# **Preventing HIV Transmission in Persons with HIV**

This is a PDF version of the following document:Module 5:Prevention of HIVLesson 2:Preventing HIV Transmission in Persons with HIV

You can always find the most up-to-date version of this document at <u>https://www.hiv.uw.edu/go/prevention/prevention-positives/core-concept/all</u>.

## **Introduction and Background**

The availability of widespread effective antiretroviral therapy has transformed HIV from a fatal infection to a manageable chronic disease. Despite major advances in antiretroviral therapy, the incidence of new HIV infections in the United States continues to occur at a significant number, with about 32,000 to 36,000 persons newly acquiring HIV in recent years (Figure 1).[1] For persons living with HIV, antiretroviral therapy can dramatically reduce HIV transmission to others.[2,3,4,5] All clinicians providing HIV services should have knowledge and awareness of effective strategies that can play a role in reducing the risk of HIV transmission from persons with HIV. This topic review will focus on biomedical, behavioral, and structural measures related to preventing transmission from persons with HIV, a concept often referred to as Treatment as Prevention (TasP).[6]

# Antiretroviral Treatment as Prevention

## HIV RNA Level and Risk of HIV Transmission

A sentinel study in Rakai, Uganda, first reported HIV RNA levels tightly correlated with the risk of heterosexual HIV transmission.[7] This study involved 415 serodifferent heterosexual couples not taking antiretroviral therapy.[7] The main finding was that mean serum HIV RNA was significantly higher in subjects whose partners seroconverted than in subjects whose partners remained HIV negative, and HIV transmission was considered rare among persons who had an HIV RNA level less than 1,500 copies/mL (Figure 2).[7] This study played a major role in spurring on studies that would examine the impact that lowering HIV RNA levels with antiretroviral therapy would have on sexual transmission of HIV.

## Antiretroviral Therapy in HIV Serodifferent Couples

Convincing data from several studies have shown that antiretroviral treatment taken by individuals with HIV dramatically reduces HIV transmission to their sex partners.[3,4,8,9]

- **HPTN-052**: The concept of treating persons with HIV to prevent transmission to others, often referred to as treatment as prevention, garnered major support following the release of data from the landmark HPTN-052 trial.[3,4] The HPTN-052 trial was a randomized, controlled study that enrolled 1,763 HIV serodifferent, predominantly heterosexual couples from 9 countries. All persons with HIV had a CD4 count of 350 to 550 cells/mm<sup>3</sup> at enrollment, and none had HIV-related symptoms. The trial demonstrated that early initiation of antiretroviral therapy (started at the time of enrollment) reduced rates of sexual transmission of HIV to the partner without HIV by 96%, when compared with deferral of antiretroviral therapy (started when the CD4 count decreased to less than 250 cells/mm<sup>3</sup> or at the onset of an AIDS-related event). (Figure 3).[3,4]
- **Observational Trials**: Several observational studies with serodifferent heterosexual couples have similarly demonstrated that antiretroviral treatment of a partner with HIV is a potent intervention for preventing sexual transmission of HIV.[10] Further, data from a meta-analysis of 5,021 heterosexual serodifferent couples reinforces this finding; no HIV transmissions were reported from persons with HIV who were treated with antiretroviral therapy if the person with HIV had an HIV RNA level below 400 copies/mL.[9]
- **PARTNER-1 Study**: In the first phase of the European PARTNER (Partners of People on ART—A New Evaluation of the Risks) study, investigators at 75 sites in 14 European countries evaluated the impact of antiretroviral therapy on HIV transmission risk in 888 HIV-serodifferent couples engaging in condomless sex, including 548 heterosexual couples and 340 gay male couples.[2] The eligibility for enrollment required the partner with HIV to be taking antiretroviral therapy and have an HIV RNA level less than 200 copies/mL.[2] Enrollment took place from September 2010 through May 2014, and during this time, there were zero phylogenetically-linked HIV transmissions that occurred in these couples, with an estimated 58,000 condomless sex acts (22,000 in gay male couples and 36,000 in heterosexual couples).[2] There were 11 new HIV infections during the study period, but none of these were phylogenetically linked.
- **PARTNER-2 Study**: In the second phase of the European PARTNER (Partners of People on ART—A New Evaluation of the Risks) study, investigators at the same 75 sites in 14 European countries continued the PARTNER study, but additional enrollment was only for serodifferent gay men.[11] The enrollment period analyzed for PARTNER-2 was September 2010 through July 31, 2017, and included a total of 972 serodifferent gay male couples.[11] The design to include only serodifferent gay male couples was to account for some of the imbalance of the predominantly heterosexual serodifferent couples enrolled in the PARTNER-1 trial. As with PARTNER-1, the eligibility for enrollment required the partner with HIV to be taking antiretroviral therapy and have an HIV RNA level less than 200 copies/mL. During the study, the serodifferent gay male couples reported condomless anal sex a total of 76,088 times, and there were zero phylogenetically-linked transmissions.[11] There were, however, 15 new HIV infections during the study period, but none of these were phylogenetically linked.[11]

• **Opposites Attract**: The Opposites Attract trial was conducted from 2012 through 2016 and enrolled 358 HIV-serodifferent gay male couples in Thailand, Brazil, and Australia.[12] Of the 358 couples that enrolled, 343 had at least one follow-up visit, and 75% (258 of 343) of the partners with HIV had an HIV RNA level less than 200 copies/mL.[12] There were zero phylogenetically-linked cases of HIV transmission from among 12,447 sex acts that involved (1) condomless anal intercourse, (2) partners with HIV taking antiretroviral therapy (and with an HIV RNA less than 200 copies/mL), and (3) partners without HIV not taking HIV preexposure prophylaxis (PrEP).[12] There were three new HIV infections during the study period, but none were phylogenetically linked.[12]

## **Undetectable Equals Untransmittable (U=U)**

Extensive data from multiple studies strongly support the concept that persons with HIV who consistently take antiretroviral therapy and maintain undetectable HIV RNA levels do not transmit HIV sexually to others, even with condomless sex.[2,11,12] This concept is commonly referred to as Undetectable equals Untransmittable or U=U.[5] The U=U concept is an extremely important message that is now widely endorsed by prominent scientists, clinicians, organizations, and societies.[5] The findings from the studies that serve as the foundation for the U=U concept underscore the tremendous impact that antiretroviral treatment can have in preventing transmission of HIV from persons with HIV. These findings also emphasize the benefit of routine HIV testing and prompt initiation of antiretroviral therapy for persons who acquire HIV. For persons with HIV who are having condomless sex, it is important to perform regular screening for sexually transmitted infections, such as chlamydia, gonorrhea, and syphilis.

### **Recommendations for Antiretroviral Treatment as Prevention**

The Adult and Adolescent ART Guidelines cite prevention of sexual transmission of HIV as one of the main reasons to recommend antiretroviral therapy for all persons with HIV.[13] This recommendation to use antiretroviral treatment as prevention is now supported by multiple studies that persons who consistently maintain plasma HIV RNA levels less than 200 copies/mL do not transmit HIV to sex partners.[9,11,12,14] The Adult and Adolescent ART Guidelines also provide specific recommendations for the use of antiretroviral therapy to prevent sexual transmission of HIV (Treatment as Prevention); these recommendations clarify that persons with HIV who are starting antiretroviral therapy should use another form of HIV prevention with sex partners (e.g., condoms, preexposure prophylaxis for the HIV-negative sex partner, sexual abstinence) for at least the first 6 months of treatment and until they have achieved an HIV RNA level less than 200 copies/mL.[6] Note, these recommendations do not address use of antiretroviral therapy with achievement of undetectable plasma HIV RNA levels should decrease bloodborne transmission of HIV. The table below summarizes the Adult and Adolescent ART Guidelines recommendations for the use of **Table to Guidelines with HIV** 

#### Use of Antiretroviral Therapy to Prevent Sexual Transmission of HIV

- All persons with HIV should be informed that maintaining a plasma HIV RNA (viral load) of <200 copies/mL, including any measurable value below this threshold value, with antiretroviral therapy (ART) prevents sexual transmission of HIV to their partners. Patients may recognize this concept as Undetectable = Untransmittable or U=U (AII).
- Persons with HIV who are starting

ART should use another form of prevention with sexual partners (e.g. condoms, pre-exposure prophylaxis [PrEP] for the HIVnegative sexual partner, sexual abstinence) for at least the first 6 months of treatment and until a viral load of <200 copies/mL has been documented (AII). Many experts would recommend confirming sustained suppression before assuming that there is no further risk of sexual HIV transmission (AIII).

- When the viral load is ≥200 copies/mL, additional methods are needed to prevent transmission of HIV to sexual partners until resuppression to <200 copies/mL has been confirmed (AIII).
- Persons with HIV who intend to rely upon ART for prevention need to maintain high levels of ART adherence (AIII). They should be informed that transmission is possible during periods of poor adherence or treatment interruption (AIII).
- At each visit for HIV care, clinicians should assess adherence to ART and counsel patients regarding the importance of ART to their own health as well as its role in preventing sexual HIV transmission (AIII).
- Providers should inform patients that maintaining a viral load of <200 copies/mL does not prevent acquisition or transmission of other sexually transmitted infections (STIs) (AII).
- Providers should also routinely screen all sexually active persons with HIV for STIs, both for their own health and to prevent transmission of STIs to others (AIII).

