Attitudes and Barriers towards Pre-Exposure Prophylaxis (PrEP) among High-Risk HIV-Seronegative Men who Have Sex with Men

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Abstract

Objective: Recent studies have demonstrated the efficacy of antiretroviral preexposure prophylaxis (PrEP) for prevention of HIV-1 infection. The purpose of this study was to identify barriers to PrEP use among high HIV-risk men who have sex with men in Denver, Colorado.

Methods: A 19 question Likert-scale survey was used to assess knowledge of PrEP; attitudes towards cost, side effects, and alternative prevention methods; and the effects of PrEP on sexual behaviors and practices. The survey was administered at study entry and six months later to HIV seronegative, men who have sex with men and male-to-female transgendered women who have sex with men and participate in high-risk sexual behavior.

Results: Between June and September 2013, 65 participants from the metropolitan Denver area completed the survey. Median age was 36 years (range: 20-52 years); 88% were white, 6% Hispanic, and 1.5% African American; 65% had at least a college degree or higher and 27% had more than high school education. The reported number of sexual partners in the past six months ranged from zero to 150 partners (median 3 partners) and 75% reported condom use during all sexual encounters. Although 72.3% reported prior knowledge of PrEP only five participants (7.7%) reported ever using PrEP in the past. Participants were most likely (93.8%) to use PrEP in the future if they were in a monogamous relationship with an HIV-infected partner and least likely to use PrEP if it required out-of-pocket costs (10.7%). Younger age was associated with decreased odds of future PrEP use even if PrEP was provided free of charge (OR 0.2, 95% CI 0.1, 1.0). Higher number of sexual partners in the preceding six months was associated with decreased odds of using condoms if taking PrEP in the future (aOR 0.2; 95% CI 0.1, 0.8). No changes in survey responses were noted between baseline and six months.

Conclusions: PrEP usage was uncommon among men at high risk for sexual acquisition of HIV infection and cost of antiretroviral drugs was a major barrier to future PrEP use. If PrEP is to have major impact on transmission of HIV-1, expanded efforts to decrease cost and increase community awareness of PrEP safety and efficacy are needed.

Keywords: Pre-exposure prophylaxis; PrEP; HIV; Vaccine; Prevention; HVTN 505; Barriers; Sexual behaviors

Introduction

Recent studies have demonstrated the protective effects of emtricitabine/tenofovir-DF as pre-exposure chemoprophylaxis (PrEP) for patients at high risk of HIV-1 infection [1,2]. The iPrex study demonstrated that co-formulated emtricitabine/tenofovir-DF reduced the incidence of HIV infection by 44% among seronegative male-to-female transgendered women or men who have sex with men (MSM) [1]. More recently, the Partners PrEP Trial demonstrated proof from HIV transmission among serodiscordant heterosexual couples in Uganda and Kenya, with a 67% relative reduction of HIV incidence with tenofovir-DF alone and 75% with emtricitabine/tenofovir-DF when compared to placebo [2]. Other trials have demonstrated similar findings; however efficacy in HIV infection rate reduction is highly dependent on adherence to the antiretroviral regimen [3,4]. Although emtricitabine/tenofovir-DF has a favorable safety profile, side effects such as nausea, renal toxicity, bone density loss, and liver toxicity can limit use [5].

Little information is available on the factors that limit PrEP use in community settings. Several small studies have demonstrated moderate interest in taking PrEP among at-risk populations, if it were available. There is modest correlation between risk behavior and preference for PrEP, and often little knowledge about PrEP [6-10]. Little is known about how other factors such as medication cost, issues affecting daily compliance, HIV education, and prevention counselling affect PrEP use. The present study evaluated attitudes and potential barriers to PrEP use among men who have sex with men or transgender women and are participants in a clinical trial of a candidate HIV vaccine. PrEP’s high cost, side effect profile, and individual sexual behaviour were hypothesized to be the greatest stated barriers for PrEP use in this setting.

