

RESEARCH ARTICLE

Population prevalence of antiretroviral therapy sharing and its association with HIV viremia in rural Uganda: a cross-sectional population-based study

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Abstract

Introduction: Antiretroviral treatment (ART) sharing has been reported among fishermen and sex workers in Uganda and South Africa. However, no population-based studies have documented ART diversion prevalence (including sharing [giving/receiving], buying and selling) or its relationship with viremia among men and women living with HIV in Africa.

Methods: In 2018–2020, we surveyed people living with HIV aged 15–49 years in 41 communities in the Rakai Community Cohort Study, a population-based cohort in south-central Uganda. We assessed the prevalence and correlates of self-reported lifetime and past-year ART diversion, stratifying by age and gender and documenting sources of diverted drugs. We used log-binomial regression to quantify the relationship between diversion patterns and viremia (viral load >40 copies/ml), reported as unadjusted and adjusted prevalence ratios (aPR) with 95% confidence intervals (CI).

Results: Of 2852 people living with HIV and self-reporting current ART use, 266 (9.3%) reported lifetime ART diversion. Giving/receiving drugs were most common; few participants reported buying, and none reported selling. Men (12.9%) were more likely to report lifetime diversion than women (7.4%), with men aged 25–34 reporting high levels of sharing (18.9%). Friends were the most common sources of shared drugs, followed by spouses/sexual partners. Patterns of lifetime and past-year diversion were similar. Among participants with viral load results, 8.6% were viraemic. In adjusted analyses, people who reported only giving ART were nearly twice as likely to be viraemic than those who reported no diversion (aPR: 1.94, 95% CI: 1.10–3.44), and those reporting only receiving ART were less likely to exhibit viremia (aPR: 0.46, 95% CI: 0.12–1.79), although the latter was not statistically significant. Reporting both giving and receiving ART was not associated with viremia (aPR: 0.79, 95% CI: 0.43–1.46). Reporting buying ART, though rare, was also correlated with higher rates of viremia, but this relationship was not statistically significant (aPR: 1.98, 95% CI: 0.72–5.45).

Conclusions: ART sharing is common among persons reporting ART use in rural Uganda, particularly among men. Sharing ART was associated with viremia, and receiving ART may facilitate viral suppression. HIV programmes may benefit from considering ART sharing in counselling messages.

Keywords: HIV care continuum; HIV adherence; ART sharing; ART division; viremia; Uganda

Additional information may be found under the Supporting Information tab of this article.

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1 | INTRODUCTION

Drug diversion is broadly conceptualized as the transfer of prescribed medication from the individual to whom it was prescribed to a different individual [1]. Drug diversion may include sharing (giving and receiving), buying or selling drugs. Drug sharing is frequently reported across classes of

prescription medication, with rates of borrowing ranging from 5% to 52% across studies, and lending from 6% to 23% [2].

Globally, 38.4 million people are estimated to be living with HIV, 75% of whom are taking daily antiretroviral treatment (ART) [3]. Despite the enormous global attention to HIV treatment, there has been limited research on ART drug diversion. Research from the United States in South Florida has

identified illicit marketplaces where ART is sold or traded [4–12]. Reported reasons for ART diversion in that setting include substance use, [4, 5, 8, 10], food and housing insecurity [6–8], barriers to accessing HIV care [4, 5, 8] and perceived efficacy of ART for HIV prevention (e.g. using ART for pre-exposure prophylaxis [PrEP]) [10, 12]. Other studies from South Africa have reported ART use in a recreational drug cocktail [13–17]. These examples of drug diversion for illicit purposes were for ART combinations that contained efavirenz, and may be less likely to occur now following the switch to dolutegravir (DTG)-based regimens.

While these studies are important, there have been few explorations of drug sharing for therapeutic HIV use, though drug sharing is common for other medications. To date, just one study has quantitatively assessed ART diversion for therapeutic use. This study among female sex workers living with HIV in eThekwini (Durban), South Africa, found that 30% reported ever sharing (giving and/or receiving) ART [18]. Factors associated with sharing ART included higher levels of alcohol use, illicit drug use, depression severity and physical/sexual violence [18], although in qualitative interviews, female sex workers reported they shared ART to avoid seeking clinical care or because they missed appointments [19]. The study found a modest dose-response relationship between the number of ART pills that participants reported giving to their peers in the last 30 days and viral suppression; however, there was no relationship between viral suppression and number of pills received or given [18].

In 2017, we conducted a qualitative study among female sex workers and fishermen living with HIV in hyperendemic fishing communities along Lake Victoria in Rakai, Uganda [20]. Participants reported frequent, episodic ART sharing with coworkers/friends, motivated by the desire to remain adherent to medication despite challenges to regular clinic attendance or stock-outs. High HIV prevalence in these communities seemed to facilitate drug diversion, as participants were likely to know other people living with HIV. However, this qualitative study did not assess ART diversion among broader populations of people living with HIV, nor did it examine the relationship between diversion and health outcomes. Although our preliminary qualitative research found that ART diversion was used to support adherence and would thus be expected to be related to improved viral suppression, sharing drugs also implies insufficient ART access and running out of medications, so it may be associated with poorer adherence and reduced viral suppression.