Rating of Recommendations: A = Strong; B = Moderate; C = Optional

Rating of Evidence: I = Data from randomized controlled trials; II = Data from well-designed nonrandomized trials or observational cohort studies with long-term clinical outcomes; III = Expert opinion Source: • Panel on Antiretroviral Guidelines for Adults and Adolescents. Guidelines for the use of antiretroviral agents in adults and adolescents living with HIV. Department of Health and Human Services. Antiretroviral therapy to prevent sexual transmission of HIV (treatment as prevention). December 18, 2019. [HIV.gov]

## **Community Viral Load and Treatment Impact at Population Level**

Several groups have shown that a reduction in community viral load is associated with decreased numbers of new HIV infections in that community, supporting the hypothesis that wide-scale use of antiretroviral therapy in persons with HIV could significantly reduce HIV transmission at the population level. For example, in British Columbia, the scale-up of antiretroviral therapy from 1996 through 2009 led to a 52% decline in new HIV infections.[15] A separate study performed in San Francisco found that reductions in mean and total community viral load from 2004 through 2008 were associated with a decrease in new HIV diagnoses.[16] Furthermore, mapping of mean and total community viral load can identify disparities. For example, a community viral load study in Washington, D.C. revealed the highest viral loads in low-income areas with low education rates;[17] a similar study in New York City showed that low-income neighborhoods had the highest that expanded utilization of antiretroviral therapy is cost-effective due to the number of new infections averted.[19]

# **Knowledge of HIV Status**

The proportion of persons with undiagnosed HIV has steadily decreased from an estimated 25% in 2003 to 12.7% in 2021.[1,20,21] Several HIV transmission modeling studies for the United States have concluded that persons unaware of their HIV diagnosis account for a disproportionately higher number of transmitted HIV infections than persons aware of their HIV diagnosis.[22,23,24] In the 2016 CDC Progression and Transmission of HIV (PATH 2.0) model, investigators estimated that among all persons with HIV in the United States in 2016, the 14.5% of persons with undiagnosed HIV accounted for 37.5% of new HIV transmissions that year (Figure 4).[24] Knowledge of HIV serostatus is the first step toward linking persons to HIV medical care whereby they can receive antiretroviral therapy. In the Ending HIV Epidemic—A Plan for the United States, diagnosing all individuals with HIV as early as possible after infection is one of the main pillars in the strategic initiative.[25] Older studies have shown that persons with a known diagnosis of HIV reduce sexual activities associated with an increased risk of HIV transmission, but these data may not be applicable to the current era.[26,27]

# **Behavioral Prevention Interventions for Persons with HIV**

## **Risk Reduction Counseling**

Counseling to reduce activities that can increase the risk of HIV transmission to others is inadequate as a primary strategy for reducing HIV transmission. For persons with HIV, the impact of consistently taking antiretroviral therapy and maintaining undetectable HIV RNA levels far exceeds the impact of prevention strategies that rely on behavioral interventions. Nevertheless, risk reduction counseling for persons with known HIV remains a complementary piece of a comprehensive prevention strategy.[28] The CDC has identified evidence-based risk reduction counseling strategies for people with HIV.[29] Moreover, older studies that examined the impact of behavioral interventions for people with HIV showed a reduction in self-reported condomless sex as well as a decline in the incidence of sexually transmitted infections in persons who received behavioral interventions.[30] In the current era, however, medical providers often do not see risk reduction counseling in preventing HIV transmission from persons with HIV is minor compared with the impact of taking antiretroviral therapy and maintaining undetectable HIV RNA levels.

## **HIV Status Disclosure and Partner Testing**

Partner counseling and referral services is a public health service that helps people with HIV disclose their HIV status to current or former sex or injection drug partners. The public health system provides a trained counselor who can work with the person newly diagnosed with HIV to support disclosure to partners, as well as to provide partner notification directly in cases where the person with newly diagnosed HIV is not able to disclose their HIV status. Partner notification and testing is important because of the high yield in HIV case finding.[32,33] In two national studies, partner counseling and referral services, including partner notification and HIV testing, effectively identified a substantial number of partners with a new HIV diagnosis; in these studies, 8% of the partners of persons newly diagnosed with HIV tested positive for HIV, and these results were consistent across a 10-year period.[34,35]

## Condom Use

Extensive data has shown that persons with HIV who are taking antiretroviral therapy and maintain HIV RNA levels consistently less than 200 copies/mL do not transmit HIV sexually to others, even with condomless sex.[2,5,11,12] Accordingly, efforts emphasizing condom use as the primary method for HIV prevention have markedly diminished in recent years. Nevertheless, condoms still have a role for persons with HIV, especially in persons newly starting on antiretroviral therapy and for persons who do not have consistently suppressed HIV RNA levels. In addition, condoms prevent the acquisition of other sexually transmitted infections. Consistent, correct condom use decreases HIV transmission by 70 to 80% among HIV-serodifferent heterosexual couples when compared with non-condom users.[36,37,38] In a separate analysis of the protective effect of condom use among HIV-serodifferent male couples who have anal sex, consistent condom use reduced the risk of HIV transmission by approximately 70%.[39] Correlates of condom failure include improper or inconsistent use of appropriate lubricants, amphetamine use, heavy alcohol use, and lower socioeconomic status; proper use of condom-compatible lubricants improves condom effectiveness by lessening the risk of condom breakage and by reducing rectal or vaginal trauma.[40] Unfortunately, negotiating condom use can be problematic in some situations.[28,41,42]

## Serosorting and Seropositioning

The practice of serosorting and seropositioning (strategic positioning) and condom serosorting are selfselected behaviors intended to reduce HIV transmission risk and are referred to as seroadaptive strategies.[43,44] Serosorting describes the practice of choosing sex partners based on concordant HIV status, typically with the practice of selectively using condoms only when sex occurs with persons of a serodifferent HIV status. Strict serosorting for gay men usually refers to men having sex only with other men who have the same HIV status as themselves. Position serosorting (also called strategic positioning or seropositioning) refers to choosing a different sexual position or practice based on the HIV serostatus of one's partner—the person with HIV typically taking the receptive role during unprotected anal sex when their partner does not have HIV. Data on the impact of serosorting have been mixed.[44,45,46,47] There are no guidelines in the United States that recommend serosorting as an impactful prevention measure.

# Importance of Diagnosing and Treating Acute HIV Infection

At the time of early HIV infection (less than 6 months after HIV acquisition), patients usually have high HIV RNA levels and lack significant neutralizing antibodies and thus are considered highly infectious.[48,49,50] Studies have evaluated the relative likelihood of transmitting HIV during acute or early HIV infection, and estimate a significantly higher risk of transmission with acute (or early) HIV than with chronic HIV infection.[51,52,53,54] Using the 2016 CDC Progression and Transmission of HIV (PATH 2.0) model, the Centers for Disease Control and Prevention estimated the highest risk of HIV transmission along the HIV continuum of care occurred in persons with acute HIV who were unaware of their HIV diagnosis (Figure 6).[24] Accordingly, it is important to diagnose individuals with acute and recent (early) HIV whenever possible. To this end, the 2014 CDC HIV testing guidelines recommend the use of the HIV-1/2 antigen-antibody immunoassay for use as the initial HIV screening test in an effort to improve the diagnosis of persons with acute HIV.[55,56] In addition, all persons diagnosed with acute HIV should immediately start antiretroviral therapy to prevent forward transmission of HIV infection, as well as to garner potential long-term immunologic benefit from early therapy.[3,57,58,59]

# Screening and Treatment of Sexually Transmitted Infections

Sexually transmitted infections can facilitate transmission and acquisition of new HIV infection, especially when the individual with HIV is not taking suppressive antiretroviral therapy.[60] Trends of increasing sexually transmitted infections, particularly syphilis, gonorrhea, and chlamydia, have been reported among men with HIV who have sex with men.[61] Non-injection drug use, particularly with methamphetamines, as well as recreational use of erectile-enhancing medications, among men with HIV who have sex with men has been implicated in transmission of sexually transmitted infections.[62,63] Available data on herpes simplex virus (HSV) has not shown a convincing reduction in HIV transmission risk with acyclovir suppressive therapy taken by persons with HIV.[64] Screening, diagnosis, and treatment of sexually transmitted infections in persons with HIV remain a priority, but in the current era, the impact of treatment of sexually transmitted infections on preventing HIV transmission is unclear, especially for persons taking fully suppressive antiretroviral therapy.

