).

Variables	Weighted count (N)	Weighted %
Sex		
Male	7,537	39.4
Female	11,599	60.6
Age group (years)		
15–24	7,547	39.4
25–34	5,664	29.6
35–44	3,136	16.4
45–54	1,651	8.6
55–64	1,138	6.0
Marital status		
Married/ cohabiting	9,418	49.5
Previously married	2,495	13.1
Never married	7,103	37.4
Educational level		
No formal education	2,400	12.6
Primary school	6,803	35.7
Secondary school	5,488	28.8
Post-secondary	4,376	23.0
Region		
Tigray	1,369	7.2
Afar	821	4.3
Amhara	2,999	15.7
Oromia	4,510	23.6
Somali	926	4.8
Benishangul Gumuz	798	4.2
SNNPR	2,665	13.9
Gambella	788	4.1
Harari	697	3.6
Addis Ababa	2,780	14.5
Dire Dawa	783	4.1
Sexual experience		
Ever had sex	14,071	74.2
Never had sex	4,892	25.8
Age at first sex ($n = 14, 071$)		
< 18 years	4,759	34.3
≥ 18 years	9,098	65.7
Number of partners in the past 12 months ($n = 14, 071$)		
None	3,763	28.9
One	8,778	67.3
More than one	497	3.8
Partner relationship (past 12 months; $n = 9,275$)		
Marital/ cohabiting	7,766	83.8
Non marital/ non-cohabiting	1,504	16.2
Condom use consistency in the last 12 months (non-marital; $n = 1,504$)		
Consistent or abstinence	312	22.8
Inconsistent use	189	13.8
Never used	866	63.4
Household wealth index		
Low	6,757	35.3
Medium	3,810	19.9
High	8,569	44.8
Ever tested for HIV		
Continued		

Variables	Weighted count (N)	Weighted %
Yes	13,186	69.7
No	5,740	30.3
History of STI (n = 14,071)		
Yes	188	1.3
No	13,879	98.7

Table 1. Demographic, socioeconomic, and behavioral characteristics of study participants in Ethiopia (N = 19,136). HIV, human immunodeficiency virus; SNNPR, Southern Nations, Nationalities, and Peoples’ Region; STI, sexually transmitted infection.

Variables	Category	Weighted N	Weighted % (95% CI)
HIV-1 infection duration (n = 614)	Recent	5	0.5 (0.2–1.5)
	Long-term	609	99.5 (98.5–99.8)
Self-report ART status	On ART	410	94.9 (91.8–96.8)
	Not on ART	21	5.1 (3.2–8.2)
ART status (self-report + ARV testing)	ARVs detected + self-reported ART status	476	76.7 (72.3–80.7)
	ARVs not detected + self-reported not on ART	133	23.3 (19.3–27.7)
Duration of ART	≥ 24 Months	352	62.1 (56.6–67.3)
	12–23 months	18	2.8 (1.7–4.4)
	< 12 months	24	3.5 (2.3–5.5)
	Not on ART	168	31.6 (26.8–36.8)
Time since ART initiation (ever treated)	≥ 12 months	379	94.7 (92.0–96.5)
	< 12 months	25	5.3 (3.5–8.0)
ARV detection	Detected	459	72.9 (68.6–76.7)
	Not detected	155	27.1 (23.3–31.4)
Specific ARV detection	Efavirenz	292	46.7 (41.8–51.6)
	Lopinavir	1	0.2 (0.0–1.6)
	Nevirapine	166	26.0 (21.7–30.7)
Viral load suppression (n = 476)	Suppressed (VL < 1,000 copies/mL)	414	87.6 (83.7–90.6)
	Not suppressed (VL ≥ 1,000 copies/ mL)	62	12.4 (9.4–16.3)
CD4 count (cells/μL) (n = 614)	< 100	20	3.3 (2.1–5.1)
	100–199	62	10.8 (8.5–13.7)
	200–349	139	21.7 (17.8–26.1)
	350–499	156	28.1 (24.0–32.7)
	≥ 500	237	36.1 (31.6–40.9)

Table 2. HIV-1 infection duration, ART adherence, and treatment outcomes in study participants. ART, antiretroviral therapy; ARV, antiretroviral; CD4, cluster of differentiation 4; CI, confidence interval; HIV, human immunodeficiency virus; VL, viral load.