Here, we sought to measure the prevalence of and factors associated with ART diversion, and to assess the relationship between ART diversion and viral suppression, in a representative population of people living with HIV in south-central Uganda.

2 | METHODS

2.1 Study setting and population

This study was conducted in Rakai and neighbouring districts in south-central Uganda, where the Rakai Health Sciences Program (RHSP) has conducted epidemiological and HIV prevention research since 1988. RHSP conducts the Rakai Com-

munity Cohort Study (RCCS), an ongoing HIV surveillance study with over 30 years of population-based, longitudinal data on HIV prevalence and incidence, risk behaviours, sexual networks, migration and service coverage [21]. Approximately every 18 months, the RCCS conducts household censuses, extensive interviews and HIV testing with all consenting individuals aged 15 and older in 41 study communities. These study communities are diverse and include inland agrarian villages, trading towns and fishing villages on Lake Victoria. Written informed consent is obtained from all adults and emancipated minors, and parental consent and minor assent is obtained for all unemancipated minors aged 15–17.

Questions about ART diversion were added to the RCCS survey round 19, conducted from 19 June 2018 to 6 November 2020. The analytic sample included all RCCS participants living with HIV aged 15–49 years who were aware of their HIV status and who self-reported current ART use. The accuracy of self-reported ART use in the cohort has previously been validated by plasma detection of antiretroviral drugs [22]. Associations with viral load were assessed among participants with viral load results.

2.2 | Measures

To measure ART diversion, we asked participants to self-report whether they had ever, or in the past 12 months, bought ART, sold ART, shared ART with someone else or had someone else share ART with them. For those who responded yes to any type of diversion, we elicited the drug recipients/sources involved in the reported diversion type (bought/sold/shared); responses included spouse/sexual partner, other family member, friend, work colleague, stranger or other.

HIV status was assessed using a validated three-test algorithm followed by confirmatory enzyme immunoassays. Viral load assays were performed on stored plasma using the Abbott RealTime HIV-1 assay (Abbott Molecular, Inc., Des Plaines, IL), and viremia was defined as having a detectable viral load (>40 copies/ml); we also ran sensitivity analyses with viral load ≥1000 copies/ml, the current World Health Organization (WHO) criteria [23].

Socio-demographic measures included age, gender, marital status (never married, currently married, previously married [i.e. separated, divorced, widowed]), highest level of education (none, primary, secondary or higher), religion, occupation, migration status (recent in-migrant to the community since the last survey round) and community of residence (fishing, trading or agrarian). Sexual behaviours in the past 12 months and other factors associated with HIV transmission were also assessed, including number and type of sexual partners (marital, non-marital), condom use with non-marital partners, exchange of money or goods for sex, alcohol use before sex (by self or partner), sex with partners outside their community and symptoms of genital ulcer disease. Current pregnancy status was assessed among women, and circumcision status was assessed among men.

2.3 | Statistical analysis

We examined the prevalence of ART diversion overall among persons reporting ART use and by type of diversion, stratified by age, gender and community of residence, in the past 12 months and ever. We compared socio-demographic characteristics and sexual behaviours among individuals who did and did not report ART diversion, and who reported different types of diversion using Pearson's Chi-square test. We summarized data on drug recipients/sources by gender and type of ART diversion reported.

We used log-binomial regression to quantify the relationship between lifetime ART diversion and viremia, reported as prevalence ratios (PR) and adjusted prevalence ratios (aPR) with 95% confidence intervals (CI). We hypothesized that individuals who reported giving or receiving ART would be more likely to be viraemic than those who did not report sharing. We first conducted unadjusted analyses for the whole sample, and stratified by gender and community of residence. We then conducted additional regression analyses adjusting for gender, age (in years) and community of residence (classified as fishing vs. non-fishing [inland] communities).

3 | RESULTS

Of 19,341 RCCS participants, 3374 (17%) were living with HIV, of whom 2852 (85%) self-reported knowledge of their serostatus and currently taking ART. Of people living with HIV self-reporting ART use, approximately two-thirds were women (n=1868/2852; 65%), and one-third were men (n=984/2852; 35%). Most were older than 35 years, had completed primary education or less and were employed in agricultural/housework, trading or fishing. Overall, 9.3% (n=266/2852) reported lifetime ART diversion, while 6.8% (n=193/2852) reported ART diversion in the past year.

The practice of giving and receiving ART were the most commonly reported types of diversion; 62 participants (2.2%) reported giving ART only, 54 (1.9%) reported receiving ART only and 132 (4.6%) reported both giving and receiving (Table 1). Few participants reported buying ART (n=18, 0.6%), and none reported selling. Men were more likely to report any diversion than women (12.9% vs. 7.4%). Stratifications by gender and age showed that men aged 25–34 years reported ART sharing most frequently (18.9%) (Figure 1). Patterns of giving, receiving, and buying ART and associations with socio-demographic factors were similar for lifetime and past-year diversion (Supplementary Table).

People who reported lifetime ART diversion were also generally more likely to report several behaviours associated with potential HIV transmission risk, including alcohol use before sex, sex with partners outside their community and symptoms of genital ulcer disease in the last 12 months (Table 2).

