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# Behavioral and Other Characteristics Associated with HIV Viral Load in an Outpatient Clinic 

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#### Abstract

Persons living with HIV (PLWH) who are engaged in care, yet not virally suppressed, represent a risk for transmission and opportunity for risk reduction interventions. This study describes characteristics of an outpatient clinic cohort of PLWH by laboratory confirmed viral suppression status and examines associations with demographics and sexual and drug use behaviors gathered through questionnaire. From a sample of 500 clinic patients, 438 were prescribed antiretroviral treatment (ART) and 62 were not. Among the 438 on ART, 72 (16.4\%) were not virally suppressed at the most recent lab draw. Compared to individuals with a suppressed viral load, those that were unsuppressed were more likely to: be black ( $79.2 \%$ vs. $64.2 \%$; $p=0.014$ ); earn below $\$ 25,000$ year ( $88.9 \%$ vs. $65.0 \%$; $p<$ 0.001 ); be of a younger age ( 47.8 vs . 50.0 mean years; $p=0.009$ ); be on opiate substitution (14.1\% vs. $6.3 \% ; p=0.023$ ); and acknowledge poly-substance (38.9\% vs. 24.4\%; $p=$ 0.012 ) and excessive alcohol use ( $13.9 \%$ vs. $6.0 \%$; $p=0.019$ ). Conversely, a smaller proportion of those with an unsuppressed viral load had multiple sex partners in the previous 30 days ( $39.8 \%$ vs. $58.5 \% ; p=0.003$ ). In multivariable regression of those on ART, the prevalence of an unsuppressed viral load was $3 \%$ lower with each increasing year of age (aPR: 0.97; 95\% CI: $0.95,0.99$ ) and $47 \%$ lower with income over $\$ 25,000 /$ year (aPR: 0.33 ; $95 \% \mathrm{CI}: 0.16,0.70$ ). In a separate analysis of all 500 subjects, ART was less frequently prescribed to blacks compared to whites, heterosexuals, those with lower education and income, and persons with active substance use. Findings confirm that a large proportion of PLWH and engaged in care were not virally suppressed and continued behaviors that risk transmission, indicating the need for screening, prevention counseling and access to ancillary services to lower the incidence of HIV infections.


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## Introduction

In the United States, $4 \%$ of persons living with HIV (PLWH) diagnosed and retained in care have not been prescribed antiretroviral treatment (ART) and are not virally suppressed, while $10 \%$ on ART have not achieved viral suppression.[1] Risky sexual behavior with a person who has an unsuppressed viral load accounts for an estimated 28.9 new infections per 100 persons. [2] Although transmission from injection drug use has declined, substance use is associated with sexual behaviors that risk transmission and may also impair adherence to ART. The purpose of this study is to describe a clinic cohort of PLWH and examine associations with viral suppression with a focus on drug use and sexual risk behaviors. Study findings can inform prevention efforts to reduce risk behaviors among PLWH with unsuppressed viral load seen in the clinical setting.

Substance use is a key factor for high-risk sexual behavior.[3-5] Nationally, $17 \%$ of heterosexual women, $16 \%$ of men who have sex with men (MSM) and $11 \%$ of heterosexual men living with HIV have an unsuppressed viral load concurrent with a sexual or drug use risk behavior. [2] Alcohol and crack or powder cocaine use have been related to multiple concurrent sex partners among heterosexuals and MSM,[6] and heroin use with fewer partners.[7] Inconsistent condom use occurs with alcohol $[6,8,9]$ and injection drug use.[10]

Limited access to treatment and low adherence to ART may be a consequence of substance use, but for those who do adhere to treatment virologic outcomes are comparable to those who do not use substances.[11] Use of stimulants, including methamphetamine [12] crack [12, 13] and cocaine [12, 14] as well as alcohol [13,15] but not heroin [14] have been associated with poorer adherence to ART compared to persons who do not use substances. Persons who actively inject drugs have had lower rates of viral suppression at ART initiation but not with continued treatment, possibly due to varying intensity of injection over time, including abstinence, as well as linkages with support services that can improve adherence for those engaged in care.[16]

## Methods

This is a secondary analysis of a cross-sectional sample of 500 PLWH from a study evaluating risk factors for methicillin-resistant Staphylococcus aureus colonization.[17] Subject recruitment, interviews and chart abstraction occurred between March 2010 and April 2011. The Johns Hopkins Medicine Institutional Review Board approved the study and consent procedure and all participants provided written informed consent that was recorded on a signed consent form.

