HIV Screening, Diagnosis, and Rapid Antiretroviral Therapy Start

Kento Sonoda, MD, AAHIVS Amy J. Kennedy, MD, MS, AAHIVS Julie Childers, MD, FASAM

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Disclosure Information

- Kento Sonoda, MD, AAHIVS
 - No Disclosures
- Amy J. Kennedy, MD, MS, AAHIVS
 - No Disclosures
- Julie Childers, MD, FASAM
 - No Disclosures



Learning Objectives

- 1. Apply HIV screening tests into clinical practice
- 2. Interpret HIV diagnostic test results
- **3.** Identify resources for starting ART immediately after the diagnosis of HIV



Target Audience

Addiction medicine clinicians in the community setting
Limited access to HIV specialists
Introductory level

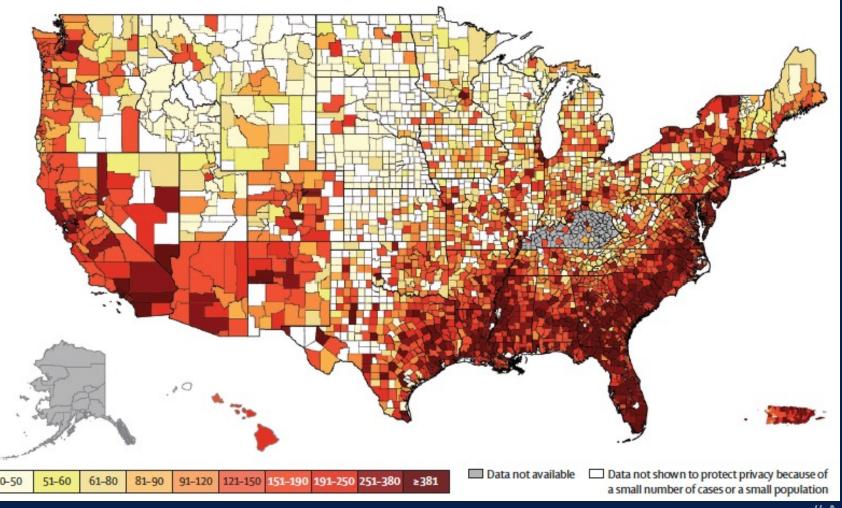


Epidemiology

- More than 1.2 M people living with HIV in the U.S.
 - ◆ 159 K (13%) unaware of HIV infection
- New HIV infections (2019): 37 K
 - Age group highest among people aged 25 to 34 (36%)
 - PWID: 7% of the new HIV diagnoses
- Estimated prevalence of HIV infection among PWID: 1.9%