Friends were the most commonly reported recipients and sources of shared drugs: 57.8% (n=108/187) of participants reporting any past-year ART diversion said they had done so with a friend (Figure 2). Spouses/sexual partners were the next most common at 21.9% (n=41/187), and other family members followed at 14.4% (n=27/187). Almost no one reported sharing with strangers. Men and women generally reported similar recipients and sources of shared ART, although women (compared to men) were more likely to report sharing with other family members, while men (com-

pared to women) were more likely to report sharing with work colleagues.

Of 2852 people living with HIV self-reporting ART use, 2725 (95.5%) had viral load results, and 8.6% of these (234/2725) were viraemic. Associations between lifetime ART diversion and viremia were generally consistent in unadjusted, adjusted and stratified analyses. In adjusted analyses, people who reported only giving ART to others were nearly twice as likely to be viraemic than those who reported no diversion (aPR: 1.94, 95% CI: 1.10-3.44), while those who reported only receiving ART were less likely to be viraemic compared to those who reported no diversion, but this difference was not statistically significant (aPR: 0.46, 95% CI: 0.12-1.79) (Table 3). Reporting both giving and receiving was not associated with viremia (aPR: 0.79, 95% CI: 0.43-1.46). Reporting buying ART, though rare, also seemed associated with viremia, but was not statistically significant (aPR: 1.98, 95% CI: 0.72-5.45). Stratifications by gender yielded similar overall patterns but were not statistically significant. Sensitivity analyses with viral load ≥1000 copies/ml also yielded similar patterns, but only remained statistically significant for people reporting giving ART to others only.

4 | DISCUSSION

To our knowledge, this is the first population-based study documenting the prevalence of ART diversion and its relationship to HIV viremia among men and women living with HIV in the general population in Africa. We found that ART sharing is fairly common, particularly among men, and is associated with viremia.

We identified a relationship between ART sharing and viremia, with worse outcomes observed among people giving pills, potentially better outcomes observed among people receiving pills and no relationship seen among people both giving and receiving. These results suggest that sharing drugs with another person could precipitate a drug shortage for the lender (thus interfering with adherence and viral suppression), or people who are poorly adherent themselves may be more likely to share because they have extra pills available to give away. Conversely, borrowing drugs may facilitate treatment adherence for the borrower. Reporting both borrowing and sharing drugs may reflect a reciprocal arrangement to address more persistent barriers to ART access, which, on balance, may have no overall impact on adherence and viral suppression. Purchasing ART, although rare, may also be associated with viremia.

These findings differ from the single prior study among female sex workers in South Africa [18], which found that only giving was associated with higher (not lower) levels of viral suppression. That study used a different ART sharing measure and timeframe (number of pills shared in the last 30 days) and focused on sharing only with sex work colleagues and not sharing with other individuals, or buying and selling ART. It is thus unclear if the differing results are due to different methodological approaches or true underlying differences in different populations. Sex workers in South Africa may have strong relationships with each other due to the multiple stigmas and difficult life circumstances they face, which may not

Table 1. Socio-demographic factors associated with ever giving, receiving or buying ART among persons living with HIV self-reporting ART use in south-central Uganda (N=2852).