## Study population

The Baltimore, Maryland, metropolitan area has the third highest prevalence of PLWH in the US.[18] Similar to national trends, the proportion of incident cases is highest in the MSM risk group (50\%), followed by heterosexual (37\%) and persons who inject drugs (PWID) (10\%).[19]

The Johns Hopkins University AIDS Service (JHUAS) has two clinics in the Baltimore area. Over 75\% of JHUAS clients receive care at the Moore Clinic for HIV at the Johns Hopkins Hospital medical campus in the East Baltimore community. The Moore Clinic provides many co-located ancillary services, including opiate substitution, counseling and mental health care, and medication support. The Moore Clinic follows an average of 2,300 mostly African-American ( $77 \%$ ) clients annually, including the uninsured with a majority having HIV transmission risks of intravenous drug use and heterosexual sex. The Green Spring Station (GSS) clinic is located in Baltimore County, serves about 650, primarily white (64\%) and exclusively insured clients, with MSM as the key affected population.

Eligible participants were current JHUAS clients with two or more visits within the previous 12 months, were 18 years of age and older and able to understand spoken English and provide consent. Clients at each clinic were approached and screened consecutively at a routine or acute care visit unrelated to the study during the recruitment period. Interviewers approached all patients at GSS and every $10^{\text {th }}$ patient entering the Moore Clinic because of its much larger clinic population. A $\$ 25$ gift card was offered for study participation.

## Behavioral assessment

Study team members administered the questionnaire (S1 Appendix) during face-to-face interviews onsite following clients' appointments with their providers. Fourteen questions developed from previously published studies [20,21] focused on characterization of sexual and substance use behaviors. Given the sensitive nature of the information, interviews took place in private consultation rooms and it was stressed that all responses were confidential. Interviewers were registered nurses or nursing students trained in use of the tool as well as sexual history taking. The time frame for any sexual or drug use behaviors was the 12 months prior to the interview. Research nurses reviewed patient medical charts to confirm demographic information and obtain medical history and viral load results from the most recent lab visit. Viral suppression was defined as viral load below detectable limits of 50 HIV-1 RNA copies/mL blood.

## Statistical analysis

The primary analysis examined associations between participant characteristics and behaviors with having an unsuppressed viral load among those prescribed ART. Given the high prevalence of unsuppressed viral load in the cohort, prevalence ratios were determined through a generalized linear model with log link, Poisson distribution and robust standard errors. Patterns of ART prescription were examined comparing persons prescribed and not prescribed ART in the full cohort. Interactions between drug use and sexual behaviors and both risk behaviors with demographic characteristics were tested. Variables with a significance of $p$ value of 0.05 or less in bivariate analyses were included in multivariable regression models, except for use of individual substances to avoid collinearity given the high levels of poly-substance use. Condom use was retained regardless of statistical significance given its importance to preventing HIV transmission. All tests were two-sided and a $p$ value of 0.05 or less was considered statistically significant. The data analysis was conducted using STATA software version 14 (Stata Corp., College Station, Texas, USA).