• Screening Recommendations for Sexually Transmitted Infections: The 2021 STI Treatment Guidelines recommend that all sexually active persons with HIV undergo routine screening for sexually transmitted infections at all exposed anatomic sites (e.g., pharynx, rectum, urethra) and that testing include serologic screening for syphilis (Table 2).[65] Despite these recommendations, several studies suggest there have been low rates of screening at rectal and pharyngeal sites in persons with HIV.[66,67] Any identified sexually transmitted infection should be promptly treated along with treatment of the partner.[68]

# Circumcision

Male circumcision studies related to HIV prevention have involved men without HIV (as a means to prevent HIV acquisition) and men with HIV (as a means to prevent HIV transmission to others). Given the extremely high impact of HIV antiretroviral therapy and HIV preexposure prophylaxis in preventing HIV transmission and acquisition, the current utility and impact of male circumcision is not known.

- Male Circumcision in Males without HIV: Three studies conducted in Africa from 2002 through 2006 evaluated more than 10,000 African heterosexual couples and addressed the impact of male circumcision on HIV acquisition among men from their female sex partners; overall, there was a 51 to 60% reduction in HIV incidence among men who were circumcised compared to men who were uncircumcised (Figure 5).[69,70,71,72] Studies have also shown that lack of circumcision increases the risk of genital ulcer diseases, which in turn can increase the risk of HIV acquisition.[73,74] A meta-analysis that examined the impact of circumcision on HIV acquisition among men who have sex with men concluded insufficient evidence exists that male circumcision protects against HIV acquisition in this group.[75]
- Male Circumcision in Males with HIV: In contrast to the benefit of circumcision in heterosexual men without HIV, there is no evidence that performing circumcision on males with HIV will reduce HIV transmission to their heterosexual or same-sex partners. In a large study in Rakai District, Uganda, that enrolled 922 uncircumcised heterosexual men with HIV who had a CD4 count of at least 350 cells/mm<sup>3</sup>, investigators randomized the men to receive immediate circumcision or delayed circumcision (24 months later), and circumcision did not reduce HIV transmission to female partners; the investigators stopped the trial early due to futility (Figure 7).[76] In the United States, there are no recommendations for using circumcision as an HIV prevention measure.

# **Prevention Strategies in Persons with Substance Use**

Alcohol and illicit drug use are common among people with HIV and are recognized cofactors for HIV transmission.[77,78,79] In addition, substance use among persons with HIV is associated with higher rates of HIV transmission risk behaviors and lower rates of antiretroviral therapy adherence.[80] Alcohol use is the most prevalent risk factor for poor HIV medication adherence and lower rates of viral suppression.[81,82] Methamphetamine and other amphetamine-type stimulant use can be an important factor in HIV transmission.[83,84] Injection of methamphetamine plays a dual role in HIV transmission—through sharing injection equipment and by altering antiretroviral medication adherence and sexual behavior while high on methamphetamine. Screening for alcohol use disorder, methamphetamine use, and opioid use can play an important role in prevention, especially if persons identified with a substance use disorder can receive counseling, treatment, and prevention services, including use of syringe services programs for individuals who are injecting drugs. Prevention efforts, including implementation of syringe services programs, have led to a major decline in HIV transmission among persons who inject drugs.[85,86]

## Harm Reduction Approach

Harm reduction is based on a set of practical strategies and ideas aimed at reducing negative consequences associated with drug use. In addition, harm reduction programs do not insist on abstinence, and these programs typically utilize a spectrum of services, including providing safe injection equipment, HIV prevention education, and opiate substitution therapy. In harm reduction programs, syringe services often provide a comprehensive set of services beyond basic needle exchange, including HIV counseling and testing, screening for sexually transmitted infections, screening for tuberculosis, vaccination services, and referral to substance use treatment programs. It is extremely important to remember that persons who inject drugs can also acquire and transmit HIV via sexual contact and should be counseled about sexual risk reduction strategies.[87]

## **Syringe Services**

In the United States, an estimated 10 to 15% of people living with HIV acquired HIV through injection drug use.[1] Thus, persons who inject drugs represent a significant source of potential HIV transmission in the United States. The use of sterile needles and clean injection equipment with each fix is an effective way for persons who inject drugs to limit their risk of acquiring and transmitting HIV and hepatitis C virus (HCV).[88] Opponents of syringe services argue that these programs condone and even encourage drug use, especially among youth; an early cohort study showed that a needle exchange program in Montreal was associated with a higher rate of HIV seroconversion, likely due to new social networks formed through the exchange, and this negative view has persisted.[89] In contrast, multiple studies and reviews have concluded that providing sterile equipment to persons who inject drugs reduces injecting risk behaviors, lowers the risk of HIV and/or HCV infection, and facilitates entry into drug treatment.[60,90,91,92]

### **Use of Federal Funds for Syringe Services Programs**

The use of federal funds for needle exchange programs in the United States has been a highly controversial issue.[93] In 1988, opponents of needle exchange programs passed a law that explicitly banned federal funding for any needle exchange programs.[93] The funding ban was in existence until late 2015, except for a brief lift in the ban during 2010 and 2011. In December 2015, the Consolidation Appropriations Act was signed into law, and it modified then existing restrictions on the use of federal funds for programs that distribute sterile needles or syringes. Although the new law continues to prohibit federal funding to purchase sterile needles or syringes, it does allow use of federal funds for other elements of a syringe service program if it is deemed appropriate by a relevant State or local health department (in consultation with the Centers for Disease Control and Prevention). In 2016, the Department of Health and Human Services issued guidance for state, local, tribal, and territorial health departments in requesting and implementing federal funds to support syringe service programs.[94] In June 2022, the Legislative Analysis and Public Policy Association published a

document that provided detailed information regarding state-specific syringe services policies; this document is titled Syringe Services Programs: Summary of State Laws.[95]

# **Summary Points**

- In the United States, integrated, evidence-based biomedical, behavioral, and structural interventions can substantially reduce transmission of HIV from persons with HIV to others.
- Antiretroviral therapy is recommended for all persons with HIV to prevent HIV transmission to others. Persons with HIV who consistently take antiretroviral therapy and maintain undetectable HIV RNA levels do not sexually transmit HIV to others, even with condomless sex.
- Persons unaware of their HIV status account for a disproportionate number of new HIV infections in the United States.
- Most persons aware of their HIV status have a substantial reduction in activities associated with increased risk of HIV transmission to others.
- For persons newly diagnosed with HIV, partner notification and contact HIV testing provides effective HIV case finding and the opportunity to decrease exposure to others.
- Consistent and correct condom use decreases HIV transmission by approximately 80% among serodifferent heterosexual couples and reduces the per-contact risk of HIV infection by 78% among men who have sex with men who practice receptive anal intercourse.
- Persons with undiagnosed acute (early) HIV have the highest relative risk of HIV transmission to others. Diagnosis, counseling, and treatment of persons with acute HIV can substantially reduce new HIV infections.
- Screening and treatment of sexually transmitted infections is an important component of overall HIV prevention services.
- Circumcision reduces the risk of acquiring HIV in heterosexual men, but there is no evidence that performing circumcision on males with HIV will reduce HIV transmission to their heterosexual partners who do not have HIV. The benefit of circumcision in preventing HIV acquisition or transmission in men who have sex with men is unclear.
- Screening for substance use can help providers identify individuals with HIV who could benefit from support to decrease or stop alcohol and illicit drug use. Providing counseling and treatment of substance use, including syringe services programs and opiate substitution therapy when indicated, can reduce injecting and sexual risk behaviors, and promote adherence to antiretroviral therapy.

## Citations

- Centers for Disease Control and Prevention. Estimated HIV Incidence and Prevalence in the United States, 2017–2021. HIV Surveillance Supplemental Report. 2023;28(3). Published May 2023.
   [CDC] -
- Rodger AJ, Cambiano V, Bruun T, et al. Sexual Activity Without Condoms and Risk of HIV Transmission in Serodifferent Couples When the HIV-Positive Partner Is Using Suppressive Antiretroviral Therapy. JAMA. 2016;316:171-81.
   [PubMed Abstract] -
- Cohen MS, Chen YQ, McCauley M, et al. Prevention of HIV-1 infection with early antiretroviral therapy. N Engl J Med. 2011;365:493-505.
   [PubMed Abstract] -
- Cohen MS, Chen YQ, McCauley M, et al. Antiretroviral Therapy for the Prevention of HIV-1 Transmission. N Engl J Med. 2016;375:830-9.
   [PubMed Abstract] -
- Eisinger RW, Dieffenbach CW, Fauci AS. HIV Viral Load and Transmissibility of HIV Infection: Undetectable Equals Untransmittable. JAMA. 2019;321:451-2.
   [PubMed Abstract] -
- Panel on Antiretroviral Guidelines for Adults and Adolescents. Guidelines for the use of antiretroviral agents in adults and adolescents living with HIV. Department of Health and Human Services. Antiretroviral therapy to prevent sexual transmission of HIV (treatment as prevention). December 18, 2019.
   [HIV.gov] -
- Quinn TC, Wawer MJ, Sewankambo N, et al. Viral load and heterosexual transmission of human immunodeficiency virus type 1. Rakai Project Study Group. N Engl J Med. 2000;342:921-9.
   [PubMed Abstract] -
- McNairy ML, El-Sadr WM. Antiretroviral therapy for the prevention of HIV transmission: what will it take? Clin Infect Dis. 2014;58:1003-11.
   [PubMed Abstract] -
- Attia S, Egger M, Müller M, Zwahlen M, Low N. Sexual transmission of HIV according to viral load and antiretroviral therapy: systematic review and meta-analysis. AIDS. 2009;23:1397-404.
   [PubMed Abstract] -
- Fisher M, Pao D, Brown AE, et al. Determinants of HIV-1 transmission in men who have sex with men: a combined clinical, epidemiological and phylogenetic approach. AIDS. 2010;24:1739-47.
   [PubMed Abstract] -
- Rodger AJ, Cambiano V, Bruun T, et al. Risk of HIV transmission through condomless sex in serodifferent gay couples with the HIV-positive partner taking suppressive antiretroviral therapy (PARTNER): final results of a multicentre, prospective, observational study. Lancet. 2019;393:2428-38.
   [PubMed Abstract] -
- 12. Bavinton BR, Pinto AN, Phanuphak N, et al. Viral suppression and HIV transmission in serodiscordant male couples: an international, prospective, observational, cohort study. Lancet HIV. 2018;5:e438-e447.