		ART diversion (N=266)							
	No ART		Received	Gave and		Any ART			
Socio-demographic	diversion	Gave only	only	received	Bought	diversion			
factors	(N=2586)	N=62	N=54	N=132	N=18	N=266	p-value*		
Age (years)									
15-24	154 (91.1%)	5 (3.0%)	4 (2.4%)	5 (3.0%)	1 (0.6%)	15 (8.9%)	0.073		
25-34	862 (88.2%)	25 (2.6%)	20 (2.1%)	63 (6.5%)	7 (0.7%)	115 (11.8%)			
35-49	1570 (92.0%)	32 (1.9%)	30 (1.8%)	64 (3.8%)	10 (0.6%)	136 (8.0%)			
Gender									
Female	1729 (92.6%)	41 (2.2%)	33 (1.8%)	56 (3.0%)	9 (0.5%)	139 (7.4%)	< 0.001		
Male	857 (87.1%)	21 (2.1%)	21 (2.1%)	76 (7.7%)	9 (0.9%)	127 (12.9%)			
Community of residence									
Agrarian	838 (94.9%)	9 (1.0%)	8 (0.9%)	24 (2.7%)	4 (0.5%)	45 (5.1%)	< 0.001		
Trading	582 (92.8%)	13 (2.1%)	13 (2.1%)	15 (2.4%)	4 (0.6%)	45 (7.2%)			
Fishing	1166 (86.9%)	40 (3.0%)	33 (2.5%)	93 (6.9%)	10 (0.8%)	176 (13.1%)			
Primary occupation									
Agriculture/housework	1008 (94.3%)	15 (1.4%)	10 (0.9%)	33 (3.1%)	3 (0.3%)	61 (5.7%)	< 0.001		
Bar/restaurant work	231 (87.8%)	9 (3.4%)	8 (3.0%)	12 (4.6%)	3 (1.1%)	32 (12.2%)			
Boda boda/trucking	22 (88.0%)	0 (0.0%)	2 (8.0%)	1 (4.0%)	0 (0.0%)	3 (12.0%)			
Fishing	338 (80.5%)	12 (2.9%)	10 (2.4%)	53 (12.6%)	7 (1.7%)	82 (19.5%)			
Student	5 (100.0%)	0 (0.0%)	0 (0.0%)	0 (0.0%)	0 (0.0%)	0 (0.0%)			
Trade/shop keeper	495 (92.4%)	10 (1.88%)	9 (1.7%)	18 (3.4%)	4 (0.8%)	41 (7.7%)			
Other	487 (91.2%)	16 (3.0%)	15 (2.8%)	15 (2.8%)	1 (0.2%)	47 (8.8%)			
Educational status	107 (711270)	10 (0.070)	10 (2.070)	10 (2.070)	1 (0.270)	., (0.0,0)			
None	479 (91.1%)	12 (2.3%)	11 (2.1%)	21 (4.0%)	3 (0.6%)	47 (8.9%)	0.240		
Primary	1679 (90.2%)	41 (2.2%)	35 (1.9%)	98 (5.3%)	8 (0.4%)	182 (9.8%)	0.2 10		
Secondary	379 (91.6%)	8 (1.9%)	8 (1.9%)	13 (3.1%)	6 (1.5%)	35 (8.5%)			
Tertiary	49 (96.1%)	1 (2.0%)	0 (0.0%)	0 (0.0%)	1 (2.0%)	2 (3.9%)			
Religion	17 (70.170)	1 (2.070)	0 (0.070)	0 (0.070)	1 (2.070)	2 (0.770)			
Catholic	1731 (90.5%)	41 (2.1%)	33 (1.7%)	97 (5.1%)	11 (0.6%)	182 (9.5%)	0.088		
Muslim	304 (90.8%)	8 (2.4%)	9 (2.7%)	13 (3.9%)	1 (0.3%)	31 (9.3%)	0.000		
Protestant	435 (91.8%)	11 (2.3%)	5 (1.1%)	17 (3.6%)	6 (1.3%)	39 (8.2%)			
Other	116 (89.2%)	2 (1.5%)	7 (5.4%)	5 (3.9%)	0 (0.0%)	14 (10.8%)			
Marital status	110 (07.270)	2 (1.570)	7 (3.470)	3 (3.770)	0 (0.070)	14 (10.070)			
Never married	149 (90.9%)	5 (3.1%)	4 (2.4%)	5 (3.1%)	1 (0.6%)	15 (9.2%)	0.230		
Married, monogamous	1237 (90.4%)	27 (2.0%)	25 (1.8%)	74 (5.4%)	6 (0.4%)	13 (9.2%)	0.230		
union	1237 (70.470)	27 (2.076)	23 (1.0%)	74 (3.470)	0 (0.4%)	132 (7.070)			
Married, polygamous union	321 (88.0%)	11 (3.0%)	6 (1.6%)	22 (6.0%)	5 (1.4%)	44 (12.1%)			
Previously married	879 (92.1%)	19 (2.0%)	19 (2.0%)	31 (3.3%)	6 (0.6%)	75 (7.9%)			
Migration status	/	,,	,/	,,	. (/	. (/			
Recent in-migrant	368 (91.5%)	7 (1.7%)	10 (2.5%)	14 (3.5%)	3 (0.8%)	34 (8.5%)	0.600		
Pregnancy status (among v		, /	,/	,,	(/	(/			
Pregnant Pregnant	117 (90.0%)	3 (2.3%)	3 (2.3%)	5 (3.9%)	2 (1.5%)	13 (10.0%)	0.418		

Note: No participants reported selling ART, so selling is not included in this table.

^{*}p-value for chi-square test by ART diversion types, comparing any antiretroviral treatment (ART) diversion (gave only, received only, gave and received, bought or sold) to no ART diversion. All p-values were two-sided and p<0.05 was considered statistically significant.

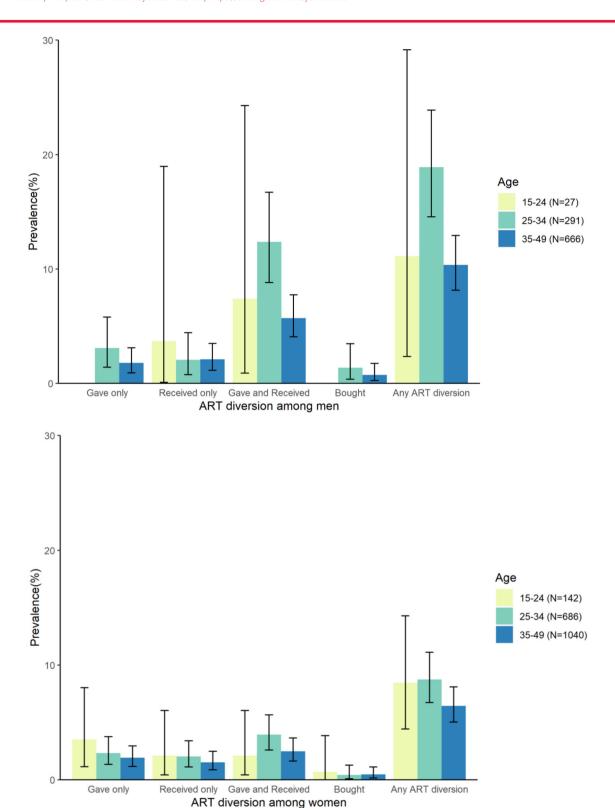


Figure 1. Prevalence of ever giving, receiving and buying antiretroviral treatment (ART) by age category and gender among people living with HIV ages 15-49 in 41 study communities in south-central Uganda.