HIV-1 drug resistance mutations and genotyping outcomes

All successfully genotyped samples (100%, 42/42) were subtype C. The most prevalent resistance mutation was K103N, observed in 45.2% of samples (19/42), followed by V106M in 21.4% (9/42) for NNRTIs. For NRTIs, M184V was the most frequent mutation at 57.1% (24/42), followed by K65R at 21.4% (9/42) (Supp. Table 1). In most samples, 78.8% (33/42) showed resistance to at least one NNRTI, while 66.7% (28/42) showed mutations impacting at least one NRTI. Mutations conferring resistance to protease inhibitors (PIs) were found in only one sample. Furthermore, co-occurring resistance mutations for NNRTI and NRTI classes were present in 64.3% (27/42) of samples. Co-occurrence of mutations conferring resistance to all three ARV classes was infrequent, observed in only one individual (Fig. 2).

Factors associated with HIV-1 prevalence

The study identified multiple factors associated with HIV-1 prevalence. Females had significantly higher odds of HIV-1 infection than males (AOR: 1.7, 95% CI: 1.26–2.26). Regarding age, participants aged 35–44 and 45–54 years had the highest odds ratios (AOR: 6.7, 95% CI: 4.02–10.54 and AOR: 6.7, 95% CI: 3.88–11.52, respectively) compared to those aged 15–24 years. The study also found that education level significantly influenced HIV-1 prevalence; participants with primary education (AOR: 2.5, 95% CI: 1.64–3.71) and secondary school (AOR: 2.2, 95% CI: 1.43–3.36) had a higher likelihood of infection than those with post-secondary education. Marital status was another determinant; previously married individuals had significantly higher odds of infection (AOR:

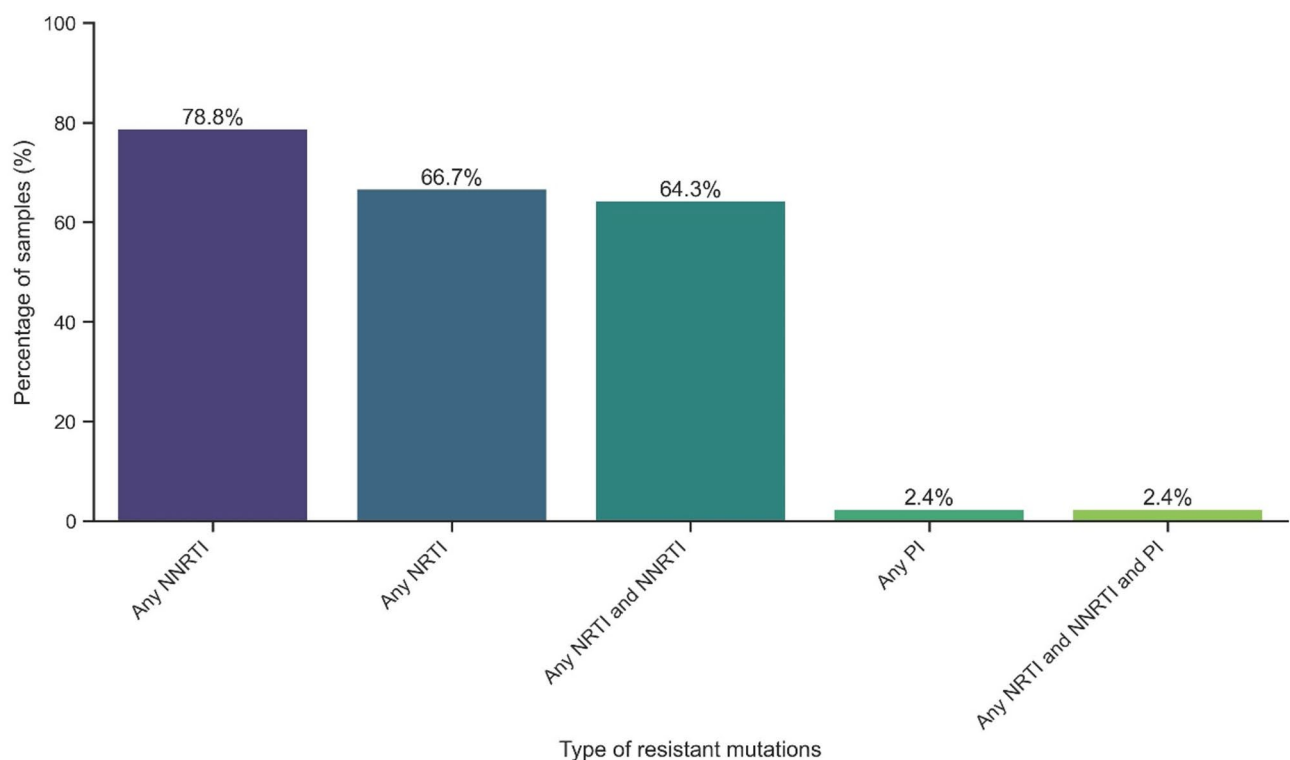


Fig. 2. HIV-1 resistance mutations in Ethiopia using the Ethiopian 2017–2018 PHIA survey datasets. NNRTIs, Non-Nucleoside Reverse Transcriptase Inhibitors; NRTIs, Nucleoside Reverse Transcriptase Inhibitors; PIs, Protease Inhibitors.

2.7, 95% CI: 1.47–4.95) compared to those never married. Regional variation was also evident; residents of Gambella had significantly higher odds of infection (AOR: 2.8, 95% CI: 1.52–5.31) than those in Tigray (the reference region). Participants whose last partner was neither marital nor cohabiting had increased odds of HIV-1 infection (AOR: 1.7, 95% CI: 1.08–2.67) compared to those in marital or cohabiting relationships. Additionally, individuals with a history of STIs had the highest odds ratio (AOR: 3.3, 95% CI: 1.85–5.78) compared to those without such a history (Table 3).

Discussion

Despite significant progress in prevention and treatment programs in Ethiopia, HIV remains a major public health challenge. In this study, the overall HIV-1 prevalence was found to be 3% (95% CI: 2.6–3.5), which aligns with the national estimates (2.9–6.0%)^{1,7} and falls within the range reported in Zambia (4.2–23.5%)³⁹. However, this result is much lower than the prevalence among female sex workers in Ethiopia, which ranges from 18.5% nationally⁴⁰ to 19.9% in Hawassa⁴¹ and 28.2% in Bahir Dar⁴². This variation may stem from the heightened vulnerability of female sex workers due to factors such as inconsistent condom use⁴⁰, stigma, discrimination⁴³, and limited access to healthcare services^{40,43,44}.

Cross-country comparisons further highlight regional variations. For instance, HIV prevalence in Lesotho (25.6%)⁴⁵ and South Africa (25.9–28.0%)⁴⁶ far exceeds Ethiopia's rates. These differences likely reflect variations in health system infrastructure, socioeconomic conditions, and cultural contexts that influence sexual behavior and HIV risk.

HIV-1 prevalence was significantly higher among women (4.1%) than among men (2.0%), representing a more than twofold difference. Several factors may contribute to this difference. Biologically, women are more susceptible to HIV acquisition during heterosexual intercourse due to anatomical differences that facilitate male-to-female transmission. Additionally, gender inequalities and cultural norms often limit women's ability to negotiate safe sex practices, while financial dependence may further heighten vulnerability, potentially leading to engagement in transactional sex. However, this prevalence is lower than the 24.1% reported in South Africa⁴⁷, likely reflecting broad differences between the two countries. These include variations in national HIV program effectiveness, socioeconomic conditions, health system infrastructures, and cultural norms governing sexual behavior and HIV risk. The HIV-1 incidence rate in this study was 0.5%, lower than the 1.5% reported in Lesotho⁴⁵, suggesting that new infections are being effectively prevented in the current study's context. Similarly, age-stratified analysis showed that HIV-1 prevalence peaked among adults aged 35–44 (6.2%) and 45–54 (6.1%). This result likely reflects cumulative exposure risk over time, as older individuals may have had more sexual partners or prolonged periods of untreated infection. Delayed diagnosis due to stigma and discrimination