Methods

Study population

Participants were recruited from those taking part in the HVTN...
505 vaccine trial at the University of Colorado Hospital Clinical Research Site, Aurora, Colorado. Complete eligibility criteria for HVTN 505 participation have been published elsewhere [11]. Participants in HVTN 505 were HIV seronegative men who have sex with men and male-to-female transgendered women who have sex with men who had unprotected anal intercourse with one or more male or male-to-female transgender partners or anal intercourse with two or more male or male-to-female transgender partners in the past six months. HVTN 505 enrollment at the University of Colorado Hospital began in November 2010 and was halted in April 2013, after enrollment of 107 participants, following recommendations by an independent data and safety monitoring board (DSMB) which found that the candidate vaccine did not prevent HIV infection nor reduce viral load after HIV infection [11,12]. All HVTN 505 study participants at the University of Colorado Hospital site who remained HIV seronegative were eligible for the current study and were invited to complete a survey about attitudes and barriers toward PrEP usage.

**Recruitment**

Recruitment took place by telephone from June 2013 to September 2013. After an attempt to contact a participant was made, the date and outcome of the call was recorded. If no contact was made after three attempts, no further follow-up was undertaken. Once verbal informed consent was obtained using a standard script, a trained interviewer administered a brief survey by telephone and recorded participant responses on paper forms. Beginning January 2014, participants were contacted again to complete a six-month follow-up survey, using the same questionnaire. This study was approved by the Colorado Multiple Institutional Review Board (COMIRB) and followed the human experimentation guidelines of the US Department of Health and Human Services.

**Survey instrument**

The 19 question Likert-scale survey took appropriately 10-15 minutes to administer by telephone (Appendix 1). The survey included questions about demographics, awareness of PrEP, prior history of PrEP use, associated costs, perceived side effects, alternative methods of HIV prevention and individual sexual behavior and practices. Specific questions were adapted from prior studies of PrEP awareness [5,13,14]. Participants verbally rated their likelihood of taking PrEP under 14 specific circumstances or conditions using a five-point Likert scale rating (“Definitely” (1), “very likely” (2), “somewhat likely” (3), “not very likely” (4), or “definitely not” (5)) to identify barriers to PrEP use. Data were collected without personal identifiable information.

**Data analysis**

Survey data were collected and managed using REDCap electronic data capture tools hosted at the University of Colorado Denver [15]. All analysis was performed using SAS, version 9.4 (SAS Institute, Cary, NC, USA). Demographic information, including age, race/ethnicity, education, and sexual practices was collected and analyzed descriptively. Likert scale questions were grouped by Likert score and frequency of answers. For each survey question, likely responses were calculated as the combination of “definitely” (1), “very likely” (2), and “somewhat likely” (3), while unlikely responses were calculated as “not very likely” (4) and “definitely not” (5). For examining likelihood to take PrEP based upon participant characteristics, a stratified analysis using Mantel-Haenszel chi-square test was undertaken in order to adjust for potential confounding (age), Both unadjusted (age) and adjusted odds ratios with 95% confidence intervals were calculated. McNemar’s test for correlated proportions was used to evaluate differences of likely and unlikely survey responses between those who completed both the initial and the follow-up surveys. The p–values were based on two-tailed test results and p ≤ 0.05 was used to define statistical significance.

**Results**

**Study population**

Among 103 eligible persons who were invited to participate, 65 (63.1%) completed the survey. Reasons for not participating in the survey included no response to telephone or email messages (32 participants); declined participation (5 participants); and incarceration (1 participant). Among survey participants, median age was 36 years (range; 20-52 years); 88% were white, 6% Hispanic, and 1.5% African American; 65% had at least a college degree or higher and 27% had more than high school education. The reported number of sexual partners in the past six months ranged from zero to 150 partners (median 3 partners) and 75% reported condom use during all sexual encounters (Table 1). During the initial survey, 47 participants (72.3%) indicated prior knowledge of PrEP and 5 participants (7.7%) reported having used PrEP in the past.

**Likelihood of using PrEP in the Future**

Participants were most likely to use PrEP in the future if they were in a monogamous relationship with an HIV-infected partner (93.8%) or had casual sexual partners whose HIV status was unknown (89.1%). Eighty percent (80%) of participants would be likely to use PrEP if it was provided free of charge, whereas 89.3% of participants were unlikely to use PrEP if it cost money out-of-pocket, approximately $500, each month. Participants were also less likely to use PrEP if there was a more effective vaccine available (89.2%) and if there were associated side effects (70.8%; Figure 1).