Table 2. Sexual behaviours associated with ever giving, receiving or buying ART among persons living with HIV self-reporting ART use in south-central Uganda (N=2852)

		ART diversion (N=266)							
Sexual behaviours	No ART diversion (N=2586)	Gave only N=62	Received only N=54	Gave and received N=132	Bought N=18	Any ART diversion N=266	p-value*		
Sexually active in the past year	2236 (90.2%)	56 (2.3%)	49 (2.0%)	122 (4.9%)	17 (0.7%)	244 (9.8%)	0.180		
Multiple sexual partners in the past year	616 (85.4%)	23 (3.2%)	19 (2.6%)	55 (7.6%)	8 (1.1%)	105 (14.6%)	<0.001		
Non-marital partners in the past year (among married persons, <i>N</i> =2480)	980 (88.3%)	31 (2.8%)	26 (2.3%)	62 (5.6%)	11 (1.0%)	130 (11.7%)	0.055		
Consistent condom use with non-marital partners (among those with non-marital partners, <i>N</i> =1110)	228 (89.1%)	5 (2.0%)	5 (2.0%)	13 (5.1%)	5 (2.0%)	28 (10.9%)	0.360		
Alcohol use by respondents or partners before sex	1605 (88.3%)	50 (2.8%)	41 (2.3%)	106 (5.8%)	15 (0.8%)	212 (11.7%)	<0.001		
Sex with partners outside community	174 (84.5%)	4 (1.9%)	6 (2.9%)	20 (9.7%)	2 (1.0%)	32 (15.5%)	0.013		
Money, gifts or favours exchanged for sex with partner	294 (85.2%)	11 (3.2%)	13 (3.8%)	21 (6.1%)	6 (1.7%)	51 (14.8%)	0.052		
Symptoms of genital ulcer disease in the last 12 months	130 (83.9%)	5 (3.2%)	7 (4.5%)	10 (6.5%)	3 (1.9%)	25 (16.1%)	0.010		
Circumcised (among men, N=983)	461 (89.3%)	9 (1.7%)	11 (2.1%)	33 (6.4%)	2 (0.4%)	55 (10.7%)	0.495		

Note: No participants reported selling ART, so selling is not included in this table.

Table 3. Regression results showing the association between sharing, giving and buying ART and viremia (viral load >40 copies/ml) among persons living with HIV self-reporting ART use in south-central Uganda (N=2725)

ART diversion	Viraemic/total (%)	PR ^a (95% CI)	p-value	aPR ^b (95% CI)	p-value
No diversion	209/2469 (8.5)	Ref	-	Ref	-
Gave only	10/58 (17.2)	2.04 (1.14-3.63)	0.016	1.94 (1.10, 3.44)	0.022
Received only	2/49 (4.1)	0.48 (0.12-1.89)	0.294	0.46 (0.12, 1.79)	0.261
Gave and received	10/131 (7.6)	0.90 (0.49-1.66)	0.740	0.79 (0.43, 1.46)	0.458
Bought	3/18 (16.7)	1.97 (0.70-5.58)	0.202	1.98 (0.72, 5.45)	0.185

Note: No participants reported selling ART, so selling is not included in this table.

apply to our more diverse sample of men and women from rural communities in Uganda with a range of occupations and ART sharing partners. Ultimately, we feel these differences call for further research to better understand patterns of, reasons for and consequences of ART sharing across settings.

Men in our study were more likely to report sharing ART than women, and younger men (aged 25–34 years) had the highest overall levels of sharing. This group is also least likely to be virally suppressed in the RCCS overall. Our results are similar to findings from multiple countries in East and

^{*}p-value for chi-square test by ART diversion types, comparing any antiretroviral treatment (ART) diversion (gave only, received only, gave and received or bought) to no ART diversion. All p-values were two-sided and p<0.05 was considered statistically significant.

^aPrevalence ratio (PR) and 95% confidence interval (CI), comparing the prevalence of viremia between a specific type of antiretroviral treatment (ART) diversion and no ART diversion.

^bAdjusted prevalence ratio (aPR), adjusting for sex, age in years and community type (fishing vs. non-fishing).