Variables	HIV-1 status		OR (95% CI)	AOR (95% CI)	p-value
	Negative	Positive			
Sex					
Male	7,384	153	Reference	Reference	
Female	11,138	461	2.0 (1.66–2.40)	1.7 (1.26–2.26)	< 0.001
Age in years					
15–24	7,485	62	Reference	Reference	
25–34	5,489	175	3.9 (2.88–5.15)	2.8 (1.74–4.41)	< 0.001
35–44	2,902	234	9.7 (7.33–12.92)	6.7 (4.02–10.54)	
45–54	1,547	104	8.1 (5.90–11.17)	6.7 (3.88–11.52)	
55–64	1,099	39	4.3 (2.86–6.42)	2.6 (1.09–5.96)	0.031
Educational level					
No formal education	2,279	121	4.0 (2.88–5.43)	1.8 (1.06–2.90)	0.029
Primary school	6,512	291	3.3 (2.50–4.42)	2.5 (1.64–3.71)	0.001
Secondary school	5,347	141	2.0 (1.44–2.67)	2.2 (1.43–3.36)	
Post-secondary	4,318	58	Reference	Reference	
Marital status					
Married/ cohabiting	9,133	285	3.1 (2.38–4.02)	1.5 (0.78–2.84)	-
Previously married	2,239	256	11.3 (8.67–14.79)	2.7 (1.47–4.95)	0.001
Never married	7,032	71	Reference	Reference	
Region					
Tigray	1,330	39	Reference	Reference	
Afar	789	32	1.4 (0.86–2.23)	1.1 (0.51–2.27)	-
Amhara	2,881	118	1.4 (0.97–2.02)	1.5 (0.89–2.63)	-
Oromia	4,361	149	1.2 (0.81–1.67)	1.1 (0.65–1.84)	-
Somali	918	8	0.3 (0.14–0.64)	0.5 (0.16–1.48)	-
Benishangul Gumuz	778	20	0.9 (0.51–1.51)	1.4 (0.64–3.01)	-
SNNPR	2,616	49	0.6 (0.42–0.98)	0.8 (0.45–1.51)	-
Gambella	744	44	2.0 (1.30–3.13)	2.8 (1.52–5.31)	0.001
Harari	665	32	1.6 (1.02–2.64)	1.9 (0.94–3.82)	-
Addis Ababa	2,692	88	1.1 (0.76–1.64)	1.0 (0.56–1.78)	-
Dire Dawa	748	35	1.6 (1.00–2.54)	1.5 (0.75–2.98)	-
Household wealth index					
Low	6,546	211	Reference	Reference	
Medium	3,667	143	1.2 (0.98–1.50)	1.2 (0.85–1.64)	-
High	8,309	260	1.0 (0.81–1.17)	0.9 (0.66–1.28)	-
Age at first sex					
< 18 years	4,505	254	1.5 (1.30–1.83)	1.1 (0.87–1.50)	-
≥ 18 years	8,777	321	Reference	Reference	
Relationship with last partner					
Marital/ cohabiting	7,531	235	Reference	Reference	
Non marital/ non-cohabiting	1,444	60	1.3 (0.99–1.78)	1.7 (1.08–2.67)	0.023
Ever tested for HIV					
Yes	12,625	561	5.0 (3.71–6.61)	2.1 (1.36–3.30)	0.001
No	5,689	51	Reference	Reference	
History of STI					
Yes	160	28	4.3 (2.82–6.41)	3.3 (1.85–5.78)	< 0.001
No	13,330	549	Reference	Reference	

Table 3. Factors associated with HIV-1 prevalence in Ethiopian urban areas using the 2017– 2018 PHIA survey dataset. AOR, Adjusted Odds Ratio; CI, Confidence Interval; HIV, Human Immunodeficiency Virus; OR, Odds Ratio; PHIA, Population-based HIV Impact Assessment; SNNPR, Southern Nations, Nationalities, and Peoples' Region; STI, Sexually Transmitted Infection.

could further exacerbate transmission in these age groups, as fear of social repercussions may deter testing and treatment-seeking.

The study found that 12.4% of PLWH on ART experienced virological failure, indicating persistent challenges in treatment adherence and effectiveness. This finding is consistent with studies from Ethiopia, which reported

virological failure rates of 15.9% and 15.95%^{48,49}. However, our result is higher than those reported in other African regions (7.3–9%)^{50,51}, suggesting potential regional variations in treatment outcomes. Even higher rates of virological failure have been documented in Ethiopia (18.0%)⁵² and Asia (17.9%)⁵³, indicating that suboptimal viral suppression remains a significant concern in multiple high-burden settings.

In the current study, a significant discrepancy was observed between self-reported HIV-1 status and laboratory-confirmed results, with 15.3% of PLWH incorrectly reported as negative status. This mismatch might be due to gaps in service delivery, including inadequate patient education, poor linkage to care, or insufficient counseling during HIV testing. It also highlights critical deficiencies in HIV-1 status awareness, which may hinder treatment initiation and retention in care.