- Panel on Antiretroviral Guidelines for Adults and Adolescents. Guidelines for the use of antiretroviral agents in adults and adolescents with HIV. Department of Health and Human Services. Initiation of antiretroviral therapy. December 18, 2019.
   [HIV.gov] -
- Reynolds SJ, Makumbi F, Nakigozi G, et al. HIV-1 transmission among HIV-1 discordant couples before and after the introduction of antiretroviral therapy. AIDS. 2011;25:473-7.
   [PubMed Abstract] -
- Montaner JS, Lima VD, Barrios R, et al. Association of highly active antiretroviral therapy coverage, population viral load, and yearly new HIV diagnoses in British Columbia, Canada: a population-based study. Lancet. 2010;376:532-9.
   [PubMed Abstract] -
- Das M, Chu PL, Santos GM, et al. Decreases in community viral load are accompanied by reductions in new HIV infections in San Francisco. PLoS One. 2010;5:e11068.
   [PubMed Abstract] -
- Castel AD, Befus M, Willis S, et al. Use of the community viral load as a population-based biomarker of HIV burden. AIDS. 2012;26:345-53.
   [PubMed Abstract] -
- Laraque F, Mavronicolas HA, Robertson MM, Gortakowski HW, Terzian AS. Disparities in community viral load among HIV-infected persons in New York City. AIDS. 2013;27:2129-39.
   [PubMed Abstract] -
- Long EF, Brandeau ML, Owens DK. The cost-effectiveness and population outcomes of expanded HIV screening and antiretroviral treatment in the United States. Ann Intern Med. 2010;153:778-89.
   [PubMed Abstract] -
- Hall HI, Frazier EL, Rhodes P, et al. Differences in human immunodeficiency virus care and treatment among subpopulations in the United States. JAMA Intern Med. 2013;173:1337-44.
   [Pub Med Abstract] -
- Hall HI, An Q, Tang T, et al. Prevalence of Diagnosed and Undiagnosed HIV Infection United States, 2008-2012. MMWR Morb Mortal Wkly Rep. 2015;64:657-62.
   [PubMed Abstract] -
- Hall HI, Holtgrave DR, Maulsby C. HIV transmission rates from persons living with HIV who are aware and unaware of their infection. AIDS. 2012;26:893-6.
   [PubMed Abstract] -
- Marks G, Crepaz N, Janssen RS. Estimating sexual transmission of HIV from persons aware and unaware that they are infected with the virus in the USA. AIDS. 2006;20:1447-50.
   [PubMed Abstract] -
- Li Z, Purcell DW, Sansom SL, Hayes D, Hall HI. Vital Signs: HIV transmission along the continuum of care - United States, 2016. MMWR Morb Mortal Wkly Rep. 2019;68:267-72.
   [PubMed Abstract] -
- 25. Fauci AS, Redfield RR, Sigounas G, Weahkee MD, Giroir BP. Ending the HIV Epidemic: A Plan for the United States. JAMA. 2019;321:844-845.

- Marks G, Crepaz N, Senterfitt JW, Janssen RS. Meta-analysis of high-risk sexual behavior in persons aware and unaware they are infected with HIV in the United States: implications for HIV prevention programs. J Acquir Immune Defic Syndr. 2005;39:446-53.
   [PubMed Abstract] -
- Noor SW, Ross MW, Lai D, Risser JM. Drug and sexual HIV risk behaviours related to knowledge of HIV serostatus among injection drug users in Houston, Texas. Int J STD AIDS. 2014;25:89-95.
   [PubMed Abstract] -
- Sullivan PS, Carballo-Diéguez A, Coates T, et al. Successes and challenges of HIV prevention in men who have sex with men. Lancet. 2012;380:388-99.
   [PubMed Abstract] -
- 29. Centers for Disease Control and Prevention, Health Resources and Services Administration, National Institutes of Health, American Academy of HIV Medicine, Association of Nurses in AIDS Care, International Association of Providers of AIDS Care, the National Minority AIDS Council, and Urban Coalition for HIV/AIDS Prevention Services. Recommendations for HIV Prevention with Adults and Adolescents with HIV in the United States, 2014. December 11, 2014 (amended December 30, 2016) [CDC] -
- Crepaz N, Lyles CM, Wolitski RJ, et al. Do prevention interventions reduce HIV risk behaviours among people living with HIV? A meta-analytic review of controlled trials. AIDS. 2006;20:143-57.
   [PubMed Abstract] -
- Rose CD, Koester KA, Kang Dufour MS, et al. Messages HIV clinicians use in prevention with positives interventions. AIDS Care. 2012;24:704-11.
   [PubMed Abstract] -
- Golden MR, Dombrowski JC, Wood RW, Fleming M, Harrington RD. A controlled study of the effectiveness of public health HIV partner notification services. AIDS. 2009;23:133-5.
   [PubMed Abstract] -
- Hogben M, McNally T, McPheeters M, Hutchinson AB. The effectiveness of HIV partner counseling and referral services in increasing identification of HIV-positive individuals a systematic review. Am J Prev Med. 2007;33:S89-100.
   [PubMed Abstract] -
- 34. Golden MR, Hogben M, Potterat JJ, Handsfield HH. HIV partner notification in the United States: a national survey of program coverage and outcomes. Sex Transm Dis. 2004;31:709-12. [PubMed Abstract] -
- Katz DA, Hogben M, Dooley SW Jr, Golden MR. Increasing public health partner services for human immunodeficiency virus: results of a second national survey. Sex Transm Dis. 2010;37:469-75.
   [PubMed Abstract] -
- Weller S, Davis K. Condom effectiveness in reducing heterosexual HIV transmission. Cochrane Database Syst Rev. 2002;:CD003255.
   [PubMed Abstract] -
- 37. Giannou FK, Tsiara CG, Nikolopoulos GK, et al. Condom effectiveness in reducing heterosexual HIV transmission: a systematic review and meta-analysis of studies on HIV serodiscordant couples. Expert Rev Pharmacoecon Outcomes Res. 2016;16:489-99.

- Hughes JP, Baeten JM, Lingappa JR, et al. Determinants of per-coital-act HIV-1 infectivity among African HIV-1-serodiscordant couples. J Infect Dis. 2012;205:358-65.
   [PubMed Abstract] -
- Smith DK, Herbst JH, Zhang X, Rose CE. Condom effectiveness for HIV prevention by consistency of use among men who have sex with men in the United States. J Acquir Immune Defic Syndr. 2015;68:337-44.
   [PubMed Abstract] -
- Stone E, Heagerty P, Vittinghoff E, et al. Correlates of condom failure in a sexually active cohort of men who have sex with men. J Acquir Immune Defic Syndr Hum Retrovirol. 1999;20:495-501.
   [PubMed Abstract] -
- Kalichman SC, Williams EA, Cherry C, Belcher L, Nachimson D. Sexual coercion, domestic violence, and negotiating condom use among low-income African American women. J Womens Health. 1998;7:371-8.
   [PubMed Abstract] -
- Raiford JL, Diclemente RJ, Wingood GM. Effects of fear of abuse and possible STI acquisition on the sexual behavior of young African American women. Am J Public Health. 2009;99:1067-71.
   [PubMed Abstract] -
- 43. Cassels S, Katz DA. Seroadaptation among men who have sex with men: emerging research themes. Curr HIV/AIDS Rep. 2013;10:305-13.
   [PubMed Abstract] -
- 44. Kennedy CE, Bernard LJ, Muessig KE, et al. J Sex Transm Dis. 2013;2013:583627. [PubMed Abstract] -
- Khosropour CM, Dombrowski JC, Swanson F, et al. Trends in Serosorting and the Association With HIV/STI Risk Over Time Among Men Who Have Sex With Men. J Acquir Immune Defic Syndr. 2016;72:189-97.
   [PubMed Abstract] -
- 46. van den Boom W, Konings R, Davidovich U, Sandfort T, Prins M, Stolte IG. Is serosorting effective in reducing the risk of HIV infection among men who have sex with men with casual sex partners? J Acquir Immune Defic Syndr. 2014;65:375-9. [PubMed Abstract] -
- Truong HM, Kellogg T, Klausner JD, et al. Increases in sexually transmitted infections and sexual risk behaviour without a concurrent increase in HIV incidence among men who have sex with men in San Francisco: a suggestion of HIV serosorting? Sex Transm Infect. 2006;82:461-6.
   [PubMed Abstract] -
- Cohen MS, Muessig KE, Smith MK, Powers KA, Kashuba AD. Antiviral agents and HIV prevention: controversies, conflicts, and consensus. AIDS. 2012;26:1585-98.
   [PubMed Abstract] -
- Pilcher CD, Joaki G, Hoffman IF, et al. Amplified transmission of HIV-1: comparison of HIV-1 concentrations in semen and blood during acute and chronic infection. AIDS. 2007;21:1723-30.
   [PubMed Abstract] -