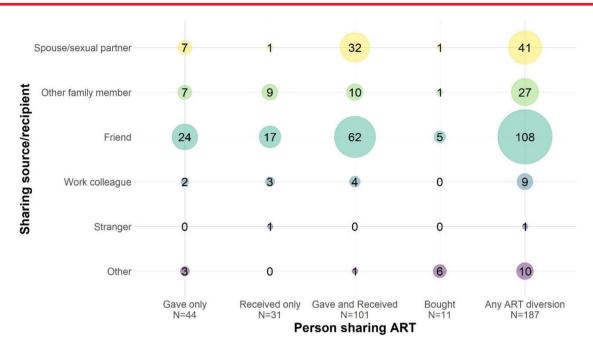


Figure 2. Recipients and sources of antiretroviral treatment (ART) given, received or bought in the past 12 months among persons living with HIV self-reporting ART use and ART diversion in south-central Uganda (N=187).

Southern Africa, where men are less likely to seek HIV testing, engage in care, adhere to ART or be virally suppressed [24]. Masculine norms likely influence these behaviours across settings [25], and health systems that are not aligned with men's health needs or social circumstances, such as work-related mobility, may also prevent engagement in care [24]. We also found that ART sharing was associated with several sexual risk behaviours, suggesting potential gendered clustering of risk. Finally, we found that ART sharing occurred more frequently in fishing communities, which also have documented different patterns of sexual risk behaviours, migration, occupational experiences and social norms that may correlate with ART diversion [26, 27].

Friends were the most reported source or recipient of shared ART. This was somewhat surprising, since studies show other drugs are commonly shared among family members [2], and HIV often clusters within families. However, ART sharing with friends was consistent with findings from our previous qualitative work among fishermen and female sex workers [20]. Another study from northern Uganda also found that neighbours were the most reported source of shared prescription drugs, primarily pain medications [28]. Communal living situations and the high prevalence of HIV may make drug sharing more common among friends in our study setting. Reciprocity has also been described as "a core cultural value" [29] in Uganda, and social norms may support patterns of drug sharing among friends and neighbours in ways that differ from drug sharing in other settings.

The associations we found between ART sharing and viremia have programmatic implications. Although there has been a recent expansion of differentiated service delivery options that offer more flexible ART access, health system inflexibilities around ART distribution remain, and the lend-

ing/borrowing patterns we observed suggest continued ART access barriers. Currently, ART sharing is not discussed in Ugandan or WHO guidelines for ART counselling. Asking clients about their experiences with drug sharing, and discussing both the potential benefits and consequences of drug sharing, may improve counselling messages. ART sharing may also raise concerns around drug resistance. These concerns may be lessened as the majority of people on ART in Uganda are now prescribed ART regimens containing DTG, which are anticipated to have a higher genetic barrier to resistance [30], although there has been recent documentation of DTG resistance in African ART programmes [31].

Some research has suggested that antiretroviral drugs meant for ART [10, 12] or PrEP [32, 33] are shared with HIV-negative individuals for HIV prevention, although other studies have shown PrEP sharing is rare [34]. We only included individuals who self-reported knowing their HIV status and self-reported being on ART, and we used the term ART rather than PrEP, which looks visibly different than ART pill formulations. We, therefore, think it was unlikely that we captured PrEP sharing. However, some proportion of participants who reported only giving ART to others may have been sharing with HIV-negative individuals for prevention.

Our research was mostly conducted prior to the COVID-19 pandemic in Uganda, during which ART distribution options became more flexible through expanded differentiated service delivery models, such as community client-led ART delivery and community drug distribution points, and networks of ART sharing may have been disrupted due to COVID-related mobility constraints during lockdown periods. However, RCCS surveys have found no change in viral suppression rates following COVID [35]. A study in Kampala, Uganda, found evidence suggesting that people living with HIV may have

stockpiled ART tablets from previous prescriptions, which allowed them to keep taking their medication even when they could not visit ART clinics during COVID lockdowns [36]. We have heard similar anecdotes in clinical care that people may be getting ART from multiple clinics and giving or sharing with friends who feel too stigmatized to go themselves. Future research could examine whether clients get ART from multiple clinics to stockpile or share, and look at changes in ART diversion patterns and associations with service delivery models following COVID.

Strengths of our study include the large, population-based sample for both men and women living with HIV across 41 study communities, and the availability of viral load biomarkers. Limitations include our reliance on self-report for ART diversion measures, which may introduce recall or social desirability bias. In particular, the fact that a few participants reported buying ART but none reported selling ART may suggest the presence of social desirability bias. However, participants in our prior qualitative study were comfortable discussing ART giving and receiving with interviewers [20], and previous work in this setting has demonstrated the accuracy of other self-reported ART-related measures [22]. Further, the presence of social desirability bias likely means that our estimates of ART diversion are potential underestimates of the true extent of these behaviours. While the RCCS is a population-based study, there may still be selection bias if non-participants are more likely to be viraemic and also to share ART; however, the direction of such bias is unclear. We may not have accounted for all potential confounders in our adjusted analyses. Our cross-sectional design cannot assess a causal relationship between ART diversion and viremia. Future longitudinal studies are needed to assess temporal relationships. Future studies could also examine associations between ART sharing and drug resistance.

5 | CONCLUSIONS

In summary, this first population-based assessment of ART diversion in Africa found that ART sharing is common, particularly among men. While receiving ART may support viral suppression, giving ART was associated with viremia. HIV programmes may benefit from considering drug sharing in counselling messaging.

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COMPETING INTERESTS

The authors have no competing interests to report.