## Results

There were 500 participants recruited for the study, 150 from GSS and 350 from the Moore Clinic. There were 438 subjects on ART, of which 366 (83.6\%) were virally suppressed and 72 (16.4\%) were unsuppressed (Table 1). The median viral load was 8,663 (IQR 250-38,380). Individuals with an unsuppressed viral load were more likely than those with a suppressed viral load to be younger (mean age 47.8 vs. 50.9 years; $p=0.009$ ), black compared to other races ( $79.2 \%$ vs. $64.2 \% ; p=0.014$ ) and earn less than $\$ 25,000$ annually ( $88.9 \%$ vs. $65.0 \% ; p<0.001$ ). Among the unsuppressed, sexual risk was lower, with fewer having multiple sex partners in the previous thirty days ( $39.8 \%$ vs. $58.5 \% ; p=0.003$ ), while substance use was greater, with more using multiple substances ( $38.9 \%$ vs. $24.4 \% ; p=0.012$ ), excessive alcohol ( $13.9 \%$ vs. $6.0 \%$; $p=0.019$ ) and benzodiazepines ( $16.7 \%$ vs. $8.5 \% ; p=0.033$ ). Benzodiazepines were the most commonly used substance in combination with others. The unsuppressed were also more likely to be taking opiate substitution therapy ( $14.1 \%$ vs. $6.3 \% ; p=0.023$ ).

Table 1. Characteristics by viral suppression among persons prescribed ART.

|  |  | Suppressed | Unsuppressed Viral | $p$-value |
| :---: | :---: | :---: | :---: | :---: |
|  | Overall $\mathrm{N}=438$ | Viral Load N = 366 | Load $\mathbf{N = 7 2}$ |  |
|  | N (\%) | N (\%) | N (\%) |  |
| Age, mean years (range) | 50.4 (22.7-80.9) | 50.9 (25.8-80.9) | 47.8 (22.7-63.4) | 0.009 |
| Sex |  |  |  |  |
| Female | 143 (32.6) | 121 (33.1) | 22 (30.6) | 0.679 |
| Male | 295 (67.4) | 245 (66.9) | 50 (69.4) |  |
| Race |  |  |  |  |
| White/other ${ }^{\text {a }}$ | 146 (33.3) | 131 (35.8) | 15 (20.8) | 0.014 |
| Black | 292 (66.7) | 235 (64.2) | 57 (79.2) |  |
| Education |  |  |  |  |
| High school or less | 229 (52.3) | 179 (48.9) | 50 (69.4) | 0.001 |
| Any college | 209 (47.7) | 187 (51.1) | 22 (30.6) |  |
| Sexual orientation |  |  |  |  |
| Heterosexual | 288 (65.7) | 235 (64.2) | 53 (73.6) | 0.124 |
| Gay/lesbian/bisexual | 150 (34.3) | 131 (35.8) | 19 (26.4) |  |
| Annual income |  |  |  |  |
| < \$25,000 | 302 (69.0) | 238 (65.0) | 64 (88.9) | < 0.001 |
| $\geq$ \$25,000 | 136 (31.0) | 128 (35.0) | 8 (11.1) |  |
| CD4+ T cell count |  |  |  |  |
| <200 | 45 (10.3) | 26 (7.1) | 19 (26.4) | <0.001 |
| 200-349 | 81 (18.5) | 60 (16.4) | 21 (29.2) |  |
| 350-500 | 96 (21.9) | 76 (20.8) | 20 (27.8) |  |
| > 500 | 216 (49.3) | 204 (55.7) | 12 (16.7) |  |
| Sexually active |  |  |  |  |
| No | 147 (35.6) | 124 (33.9) | 23 (31.9) | 0.751 |
| Yes | 291 (66.4) | 242 (66.1) | 49 (68.1) |  |
| Condom use |  |  |  |  |
| Inconsistent/never | 152 (52.2) | 125 (51.7) | 27 (55.1) | 0.659 |
| Always | 139 (47.8) | 117 (48.3) | 22 (44.9) |  |
| $>1$ sex partner ${ }^{\text {b }}$ |  |  |  |  |
| No | 275 (46.4) | 103 (41.5) | 53 (60.2) | 0.003 |
| Yes | 29 (53.6) | 145 (58.5) | 35 (39.8) |  |
| Poly-substance use ${ }^{\text {c }}$ |  |  |  |  |
| No | 319 (73.2) | 275 (75.6) | 44 (61.1) | 0.012 |
| Yes | 117 (26.8) | 89 (24.4) | 28 (38.9) |  |
| Excessive alcohol ${ }^{\text {d }}$ |  |  |  |  |
| No | 405 (92.7) | 343 (94.0) | 62 (86.1) | 0.019 |
| Yes | 32 (7.3) | 22 (6.0) | 10 (13.9) |  |
| IV heroin and/or cocaine |  |  |  |  |
| No | 421 (96.1) | 353 (96.5) | 68 (94.4) | 0.421 |
| Yes | 17 (3.9) | 13 (3.5) | 4 (5.6) |  |
| Crack cocaine |  |  |  |  |
| No | 361 (82.6) | 303 (83.0) | 58 (80.6) | 0.615 |
| Yes | 76 (17.4) | 62 (17.0) | 14 (19.4) |  |
| Benzodiazepines |  |  |  |  |
| No | 394 (90.2) | 334 (91.5) | 60 (83.3) | 0.033 |
| Yes | 43 (9.8) | 31 (8.5) | 12 (16.7) |  |