- 50. Pilcher CD, Tien HC, Eron JJ Jr, et al. Brief but efficient: acute HIV infection and the sexual transmission of HIV. J Infect Dis. 2004;189:1785-92.
   [PubMed Abstract] -
- Hollingsworth TD, Anderson RM, Fraser C. HIV-1 transmission, by stage of infection. J Infect Dis. 2008;198:687-93.
   [PubMed Abstract] -
- Wawer MJ, Gray RH, Sewankambo NK, et al. Rates of HIV-1 transmission per coital act, by stage of HIV-1 infection, in Rakai, Uganda. J Infect Dis. 2005;191:1403-9.
   [PubMed Abstract] -
- Brenner BG, Roger M, Routy JP, et al. High rates of forward transmission events after acute/early HIV-1 infection. J Infect Dis. 2007;195:951-9.
   [PubMed Abstract] -
- 54. Powers KA, Ghani AC, Miller WC, et al. The role of acute and early HIV infection in the spread of HIV and implications for transmission prevention strategies in Lilongwe, Malawi: a modelling study. Lancet. 2011;378:256-68. [PubMed Abstract] -
- 55. Centers for Disease Control and Prevention (CDC). National HIV Testing Day and new testing recommendations. MMWR Morb Mortal Wkly Rep. 2014;63:537. [PubMed Abstract] -
- 56. Centers for Disease Control and Prevention and Association of Public Health Laboratories. Laboratory testing for the diagnosis of HIV infection: updated recommendations. June 27, 2014.
  [CDC] -
- 57. Smith MK, Rutstein SE, Powers KA, et al. The detection and management of early HIV infection: a clinical and public health emergency. J Acquir Immune Defic Syndr. 2013;63 Suppl 2:S187-99. [PubMed Abstract] -
- Hamlyn E, Jones V, Porter K, Fidler S. Antiretroviral treatment of primary HIV infection to reduce onward transmission. Curr Opin HIV AIDS. 2010;5:283-90.
   [PubMed Abstract] -
- 59. Panel on Antiretroviral Guidelines for Adults and Adolescents. Guidelines for the Use of Antiretroviral Agents in Adults and Adolescents with HIV. Department of Health and Human Services. Special Populations. Early (Acute and Recent) HIV Infection. September 12, 2024. [HIV.gov] -
- 60. Marrazzo JM, del Rio C, Holtgrave DR, et al. HIV prevention in clinical care settings: 2014 recommendations of the International Antiviral Society-USA Panel. JAMA. 2014;312:390-409. [PubMed Abstract] -
- Mayer KH. Introduction: Linkage, engagement, and retention in HIV care: essential for optimal individual- and community-level outcomes in the era of highly active antiretroviral therapy. Clin Infect Dis. 2011;52 Suppl 2:S205-7.
   [PubMed Abstract] -
- 62. Garg S, Taylor LE, Grasso C, Mayer KH. Prevalent and incident hepatitis C virus infection among HIVinfected men who have sex with men engaged in primary care in a Boston community health center. Clin Infect Dis. 2013;56:1480-7.

- 63. Spindler HH, Scheer S, Chen SY, et al. Viagra, methamphetamine, and HIV risk: results from a probability sample of MSM, San Francisco. Sex Transm Dis. 2007;34:586-91. [PubMed Abstract] -
- 64. Celum C, Wald A, Lingappa JR, et al. Acyclovir and transmission of HIV-1 from persons infected with HIV-1 and HSV-2. N Engl J Med. 2010;362:427-39. [PubMed Abstract] -
- Workowski KA, Bachmann LH, Chan PA, et al. Sexually transmitted infections treatment guidelines, 2021. Screening recommendations referenced in treatment guidelines and original recommendation sources. MMWR Recomm Rep. 2021;70(No. RR-4):1-187.
   [2021 STD Treatment Guidelines] -
- Patton ME, Kidd S, Llata E, et al. Extragenital gonorrhea and chlamydia testing and infection among men who have sex with men--STD Surveillance Network, United States, 2010-2012. Clin Infect Dis. 2014;58:1564-70.
   [PubMed Abstract] -
- 67. Hoover KW, Butler M, Workowski K, et al. STD screening of HIV-infected MSM in HIV clinics. Sex Transm Dis. 2010;37:771-6. [PubMed Abstract] -
- Workowski KA, Bachmann LH, Chan PA, et al. Sexually transmitted infections treatment guidelines, 2021. MMWR Recomm Rep. 2021;70(No. RR-4):1-187.
   [2021 STI Treatment Guidelines] -
- Siegfried N, Muller M, Deeks JJ, Volmink J. Male circumcision for prevention of heterosexual acquisition of HIV in men. Cochrane Database Syst Rev. 2009;:CD003362.
   [PubMed Abstract] -
- Auvert B, Taljaard D, Lagarde E, Sobngwi-Tambekou J, Sitta R, Puren A. Randomized, controlled intervention trial of male circumcision for reduction of HIV infection risk: the ANRS 1265 Trial. PLoS Med. 2005;2:e298.
   [PubMed Abstract] -
- Bailey RC, Moses S, Parker CB, et al. Male circumcision for HIV prevention in young men in Kisumu, Kenya: a randomised controlled trial. Lancet. 2007;369:643-56.
   [PubMed Abstract] -
- 72. Gray RH, Kigozi G, Serwadda D, et al. Male circumcision for HIV prevention in men in Rakai, Uganda: a randomised trial. Lancet. 2007;369:657-66. [PubMed Abstract] -
- 73. Weiss HA, Thomas SL, Munabi SK, Hayes RJ. Male circumcision and risk of syphilis, chancroid, and genital herpes: a systematic review and meta-analysis. Sex Transm Infect. 2006;82:101-10. [PubMed Abstract] -
- 74. Van Howe RS. Genital ulcerative disease and sexually transmitted urethritis and circumcision: a metaanalysis. Int J STD AIDS. 2007;18:799-809.
   [PubMed Abstract] -
- 75. Millett GA, Flores SA, Marks G, Reed JB, Herbst JH. Circumcision status and risk of HIV and sexually

transmitted infections among men who have sex with men: a meta-analysis. JAMA. 2008;300:1674-84. [PubMed Abstract] -

- 76. Wawer MJ, Makumbi F, Kigozi G, et al. Circumcision in HIV-infected men and its effect on HIV transmission to female partners in Rakai, Uganda: a randomised controlled trial. Lancet. 2009;374:229-37.
  [PubMed Abstract] -
- Morin SF, Myers JJ, Shade SB, Koester K, Maiorana A, Rose CD. Predicting HIV transmission risk among HIV-infected patients seen in clinical settings. AIDS Behav. 2007;11:S6-16.
   [PubMed Abstract] -
- Mimiaga MJ, Reisner SL, Grasso C, et al. Substance use among HIV-infected patients engaged in primary care in the United States: findings from the Centers for AIDS Research Network of Integrated Clinical Systems cohort. Am J Public Health. 2013;103:1457-67.
   [PubMed Abstract] -
- 79. Samet JH, Cheng DM, Libman H, Nunes DP, Alperen JK, Saitz R. Alcohol consumption and HIV disease progression. J Acquir Immune Defic Syndr. 2007;46:194-9. [PubMed Abstract] -
- Clark T, Marquez C, Hare CB, John MD, Klausner JD. Methamphetamine use, transmission risk behavior and internet use among HIV-infected patients in medical care, San Francisco, 2008. AIDS Behav. 2012;16:396-403.
   [PubMed Abstract] -
- Kalichman SC, Grebler T, Amaral CM, et al. Intentional non-adherence to medications among HIV positive alcohol drinkers: prospective study of interactive toxicity beliefs. J Gen Intern Med. 2013;28:399-405.
   [PubMed Abstract] -
- Kalichman SC, Grebler T, Amaral CM, et al. Viral suppression and antiretroviral medication adherence among alcohol using HIV-positive adults. Int J Behav Med. 2014;21:811-20.
   [PubMed Abstract] -
- Forrest DW, Metsch LR, LaLota M, Cardenas G, Beck DW, Jeanty Y. Crystal methamphetamine use and sexual risk behaviors among HIV-positive and HIV-negative men who have sex with men in South Florida. J Urban Health. 2010;87:480-5.
   [PubMed Abstract] -
- Semple SJ, Zians J, Grant I, Patterson TL. Sexual risk behavior of HIV-positive methamphetamine-using men who have sex with men: the role of partner serostatus and partner type. Arch Sex Behav. 2006;35:461-71.
   [PubMed Abstract] -
- Centers for Disease Control and Prevention (CDC). Syringe exchange programs --- United States, 2008. MMWR Morb Mortal Wkly Rep. 2010;59:1488-91.
   [PubMed Abstract] -
- 86. Aspinall EJ, Nambiar D, Goldberg DJ, et al. Are needle and syringe programmes associated with a reduction in HIV transmission among people who inject drugs: a systematic review and meta-analysis. Int J Epidemiol. 2014;43:235-48.
  [PubMed Abstract] -