Younger participants (<36 years) were less likely to take PrEP if it cost money out-of-pocket each month (OR 0.1; 95% CI 0.0, 1.0), if PrEP was provided free of charge (OR 0.2; 95% CI 0.1, 1.0) or if there were possible side effects (OR 0.3; 95% CI 0.1, 1.0). Higher number of partners (>5 partners) in the preceding six months was associated with decreased likelihood of anticipating use of condoms if taking

| Race/Ethnicity               | Number (n) | Percent (%) |
|-----------------------------|------------|-------------|
| White/Non-Hispanic          | 57         | 87.7        |
| White/Hispanic              | 4          | 6.2         |
| Black                       | 1          | 1.5         |
| Other                       | 3          | 4.6         |

| Education                   | Number (n) | Percent (%) |
|-----------------------------|------------|-------------|
| HS diploma or GED           | 5          | 7.7         |
| More than HS                | 18         | 27.7        |
| College graduate or higher  | 42         | 64.6        |
| Ever heard about PrEP       | 47         | 72.3        |
| Ever taken PrEP             | 5          | 7.7         |

| Sexual activity             | Number (n) | Percent (%) |
|-----------------------------|------------|-------------|
| Receptive “bottom” partner  | 11         | 16.9        |
| Insertive “top” partner     | 11         | 16.9        |
| Both                        | 39         | 60.0        |
| Neither                     | 4          | 6.2         |

| Age (in years)              | Median     | Range       |
|-----------------------------|------------|-------------|
| No. partners in past 6 months | 3          | 0 - 150    |
| Percentage (%) of condom use in past 6 months | 75         | 0 - 100    |

Table 1: Demographics of participants who completed the initial survey.
PrEP (aOR 0.2; 95% CI 0.1, 0.8). There were no associations between likelihood to use PrEP and level of education, prior knowledge of PrEP or number of sexual partners (Table 2). Fifty-four participants (83.1%) completed both the initial and six month follow-up surveys. There were no statistically significant changes in survey responses between the initial and follow-up surveys (data not shown).

**Discussion**

The present study sought to uncover perceived barriers to PrEP use among high-risk HIV-seronegative men. We found that prior PrEP use was uncommon and cost, side effect profile, and potential availability of an effective vaccine were found to be associated with decreased willingness to use PrEP in the future. Presumed PrEP use was found to have some influence on anticipated condom use in the future among those with greater than five partners in the past six months. Importantly, younger age was associated with decreased willingness to use PrEP regardless of costs or side effects. This finding is in agreement with a prior study which found that older age was associated with greater PrEP interest [16]. Since starting PrEP at an early age would result in a longer total duration of PrEP usage and greater cumulative exposure, decrease willingness to use PrEP in the younger age group could reflect concerns about potential toxicities over one’s lifetime. Additional studies are needed to better understand why younger sexually active men are less willing to consider PrEP usage and evaluate interventions to increase PrEP use in this high-HIV-risk group.

Few studies to date have examined multiple social, economic, and other factors influencing the intent to take PrEP among those at risk for HIV infection. Krakower, et al. found that shortly after the iPrEx study results were released awareness of PrEP was limited and usage was uncommon [16]. Fuchs, et al. previously examined interest in PrEP, financial barriers, and the effect of taking PrEP on participation in the HVTN 505 trial. They found that a minority of participants expressed interest to use PrEP, however that interest significantly increased if PrEP was provided free of charge [17]. Other survey studies have investigated awareness of PrEP among high-risk populations, and demonstrated widely varying levels of awareness across various risk groups with similar attitudes towards cost and side effects [6,7,9,18]. Similar to these studies, our study confirms a general awareness of PrEP, and concern for cost and side effects among potential PrEP users as well. Our study is unique in its examination of the influence of sexual practices and behaviour, personal relationships, and the effect of future PrEP usage on the use of other prevention methods such as condoms. Taken together, these findings confirm the relative lack of uptake of PrEP as a prevention strategy among high risk men and point toward realistic targets to increase PrEP usage for effective prevention of HIV transmission.