(Continued)

Table 1. (Continued)

|  |  | Suppressed | Unsuppressed Viral | $p$-value |
| :---: | :---: | :---: | :---: | :---: |
|  | Overall $\mathrm{N}=438$ | Viral Load N = 366 | Load $\mathbf{N = 7 2}$ |  |
|  | N (\%) | N (\%) | N (\%) |  |
| Opiate substitution |  |  |  |  |
| No | 404 (92.5) | 343 (93.7) | 61 (85.9) | 0.023 |
| Yes | 33 (7.5) | 23 (6.3) | 10 (14.1) |  |

Suppressed below 40 copies HIV-1 RNA particles from most recent laboratory results within the prior 12 months.
${ }^{\text {a }}$ Latino/Hispanic: 8 (1.6\%); Asian/Pacific Islander: 2 (0.4\%); Native American: 3 (0.6\%).
${ }^{\text {b }}$ Previous 30 days.
${ }^{c}$ Any marijuana; crack cocaine; prescription pain medication; benzodiazepines; excessive alcohol; intranasal heroin or cocaine; IDU heroin or cocaine; ecstasy; smoked heroin; crystal (methamphetamine); poppers over prior 12 months.
${ }^{d}$ Excessive alcohol use was self-defined by the participant.
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Associations modeled with viral suppression are given for those taking ART in Table 2. In univariate regression, lower prevalence of an unsuppressed viral load was associated with each year of increasing age (PR: $0.97 ; 95 \%$ CI: $0.95,0.99$ ), any college (PR: $0.48 ; 95 \% \mathrm{CI}: 0.30,0.77$ ), annual income equal to or over $\$ 25,000 /$ year (PR: $0.28 ; 95 \% \mathrm{CI}: 0.14,0.56$ ) and with use of benzodiazepines (PR: 0.55 ; $95 \%$ CI: $0.32,0.93$ ). Greater prevalence of unsuppressed viral load was associated with black race (PR: 1.9; 95\% CI: 1.11, 3.24), poly-substance use (PR: 1.74; 95\% CI: $1.14,2.65$ ) and opiate substitution ( $2.01 ; 95 \%$ CI: 1.14, 3.54). In multivariable regression, only the lower prevalence of being unsuppressed with increasing age (aPR: $0.97 ; 95 \% \mathrm{CI}: 0.95,0.99$ ) and income equal to or over $\$ 25,000 /$ year (aPR: $0.33 ; 95 \% \mathrm{CI}: 0.16,0.70$ ) remained significant.