- Kral AH, Bluthenthal RN, Lorvick J, Gee L, Bacchetti P, Edlin BR. Sexual transmission of HIV-1 among injection drug users in San Francisco, USA: risk-factor analysis. Lancet. 2001;357:1397-401.
   [PubMed Abstract] -
- 88. Centers for Disease Control and Prevention (CDC). Integrated prevention services for HIV infection, viral hepatitis, sexually transmitted diseases, and tuberculosis for persons who use drugs illicitly: summary guidance from CDC and the U.S. Department of Health and Human Services. MMWR Recomm Rep. 2012;61:1-40. [PubMed Abstract] -
- Vlahov D, Junge B. The role of needle exchange programs in HIV prevention. Public Health Rep. 1998;113 Suppl 1:75-80.
   [PubMed Abstract] -
- Abdul-Quader AS, Feelemyer J, Modi S, et al. Effectiveness of structural-level needle/syringe programs to reduce HCV and HIV infection among people who inject drugs: a systematic review. AIDS Behav. 2013;17:2878-92.
   [PubMed Abstract] -
- Des Jarlais DC. Structural interventions to reduce HIV transmission among injecting drug users. AIDS. 2000;14 Suppl 1:S41-6.
   [PubMed Abstract] -
- 92. Palmateer N, Kimber J, Hickman M, Hutchinson S, Rhodes T, Goldberg D. Evidence for the effectiveness of sterile injecting equipment provision in preventing hepatitis C and human immunodeficiency virus transmission among injecting drug users: a review of reviews. Addiction. 2010;105:844-59. [PubMed Abstract] -
- 93. Weinmeyer R. Needle Exchange Programs' Status in US Politics. AMA J Ethics. 2016;18:252-7. [PubMed Abstract] -
- 94. United States Health and Human Services. Department of Health and Human Services Implementation Guidance to Support Certain Components of Syringe Services Programs, 2016. [CDC] -
- 95. Legislative Analysis and Public Policy Association. Syringe services programs: summary of State laws. June 2022.
   [Legislative Analysis and Public Policy Association] -

## References

- Boily MC, Baggaley RF, Wang L, et al. Heterosexual risk of HIV-1 infection per sexual act: systematic review and meta-analysis of observational studies. Lancet Infect Dis. 2009;9:118-29.
   [PubMed Abstract] -
- Branson BM, Handsfield HH, Lampe MA, et al. Revised recommendations for HIV testing of adults, adolescents, and pregnant women in health-care settings. MMWR Recomm Rep. 2006;55:1-17.
   [PubMed Abstract] -
- Centers for Disease Control and Prevention. Monitoring selected national HIV prevention and care objectives by using HIV surveillance data--United States and 6 U.S. dependent areas, 2014. HIV Surveillance Supplemental Report. 2016;21(No. 4):1-87. Published July 2016.

[<u>CDC</u>] -

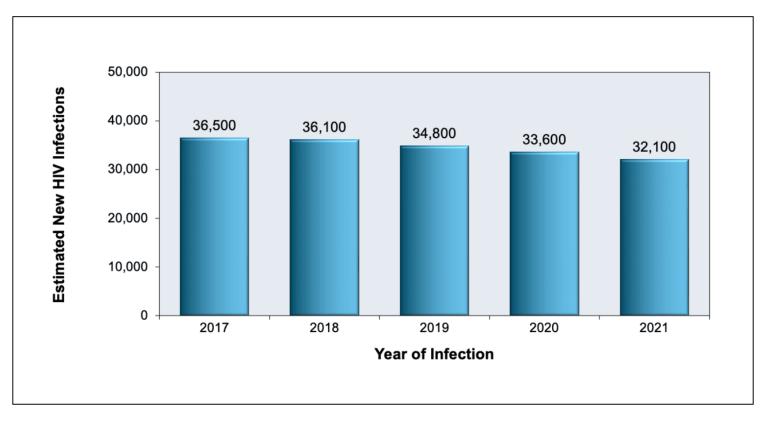
- Crosby RA, Mena L, Geter A. Favourable attitudes towards serosorting are associated with overall less frequent condom use among young Black men having sex men. Sex Health. 2016;13:91-2.
   [PubMed Abstract] -
- Frieden TR, Foti KE, Mermin J. Applying Public Health Principles to the HIV Epidemic--How Are We Doing? N Engl J Med. 2015;373:2281-7.
   [PubMed Abstract] -
- Galvin SR, Cohen MS. The role of sexually transmitted diseases in HIV transmission. Nat Rev Microbiol. 2004;2:33-42.
   [PubMed Abstract] -
- Harris NS, Johnson AS, Huang YA, et al. Vital Signs: Status of Human Immunodeficiency Virus Testing, Viral Suppression, and HIV Preexposure Prophylaxis - United States, 2013-2018. MMWR Morb Mortal Wkly Rep. 2019;68:1117-23.
   [PubMed Abstract] -
- Holt M, Lea T, Schmidt HM, et al. Increasing Belief in the Effectiveness of HIV Treatment as Prevention: Results of Repeated, National Surveys of Australian Gay and Bisexual Men, 2013-15. AIDS Behav. 2016;20:1564-71.
   [PubMed Abstract] -
- Koff A, Goldberg C, Ogbuagu O. Condomless sex and HIV transmission among serodifferent couples: current evidence and recommendations. Ann Med. 2017;49:534-544.
   [PubMed Abstract] -
- Miller WC, Powers KA, Smith MK, Cohen MS. Community viral load as a measure for assessment of HIV treatment as prevention. Lancet Infect Dis. 2013;13:459-64.
   [PubMed Abstract] -
- Pinkerton SD, Abramson PR. Effectiveness of condoms in preventing HIV transmission. Soc Sci Med. 1997;44:1303-12.
   [PubMed Abstract] -
- Wilson DP, Regan DG, Heymer KJ, Jin F, Prestage GP, Grulich AE. Serosorting may increase the risk of HIV acquisition among men who have sex with men. Sex Transm Dis. 2010;37:13-7.
   [PubMed Abstract] -

# Figures

#### Figure 1 Estimated HIV Incidence in United States, 2017-2021

Note: Data for the year 2020 should be interpreted with caution due to the impact of the COVID-19 pandemic on access to HIV testing, care-related services, and case surveillance activities in state/local jurisdictions.

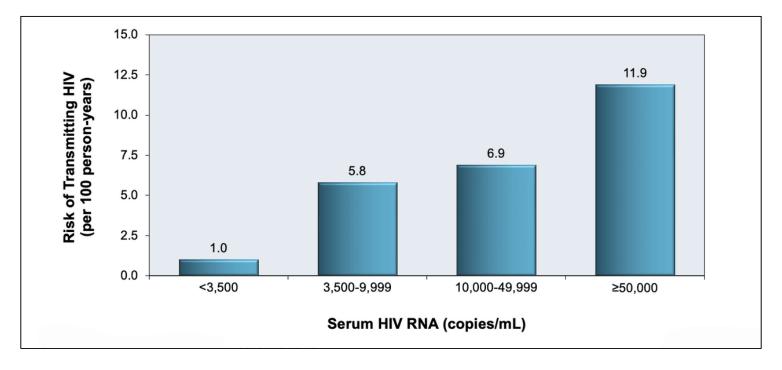
Source: Centers for Disease Control and Prevention. Estimated HIV Incidence and Prevalence in the United States, 2017–2021. HIV Surveillance Supplemental Report. 2023;28(3). Published May 2023.