AUTHORS' CONTRIBUTIONS

CEK, MKG, FN, SJR and LWC conceived and designed the study. In collaboration with MKG and CEK, JS and XF conducted statistical analysis and directly accessed

and verified the underlying data reported in the manuscript. RS, GK and JK over-saw data collection. All authors had full access to the data in the study, participated in the interpretation of data and revising the manuscript, and had final responsibility for the decision to submit for publication.

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DATA AVAILABILITY STATEMENT

A de-identified version of the Rakai Community Cohort Study data can be provided to interested parties subject to the completion of the Rakai Health Sciences Program data request form and signing of a Data Transfer Agreement. Inquiries should be directed to datarequests@rhsp.org.

REFERENCES

- 1. Berge KH, Dillon KR, Sikkink KM, Taylor TK, Lanier WL. Diversion of drugs within health care facilities, a multiple-victim crime: patterns of diversion, scope, consequences, detection, and prevention. Mayo Clin Proc. 2012;87(7):674–82.
- 2. Beyene KA, Sheridan J, Aspden T. Prescription medication sharing: a systematic review of the literature. Am J Public Health. 2014;104(4):e15–26.
- 3. World Health Organization. Key facts and latest estimates on the global HIV epidemic 2021. 2022. https://cdn.who.int/media/docs/default-source/hq-hiv-hepatitis-and-stis-library/key-facts-hiv-2021-26july2022.pdf?sfvrsn= 8f4e7c93. 5 (accessed 28 September 2022).
- 4. Tsuyuki K, Surratt HL, Levi-Minzi MA, O'Grady CL, Kurtz SP. The demand for antiretroviral drugs in the illicit marketplace: implications for HIV disease management among vulnerable populations. AIDS Behav. 2015;19(5):857–68.
- 5. Tsuyuki K, Surratt HL. Antiretroviral drug diversion links social vulnerability to poor medication adherence in substance abusing populations. AIDS Behav. 2015;19(5):869–81.
- 6. Surratt HL, O'Grady CL, Levi-Minzi MA, Kurtz SP. Medication adherence challenges among HIV positive substance abusers: the role of food and housing insecurity. AIDS Care. 2015;27(3):307–14.
- 7. Surratt HL, Kurtz SP, Levi-Minzi MA, Chen M. Environmental influences on HIV medication adherence: the role of neighborhood disorder. Am J Public Health. 2015;105(8):1660–6.
- 8. Surratt HL, Kurtz SP, Cicero TJ, O'Grady C, Levi-Minzi MA. Antiretroviral medication diversion among HIV-positive substance abusers in South Florida. Am J Public Health. 2013;103(6):1026–8.
- 9. O'Grady CL, Kurtz SP, Surratt HL. Antiretroviral medication diversion in South Florida: prescription types and motivations. Drug Alcohol Depend. 2014:140:e164.
- 10. Kurtz SP, Buttram ME, Surratt HL. Vulnerable infected populations and street markets for ARVs: potential implications for PrEP rollout in the USA. AIDS Care. 2014;26(4):411–5.
- 11. Chibi B, Torres NF, Sokhela ZP, Mashamba-Thompson TP. Risk factors for prescription drug diversion among people living with HIV: a systematic scoping review. Int J Evid Based Healthc. 2020;18(1):5-23.
- 12. Buttram ME, Kurtz SP. Preliminary evidence of HIV seroconversion among HIV-negative men who have sex with men taking non-prescribed antiretroviral medication for HIV prevention in Miami, Florida, USA. Sex Health. 2016;14(2):193–5.
- 13. Chinuoya M, Rikhotso R, Ngunyulu RN, Peu MD, Mataboge MLS, Mulaudzi F, et al. 'Some mix it with other things to smoke': perceived use and misuse of ARV by street thugs in Tshwane District, South Africa. Afr J Phys Health Edu Recreat Dance. 2014;Supplement 1(1):113–26.