Among the full cohort of 500 participants, 62 were not prescribed ART. Patterns of ART prescription revealed greater proportions prescribed ART with any college compared to high school education ( $92.5 \%$ vs. $83.6 \% ; p=0.003$ ) and with incomes over $\$ 25,000 /$ year ( $92.5 \%$ vs. $85.6 \% ; p=0.031)$. Prescription of ART was lower among blacks compared to other races ( $84.6 \%$ vs. $94.2 \% ; p=0.003$ ), persons who use multiple substances ( $81.3 \%$ vs. $90.1 \% ; p=0.007$ ), injection drugs ( $65.4 \%$ vs. $88.8 \%$; p $<0.001$ ), excessive alcohol ( $74.4 \%$ vs. $88.8 \% ; p=0.006$ ) and crack cocaine ( $78.4 \%$ vs. $89.8 \% ; p=0.002$ ), as well as among heterosexuals compared to gay/ lesbian/bisexuals ( $85.2 \%$ vs. $92.6 \% ; p=0.019$ ). The differences between sexual orientation may be related to socioeconomic characteristics, with persons identifying as gay/lesbian/bisexual more likely to have some college education ( $75.9 \%$ vs. $30.5 \% ; p<0.001$ ) and earn more than $\$ 25,000$ annually ( $58.6 \%$ vs. $15.4 \%$; $p<0.001$ ).

## Discussion

While sex and drug use behaviors were not associated with an unsuppressed viral load, they did occur in the presence of viremia and consequently represent a risk for transmission. Compared to a nationally representative survey of PLWH receiving outpatient care in the United States, [22] the study population had greater transmission risk behaviors, with more reporting unprotected sex ( $37.8 \%$ vs. $23 \%$ ), crack cocaine use ( $38.3 \%$ vs. $4.3 \%$ ), and any non-injection drug ( $37.6 \%$ vs. $26.6 \%$ ) or intravenous drug use ( $10.2 \%$ vs. $2 \%$ ). Our study also had fewer with a high school or higher education ( $45.2 \%$ vs. $53.2 \%$ ) and a greater proportion of black race ( $69 \%$ vs. $41.7 \%$ ), indicating a more socioeconomically disadvantaged group.

Guidelines to initiate ART in treatment naïve patients at the time of the study recommended starting treatment when CD4+ T cell counts declined below 500 cells $/ \mathrm{mm} 3$ or there was a history of AIDS.[23] In practice, many JHUAS providers were offering ART to all

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Table 2. Univariate and multivariable regression results for the outcome of having an unsuppressed viral load for those taking ART.

| Variable | Univariate regression |  | Multivariable regression |  |
| :---: | :---: | :---: | :---: | :---: |
|  | PR (95\% CI) | $p$-value | PR ratio (95\% CI) | $p$-value |
| Sexual orientation |  |  |  |  |
| Gay/lesbian/bisexual | 1.00 | 0.132 |  |  |
| Heterosexual | 1.45 (0.89, 2.36) |  |  |  |
| Age (years) | 0.97 (0.95, 0.99) | 0.010 | 0.97 (0.95, 0.99) | 0.009 |
| Sex |  |  |  |  |
| Female | 1.00 |  |  |  |
| Male | 0.91 (0.57, 1.44) | 0.680 |  |  |
| Race |  |  |  |  |
| White/other | 1.00 | 0.018 | 1.00 | 0.228 |
| Black | $1.9(1.11,3.24)$ |  | 1.40 (0.81, 2.41) |  |
| Education |  |  |  |  |
| High school or less | 1.00 | 0.002 |  |  |
| Any college | 0.48 (0.30, 0.77) |  |  |  |
| Income |  |  |  |  |
| < \$25,000/year | 1.00 | <0.001 | 1.00 | 0.004 |
| $\geq$ \$25,000/year | 0.28 (0.14, 0.56) |  | 0.33 (0.16, 0.70) |  |
| Sexually active ${ }^{\text {a }}$ | 0.93 (0.59, 1.46) | 0.751 |  |  |
| $\geq 1$ sex partner ${ }^{\text {b }}$ | 0.75 (0.29, 1.92) | 0.543 |  |  |
| Condom use |  |  |  |  |
| Always | 1.00 | 0.445 | 1.00 | 0.356 |
| Inconsistent | 0.85 (0.56, 1.29) |  | 0.82 (0.54, 1.24) |  |
| Poly-substance use | 1.74 (1.14, 2.65) | 0.011 | 1.18 (0.75, 1.88) | 0.470 |
| Injection drugs ${ }^{\text {c }}$ | 1.46 (0.60, 3.53) | 0.405 |  |  |
| Excessive alcohol ${ }^{\text {d }}$ | 0.49 (0.28, 0.86) | 0.013 | 1.21 (0.67, 2.19) | 0.528 |
| Opiate substitution | 2.01 (1.14, 3.54) | 0.016 |  |  |
| Benzodiazepines | 0.55 (0.32, 0.93) | 0.026 |  |  |
| Crack cocaine | 0.87 (0.51, 1.48) | 0.613 |  |  |