#### Figure 2 Adjusted Rate Ratio of Heterosexual Transmission of HIV-1 According to Serum HIV-1 RNA Level of the HIV-1 Positive Partner

This graphic illustrates the correlation of risk of HIV transmission and serum HIV-1 levels in the person infected with HIV. No HIV transmissions occurred from persons with HIV who had serum HIV RNA-1 levels less than 1,500 copies/mL.

Source: Quinn TC, Wawer MJ, Sewankambo N, et al. Viral load and heterosexual transmission of human immunodeficiency virus type 1. Rakai Project Study Group. N Engl J Med. 2000;342:921-9.

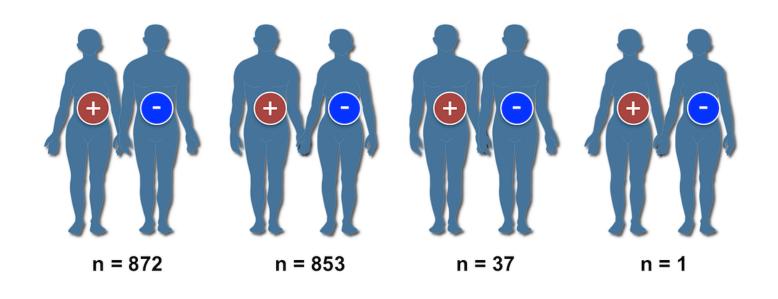


#### Figure 3 (Image Series) - HPTN 052 and Antiretroviral Therapy for the Prevention of HIV-1 (Image Series) - Figure 3 (Image Series) - HPTN 052 and Antiretroviral Therapy for the Prevention of HIV-1 Image 3A: HPTN 052 Patient Population

The HPTN 052 trial enrolled 1,763 HIV serodifferent couples and 97% of the couples were heterosexual.

Source: Cohen MS, Chen YQ, McCauley M, et al. Prevention of HIV-1 infection with early antiretroviral therapy. N Engl J Med. 2011;365:493-505.

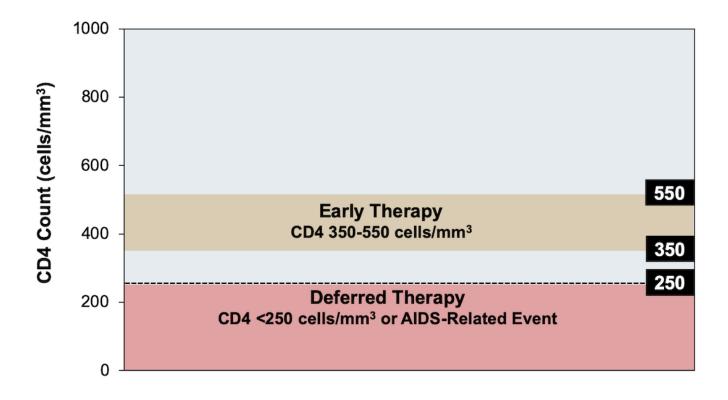
## 1,763 HIV Serodifferent Couples (97% heterosexual)



#### Figure 3 (Image Series) - HPTN 052 and Antiretroviral Therapy for the Prevention of HIV-1 Image 3B: HPTN 052 Study Design

The HIV-positive participants in the early therapy arm received combination antiretroviral therapy and those in the deferred therapy arm started when their CD4 decreased to less than 250 cells/mm<sup>3</sup> or they had an AIDS-related event.

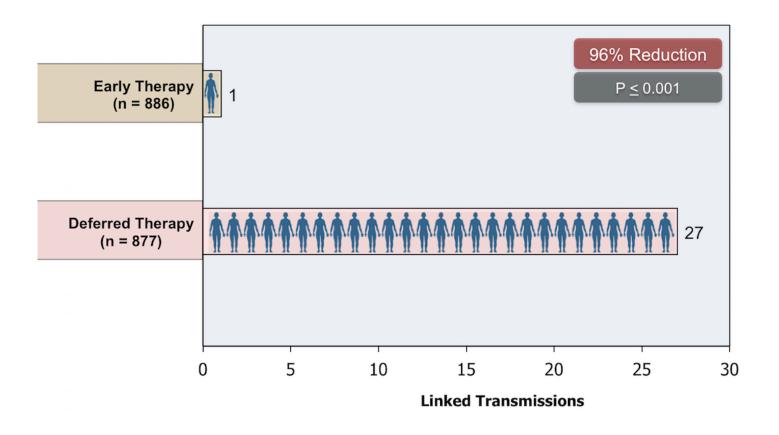
Source: Cohen MS, Chen YQ, McCauley M, et al. Prevention of HIV-1 infection with early antiretroviral therapy. N Engl J Med. 2011;365:493-505.



#### Figure 3 (Image Series) - HPTN 052 and Antiretroviral Therapy for the Prevention of HIV-1 Image 3C: HPTN 052 Results

This graphic shows linked transmissions in the two study groups. Couples in the early therapy arm had a 96% reduction in new HIV transmission events.

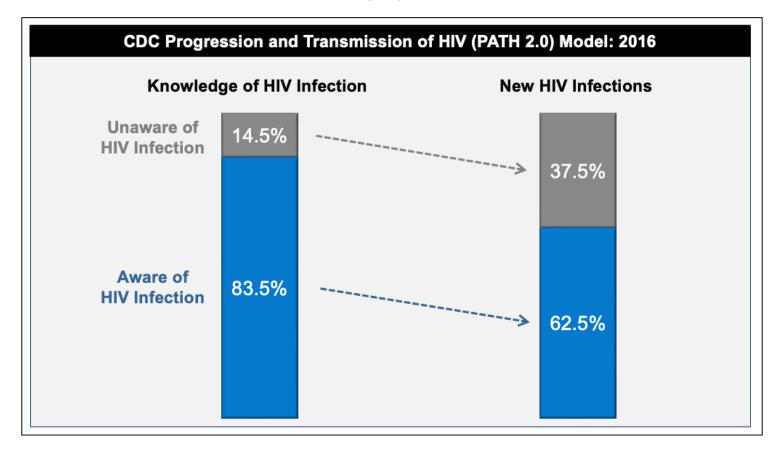
Source: Cohen MS, Chen YQ, McCauley M, et al. Prevention of HIV-1 infection with early antiretroviral therapy. N Engl J Med. 2011;365:493-505.



#### Figure 4 Transmission of HIV and Awareness of HIV Status

This graph shows the estimated percentage of HIV transmissions based on awareness of HIV diagnosis. These estimates were based on the 2016 Center for Disease Control and Prevention (CDC) Progression and Transmission of HIV (PATH 2.0) model.

Source: Li Z, Purcell DW, Sansom SL, Hayes D, Hall HI. Vital Signs: HIV transmission along the continuum of care - United States, 2016. MMWR Morb Mortal Wkly Rep. 2019;68:267-72.

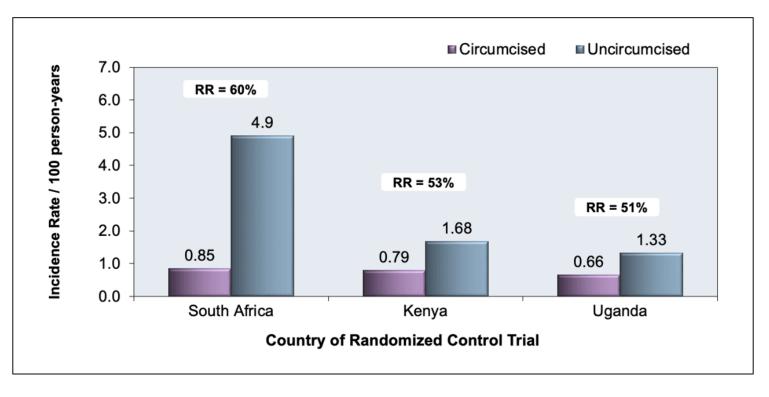


# Figure 5 Risk of Heterosexual Male HIV Acquisition Based on Circumcision Status of Males Living in Africa

#### Abbreviation: RR = relative risk

These three studies conducted in Africa addressed the risk of heterosexual HIV acquisition in men based on their circumcision status. As shown, the risk of HIV acquisition was significantly lower in men who were circumcised when compared with those who were uncircumcised.

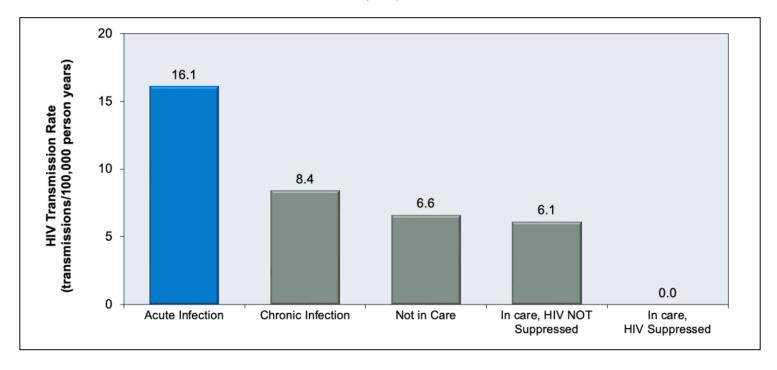
Source: (1) Auvert B, Taljaard D, Lagarde E, Sobngwi-Tambekou J, Sitta R, Puren A. Randomized, controlled intervention trial of male circumcision for reduction of HIV infection risk: the ANRS 1265 Trial. PLoS Med. 2005;2:e298. / (2) Bailey RC, Moses S, Parker CB, et al. Male circumcision for HIV prevention in young men in Kisumu, Kenya: a randomised controlled trial. Lancet. 2007;369:643-56. / (3) Gray RH, Kigozi G, Serwadda D, et al. Male circumcision for HIV prevention in men in Rakai, Uganda: a randomised trial. Lancet. 2007;369:657-66.