Responses from our initial survey demonstrated that those at risk of HIV might be more likely to take PrEP under certain circumstances, such as monogamous discordance or unknown status of numerous
Limitations of the present study include a relatively small sample size and a single site design, which potentially limits generalizability to other high-risk populations. Other limitations include bias by the interviewers in conducting phone surveys with preferential intonation or further explanations of questions. An in-person written survey could have avoided this potential interviewer bias. The survey question concerning prior PrEP use could have been interpreted as also including post-exposure prophylaxis, and thus our finding of 7.7% prior PrEP usage might be an over estimate. The survey question concerning out-of-pocket expenses assumed minimal coverage by third party providers (out-of-pocket costs of $500 per month) and we did not assess the potential impact of lesser out-of-pocket costs. Participants for the PrEP knowledge regarding self or anticipated basis, rather than daily as continuous prophylaxis. The barriers to this two-pronged approach, an as-needed prophylactic medication may bring greater risk than benefit.

Table 2: Analysis of participant characteristics and the likelihood to take PrEP.

| How likely would you be to take PrEP if… | Odds Ratios (95% confidence intervals) |
|----------------------------------------|----------------------------------------|
| (definitely/very likely/somewhat likely) | Age | Education | PrEP knowledge | Sexual risk |
| ...it was provided to you free of charge? | 1.0 (ref): 36 years [median] | 1.0 (ref): HS education or less | 1.0 (ref): no prior knowledge | 1.0 (ref): ≤5 partners, last 6 months |
| ...it cost you money out-of-pocket each month? | 0.7 (0.2, 2.6) | 0.7 (0.2, 3.2) | 0.4 (0.1, 3.5) | 0.4 (0.0, 3.7) |
| ...it had side effects? | 0.7 (0.2, 2.0) | 1.2 (0.4, 3.7) | 0.5 (0.2, 1.7) | 0.7 (0.2, 2.5) |
| ...it had to be taken every day, missing as few pills as possible? | 0.6 (0.2, 8.5) | 2.6 (0.8, 8.5) | 1.4 (0.4, 5.0) | 2.1 (0.4, 10.9) |
| ...this medication was 50% effective at preventing HIV infection? | 0.7 (0.2, 2.1) | 0.4 (0.1, 1.2) | 0.4 (0.1, 1.2) | 0.6 (0.2, 1.7) |
| ...there was an HIV vaccine available that was less effective than this medication treatment? | 0.6 (0.2, 7.8) | 2.6 (0.8, 8.5) | 1.4 (0.4, 5.0) | 2.1 (0.4, 10.9) |
| ...there was an HIV vaccine available that was more effective than this medication treatment? | 0.7 (0.2, 8.8) | 2.6 (0.8, 8.5) | 1.4 (0.4, 5.0) | 2.1 (0.4, 10.9) |
| ...you were in a monogamous relationship with a partner who was HIV positive? | 0.7 (0.2, 2.1) | 0.4 (0.1, 1.2) | 0.4 (0.1, 1.2) | 0.6 (0.2, 1.7) |
| ...you had casual sexual partners whose HIV status you didn’t know? | 0.7 (0.2, 2.1) | 0.4 (0.1, 1.2) | 0.4 (0.1, 1.2) | 0.6 (0.2, 1.7) |
| ...it is possible that taking this medication now may mean they become less effective as a treatment option should you become infected with HIV in the future? | 0.7 (0.2, 2.1) | 0.4 (0.1, 1.2) | 0.4 (0.1, 1.2) | 0.6 (0.2, 1.7) |
| ...to use condoms if you were taking PrEP? | 0.7 (0.2, 2.1) | 0.4 (0.1, 1.2) | 0.4 (0.1, 1.2) | 0.6 (0.2, 1.7) |
| ...to engage in less cautious sexual behavior, such as use condoms less or have more sexual partners? | 0.7 (0.2, 2.1) | 0.4 (0.1, 1.2) | 0.4 (0.1, 1.2) | 0.6 (0.2, 1.7) |

† Adjusted for age
based form of prevention. Regardless of how PrEP is delivered, if PrEP is to have major impact on transmission of HIV-1, expanded efforts to decrease cost and increase community awareness of PrEP safety and efficacy are needed.

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