Cl , confidence interval; PR, prevalence ratio
Note: Bold text denotes statistically significant findings at $p \leq 0.05$.
${ }^{\text {a }}$ Previous 12 months.
${ }^{\mathrm{b}}$ Previous 30 days.
${ }^{\text {c }}$ Heroin, cocaine or both.
${ }^{d}$ Excessive alcohol use was self-defined by the participant.
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patients, even those with CD4+ T cell levels over 500, and some patients not on ART during the study period had been previously, but did not continue due to low medication adherence. Prior treatment adherence and provider prescribing preferences were not available and consequently it was not possible to distinguish how these factors may have impacted the decision not to treat.

It was also found that ART was less likely to be prescribed to persons of black race, having high school or less education and reporting substance use. Abstinence or participation in mental health and substance use treatment services supports adherence [24] and indicates the importance of screening for psychosocial needs and referral to ancillary care. Although substance use was common in this cohort, it was not found to be associated with viral suppression in the adjusted model, suggesting that patients using substances can be adherent and bias
against offering treatment due to substance use behaviors should be avoided. With younger age and lower income related to an unsuppressed viral load, adherence counseling should be culturally sensitive and educationally appropriate for youth, and barriers to accessing care such as transportation needs and unstable housing should be addressed.

Although just $40 \%$ of the 1.2 million PLWH in the U.S. are engaged in care,[1] they represent a substantial opportunity to implement risk reduction interventions given that many are not virally suppressed.[22] Interventions to change drug use and sexual risk behaviors can produce significant and lasting reductions in risk behaviors and improve ART adherence.[25-27] Yet among PLWH receiving regular care, prevention counseling to reduce transmission from sexual and substance use behaviors is provided half or less of the time.[28, 29] With high proportions engaging in high-risk drug use and sexual behaviors, our study confirms the need for these services for PLWH seen in the clinical setting. However, risk behavior(s) or viremia alone are necessary but not individually sufficient for transmission to occur, and consequently must be considered together when evaluating transmission risks. [2,30]

While many HIV specialty clinics provide co-located mental health, drug treatment and other supportive services, [31] low rates of screening, counseling and referrals by clinicians remains a challenge.[28,29] Screening for risk behaviors and providing prevention messages are limited by time constraints and competing clinical priorities even where these interventions are the standard of care for a clinical practice. Additionally, with healthcare reform and the move away from specialty HIV to general primary care clinics, patients face new challenges in navigating the healthcare system to connect with ancillary services.[32]

This study has described the characteristics of clients with an unsuppressed viral load and considered patterns of ART prescription. Limitations included the use of self-reported risk behaviors that are subject to social desirability and recall bias. Important factors to consider in future studies include length of time since diagnosis, stage of HIV disease at diagnosis, duration of ART, and HIV or treatment status of stable partners, which were not determined given the degree of missing data for these particular characteristics and inability to confirm reliability of the information that was available.

## Conclusion

A proportion of persons in HIV medical care on ART were not virally suppressed and continued to participate in transmission risk behaviors, indicating the need for screening, prevention counseling and access to ancillary services, particularly for substance use and safer sex counseling. Substance use itself was not a risk factor for unsuppressed viral load, indicating persons actively using can be adherent to treatment.

## Supporting Information

S1 Appendix. Survey questionnaire instrument from the primary study. (DOC)

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## Author Contributions

## Conceptualization: PLS JEF.

Formal analysis: PLS.
Funding acquisition: JEF.
Methodology: PLS JEF.
Validation: PLS.
Writing - original draft: PLS JEF.
Writing - review \& editing: PLS JEF.

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