#### Figure 6 Relative Risk of HIV Transmission Along the HIV Care Continuum

This graph shows the estimated transmission rate based on transmissions per 100,000 person-years. These estimates were based on the 2016 Center for Disease Control and Prevention (CDC) Progression and Transmission of HIV (PATH 2.0) model.

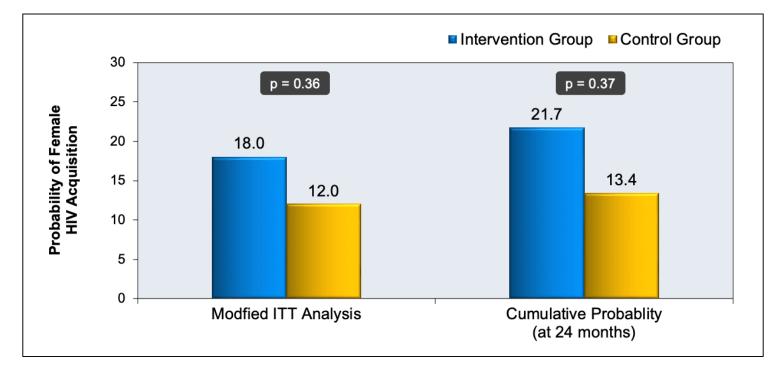
Source: Li Z, Purcell DW, Sansom SL, Hayes D, Hall HI. Vital Signs: HIV transmission along the continuum of care - United States, 2016. MMWR Morb Mortal Wkly Rep. 2019;68:267-72.



# Figure 7 Risk of Heterosexual Female HIV Acquisition Based on Circumcision Status of Male Sex Partner

In this study, investigators in Rakai, Uganda enrolled 922 uncircumcised males with HIV infection who were randomized to undergo immediate circumcision (intervention group) or have circumcision delayed for 24 months (control group). The trial was stopped early because of futility and there was no reduction in risk of female HIV acquisition from their male partners who had been circumcised.

Source: Wawer MJ, Makumbi F, Kigozi G, et al. Circumcision in HIV-infected men and its effect on HIV transmission to female partners in Rakai, Uganda: a randomised controlled trial. Lancet. 2009;374:229-37.



Jse of Antiretroviral Therapy to Prevent Sexual Transmission of HIV		
	<ul> <li>All persons with HIV should be informed that maintaining a plasma HIV RNA (viral load) of &lt;200 copies/mL, including any measurable value below this threshold value, with antiretroviral therapy (ART) prevents sexual transmission of HIV to their partners. Patients may recognize this concept as Undetectable = Untransmittable or U=U (AII).</li> <li>Persons with HIV who are starting ART should use another form of prevention with sexual partners (e.g. condoms, pre-exposure prophylaxis [PrEP] for the HIV-negative sexual partner, sexual abstinence) for at least the first 6 months of treatment and until a viral load of &lt;200 copies/mL has been documented (AII). Many experts would recommend confirming sustained suppression before assuming that there is no further risk of sexual HIV transmission (AIII).</li> <li>When the viral load is ≥200 copies/mL, additional methods are needed to prevent transmission of H to sexual partners until resuppression to &lt;200 copies/mL has been confirmed (AIII).</li> <li>Persons with HIV who intend to rely upon ART for prevention need to maintain high levels of ART adherence or treatment interruption (AIII).</li> <li>At each visit for HIV care, clinicians should assess adherence to ART and counsel patients regarding the importance of ART to their own health as well as its role in preventing sexual HIV transmission (AIII).</li> <li>Providers should inform patients that maintaining a viral load of &lt;200 copies/mL does not prevent acquisition or transmission of other sexually transmitted infections (STIs) (AII).</li> <li>Providers should also routinely screen all sexually active persons with HIV for STIs, both for their ow health and to prevent transmission of STIs to others (AIII).</li> </ul>	
٦atir	g of Recommendations: $A = Strong; B = Moderate; C = Optional$	
	ng of Evidence: I = Data from randomized controlled trials; II = Data from well-designed nonrandomized or observational cohort studies with long-term clinical outcomes; III = Expert opinion	

 Panel on Antiretroviral Guidelines for Adults and Adolescents. Guidelines for the use of antiretroviral agents in adults and adolescents living with HIV. Department of Health and Human Services. Antiretroviral therapy to prevent sexual transmission of HIV (treatment as prevention). December 18, 2019. [HIV.gov]

Table 2.         STI Screening Recommendations in Persons with HIV			
Chlamydia	<ul> <li>For sexually active individuals, screen at first HIV evaluation, and at least annually thereafter</li> <li>More frequent screening might be appropriate depending on individual risk behaviors and the local epidemiology</li> </ul>		
Gonorrhea	<ul> <li>For sexually active individuals, screen at first HIV evaluation, and at least annually thereafter</li> <li>More frequent screening might be appropriate depending on individual risk behaviors and the local epidemiology</li> </ul>		
Syphilis	<ul> <li>For sexually active individuals, screen at first HIV evaluation, and at least annually thereafter</li> <li>More frequent screening might be appropriate depending on individual risk behaviors and the local epidemiology</li> </ul>		
Herpes	<ul> <li>Type-specific herpes simplex virus (HSV) serologic screening for HSV-2 should be considered for persons presenting for an STI evaluation (especially for those persons with multiple sex partners)</li> </ul>		
Trichomoniasis	<ul> <li>Recommended for sexually active women at entry to care and at least annually thereafter</li> </ul>		
HPV, Cervical Cancer	<ul> <li>Sexually active women with HIV who are at least 21 years of age should undergo cervical cancer screening at initial entry to HIV care and again 12 months later.</li> <li>Annual Pap testing is recommended in women with HIV younger than 30 years of age, but if 3 consecutive annual screens are normal, Pap tests can be performed every 3 years.</li> <li>Women with HIV who are 30 years of age and older should have either (1) cervical cancer screening by Pap testing alone or (2) Pap testing plus simultaneous HPV co-testing. If Pap testing alone is used, it should be performed at baseline and every 12 months; if the results of 3 consecutive Pap tests are normal, then follow-up testing can occur every 3 years. If Pap and HPV co-testing is performed and both are negative, follow-up screening can be performed in 3 years.</li> <li>Cervical cancer screening should continue throughout the life in women with HIV.</li> </ul>		
Anal Cancer	<ul> <li>Digital anorectal rectal exam</li> <li>Screen men who have sex with men with HIV</li> </ul>		

STI	Screening Indications and Frequency	
	who are 35 years of age and older and screen all others with HIV who are 45 years of age and older. Specific screening recommendations depend on availability of high resolution anoscopy (HRA).	
Hepatitis B Screening	<ul> <li>At the initial evaluation, test for hepatitis B surface antigen (HBsAg), antibody to hepatitis B core (anti-HBc), and hepatitis B surface antibody (anti-HBs)</li> </ul>	
Hepatitis C Screening	<ul> <li>At the initial evaluation, perform serologic testing for antibody to HCV (anti-HCV), with reflex to HCV RNA for all positive anti-HCV tests</li> <li>Annual HCV serologic testing in men who have sex with men</li> <li>For persons with prior spontaneous or treatment clearance of HCV, screening should be conducted with HCV RNA</li> </ul>	
<b>NOTE</b> : This table is modified from recommendations in the Centers for Disease Control 2021 Sexually Transmitted Infections Treatment Guidelines and the Panel on Opportunistic Infections in Adults and Adolescents with HIV Guidelines for the Prevention and Treatment of Opportunistic Infections in Adults and Adolescents with HIV.		

Source:

- Panel on Opportunistic Infections in Adults and Adolescents with HIV. Guidelines for the prevention and treatment of opportunistic infections in adults and adolescents with HIV: recommendations from the Centers for Disease Control and Prevention, the National Institutes of Health, and the HIV Medicine Association of the Infectious Diseases Society of America. Human papillomavirus disease. July 9, 2024. [HIV.gov]
- Workowski KA, Bachmann LH, Chan PA, et al. Sexually transmitted infections treatment guidelines, 2021. Screening Recommendations and Considerations Referenced in Treatment Guidelines and Original Sources. MMWR Recomm Rep. 2021;70(No. RR-4):1-187. [2021 STI Treatment Guidelines]

National HIV Curriculum