- 14. Grelotti DJ, Closson EF, Smit JA, Mabude Z, Matthews LT, Safren SA, et al. Whoonga: potential recreational use of HIV antiretroviral medication in South Africa. AIDS Behav. 2014;18(3):511–8.
- 15. Larkan F, Wyk BV, Saris J. Of remedies and poisons: recreational use of antiretroviral drugs in the social imagination of South African carers. Afr Sociol Review. 2010;14(2):62–73.
- 16. Magidson JF, Iyer HS, Regenauer KS, Grelotti DJ, Dietrich JJ, Courtney I, et al. Recreational ART use among individuals living with HIV/AIDS in South Africa: examining longitudinal ART initiation and viral suppression. Drug Alcohol Depend. 2019;198:192–8.
- 17. Rough K, Dietrich J, Essien T, Grelotti DJ, Bansberg DR, Gray G, et al. Whoonga and the abuse and diversion of antiretrovirals in Soweto, South Africa. AIDS Behav. 2014;18(7):1378–80.
- 18. Rock A, Comins C, Mulumba N, Young K, Mcingana M, Guddera V, et al. Antiretroviral treatment sharing among female sex workers living with HIV in eThekwini (Durban), South Africa: drivers and implications for treatment success. J Int Assoc Provid AIDS Care. 2022;21.
- 19. Parmley LE, Comins CA, Young K, , Mcingana M, Phetlhu DR, Guddera V, et al. Occupational barriers to accessing and adhering to antiretroviral therapy for female sex workers living with HIV in South Africa. Occup Environ Med. 2020;77(2):100–6.
- 20. Rosen JG, Nakyanjo N, Isabirye D, Wawer MJ, Nalugoda F, Reynolds SJ, et al. Antiretroviral treatment sharing among highly mobile Ugandan fisherfolk living with HIV: a qualitative study. AIDS Care. 2020;32(7):912–5.
- 21. Chang LW, Grabowski MK, Ssekubugu R, Nalugoda F, Kigozi G, Nantume B, et al. Heterogeneity of the HIV epidemic in agrarian, trading, and fishing communities in Rakai, Uganda: an observational epidemiological study. Lancet HIV. 2016;3(8):e388–96.
- 22. Grabowski MK, Reynolds SJ, Kagaayi J, Clarke W, Chang LC, Nakigozi G, et al. The validity of self-reported antiretroviral use in persons living with HIV: a population-based study. AIDS. 2018;32(3):363–9.
- 23. World Health Organization. Consolidated guidelines on HIV prevention, testing, treatment, service delivery and monitoring: recommendations for a public health approach, 2021.
- 24. Joint United Nations Programme on HIV/AIDS. Male engagement in HIV testing, treatment and prevention in eastern and southern Africa a framework for action 2022
- 25. Sileo KM, Fielding-Miller R, Dworkin SL, Fleming PJ. A scoping review on the role of masculine norms in men's engagement in the HIV care continuum in sub-Saharan Africa. AIDS Care. 2019;31(11):1435–46.
- 26. Lubega M, Nakyaanjo N, Nansubuga S, Hiire E, Kigozi G, Nakigozi G, et al. Risk denial and socio-economic factors related to high HIV transmission in a fishing community in Rakai, Uganda: a qualitative study. PLoS One. 2015;10(8):e0132740.
- 27. Lubega M, Nakyaanjo N, Nansubuga S, Hiire E, Kigozi G, Nakigozi G, et al. Understanding the socio-structural context of high HIV transmission in Kasensero

- fishing community, South Western Uganda. BMC Public Health. 2015;15: 1033
- 28. Obol JH, Akera P, Atim PO, Awor S, Wanyama R, Moi KL, et al. Prevalence of borrowing and sharing prescription medicines and associated socio-demographic factors: findings from COBERS health centres in northern Uganda. BMC Pharmacol Toxicol. 2018;19(1):17.
- 29. Wamoyi J, Heise L, Meiksin R, Kyegombe N, Nyato D, Buller AM. Is transactional sex exploitative? A social norms perspective, with implications for interventions with adolescent girls and young women in Tanzania. PLoS One. 2019;14(4):e0214366.
- 30. Inzaule SC, Hamers RL, Doherty M, Shafer RW, Bertagnolio S, de Wit TFR. Curbing the rise of HIV drug resistance in low-income and middle-income countries: the role of dolutegravir-containing regimens. Lancet Infect Dis. 2019;19(7):e246–52.
- 31. van Oosterhout JJ, Chipungu C, Nkhoma L, Kanise H, Hosseinipour MC, Sagno JB, et al. Dolutegravir resistance in Malawi's National HIV Treatment Program. Open Forum Infect Dis. 2022;9(5).
- 32. Mansergh G, Mayer K, Hirshfield S, Stephenson R, Sullivan P. HIV pre-exposure prophylaxis medication sharing among HIV-negative men who have sex with men. JAMA Netw Open. 2020;3(9):e2016256.
- 33. Moodley J, Naidoo S, Moodley J, Ramjee G. Sharing of investigational drug among participants in the voice trial. AIDS Behav. 2016;20(11):2709-14.
- 34. Thomson KA, Haberer JE, Marzinke MA, Mujugira A, Hendrix CW, Celum C, et al. Brief Report: medication sharing is rare among African HIV-1 serodiscordant couples enrolled in an efficacy trial of oral pre-exposure prophylaxis (PrEP) for HIV-1 prevention. J Acquir Immune Defic Syndr. 2017;75(2):184–9.
- 35. Ssempijja V, Ndyanabo A, Ssekubugu R, Chang L, Nakawooya H, Nalugoda F, et al. Population trends in HIV service delivery, viral suppression, and incidence before and during the COVID-19 era in Rakai, Uganda. International AIDS Society. Montreal, Canada, 2022.
- 36. Wagner Z, Mukasa B, Nakakande J, Stecher C, Saya U, Linnemayr S. Impact of the COVID-19 pandemic on use of HIV care, antiretroviral therapy adherence, and viral suppression: an observational cohort study from Uganda. J Acquir Immune Defic Syndr. 2021;88(5):448–56.

SUPPORTING INFORMATION

Additional information may be found under the Supporting Information tab for this article:

Supplemental Table. Sociodemographic factors and sexual behaviors associated with giving, receiving, or buying ART in the past 12 months among persons living with HIV self-reporting ART use in south-central Uganda (N = 2852)