Though Ethiopia has made significant progress toward the UNAIDS 90-90-90 targets, with 97.1% of diagnosed HIV patients receiving ART and 87.6% achieving viral suppression⁵⁴, the emerging drug resistance poses a critical challenge, particularly in patients requiring second-line therapy. Our analysis of 42 HIV-positive samples revealed a high prevalence of DRMs, with NNRTIs and NRTIs being the most affected. NNRTI resistance was prevalent, observed in 76.7% of samples, with K103N (44.2%) and V106M (23.3%) being the most frequent mutations. The high prevalence of K103N is concerning, as it confers resistance to key first-line NNRTIs, including efavirenz and nevirapine, potentially undermining treatment efficacy. Similarly, NRTI resistance was detected in 65.1% of samples, with M184V (55.8%) and K65R (20.9%) being the predominant mutations. While M184V is associated with resistance to lamivudine and emtricitabine, it also increases susceptibility to tenofovir and zidovudine, which can be leveraged in regimen optimization⁵⁵. The NADIA trial further supports the efficacy of tenofovir-containing regimens even in the presence of M184V, demonstrating high virological suppression rates when combined with dolutegravir⁵⁶. These findings suggest that a substantial proportion of individuals in the study population may be experiencing treatment failure due to ineffective drug regimens, necessitating a transition to second-line therapies. On the other hand, resistance to PIs was low, with only a single (M46I) mutation identified. This indicates that PIs remain a viable and effective alternative for patients who resist NNRTIs and NRTIs. Given the high resistance rates to first-line regimens, enhanced resistance monitoring, optimized ART regimens, and timely switches to second-line therapies are essential to maintaining treatment success and preventing further transmission of resistant strains.

The current study identified multiple factors associated with HIV-1 prevalence in Ethiopia. Female participants were 1.7 times more likely to be HIV-1 positive than male participants. This finding is supported by a previous study from Ethiopia⁵⁷. The variation may be due to gender inequality that significantly contributes to higher rates of HIV infection among women⁵⁷. In terms of age, participants aged 35–44 and 45–54 years each 6.7 times more likely to be HIV-1 positive compared to those aged 15–24 years. A similar pattern was observed in China, where older age groups faced an increased HIV risk⁵⁸. This difference could reflect factors such as prolonged exposure to risk behaviors, lower testing rates, or delayed diagnosis in older populations. The study also found that participants with primary or secondary school education were 2.5 and 2.2 times more likely, respectively, to have HIV-1 compared to participants with post-secondary education. This suggests that higher education may protect against HIV infection. Consistent with this, studies in Africa have also linked higher education levels to lower HIV prevalence⁵⁹. Regarding marital status, participants who were previously married (divorced, separated, or widowed) were 2.7 times more likely to contract HIV-1, while currently married/cohabiting participants were 1.5 times more likely, respectively, compared to never-married individuals. This aligns with a Nigerian study where previously married individuals had⁶⁰. Their vulnerability may stem from disrupted social networks, economic instability, or engagement in transactional sex, as seen in Malawi and Uganda^{61,62}. Geographically, residents of the Gambella region were 2.8 times more likely to be HIV-1 positive than those in Tigray. This may reflect inadequate government focus on HIV prevention, including the discontinuation of community programs, which reduced adherence to prevention⁶³. Additionally, individuals with an STI history had a 3.3-fold higher likelihood of HIV-1 infection. STIs, especially those causing genital ulcers, increase susceptibility by damaging mucosal barriers. For example, syphilis, chlamydia, or gonorrhea elevate risk due to inflammation-driven immune responses⁶⁴.

Conclusions

This study identified a high prevalence of HIV-1 in Ethiopia. The findings showed major challenges to treatment effectiveness, including a high virological failure rate, a significant proportion of people living with HIV being unaware of or misreporting their status, and widespread drug resistance to common first-line NNRTI and NRTI antiretrovirals. Multiple factors were associated with HIV-1 prevalence, including sex, age, education level, marital status, region, and history of STIs. To improve treatment outcomes and curb transmission, Ethiopia should strengthen HIV prevention strategies focused on vulnerable populations, enhance testing and linkage to care, and implement routine drug resistance monitoring.

Data availability

The datasets used or analyzed for this study are available and can be accessed with the necessary access credentials from the PHIA program (https://phia-data.icap.columbia.edu/datasets?country_id=12).

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Author contributions

All authors participated in formulating the research question(s) and designing the study. TD contributed to the data analysis and interpreting the results, to writing the original draft, and to reviewing and editing. MG contributed to the supervision of the project, interpretation of findings, and to reviewing and editing. AB and AW contributed to the interpretation of the findings and to reviewing and editing. All authors read and approved the final version of the manuscript before submission.

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Declarations

Competing interests

The authors declare no competing interests.

Additional information

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