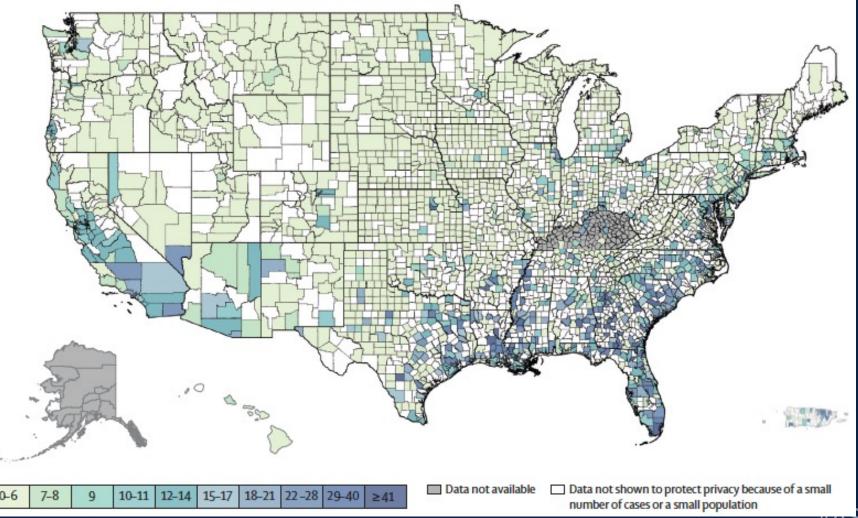


HIV prevalence by county





New HIV diagnoses by county

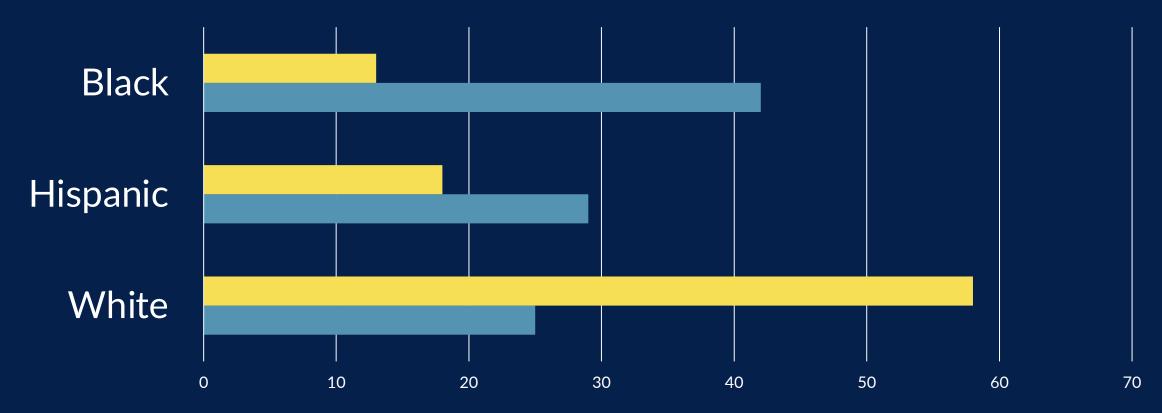




New HIV Diagnoses (2019)

Population (%)

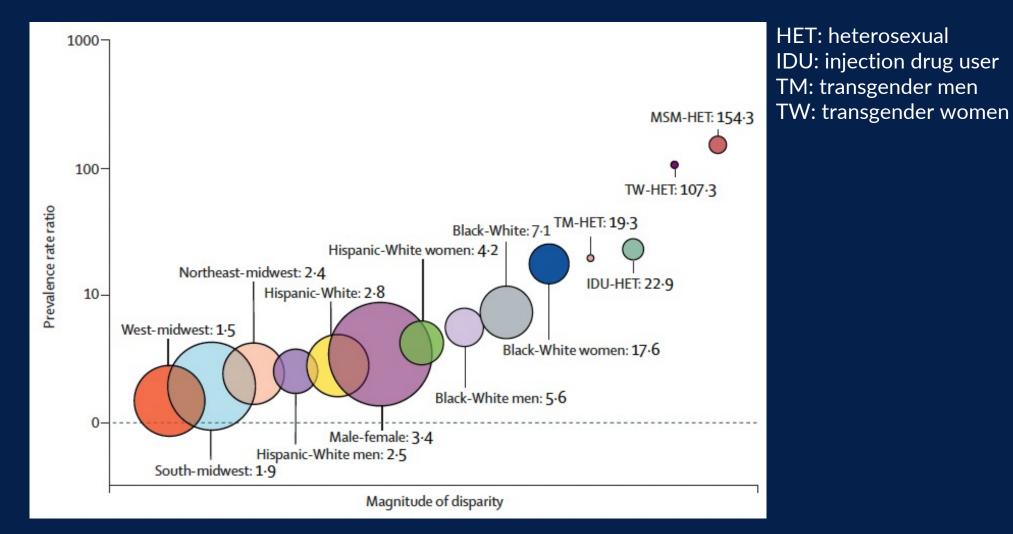
HIV Diagnoses (%)





CDC. HIV Surveillance Report 2019; vol.32. Published May 2021.

Disparities in HIV prevalence

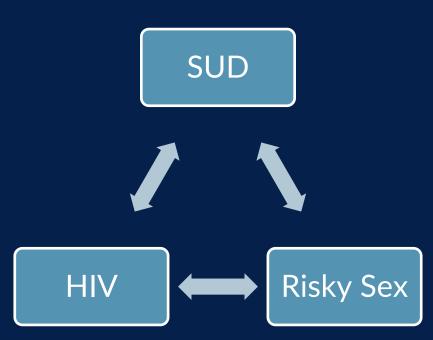




Sullivan PS, Satcher Johnson A, Pembleton ES, et al. Epidemiology of HIV in the USA: epidemic burden, inequities, contexts, and responses. Lancet. 2021;397(10279):1095-1106.

Epidemiology

Estimated prevalence of HIV infection among PWID: 1.9% Unprotected sex: more common in PWID





CDC. HIV Surveillance Report 2019; vol.32. Published May 2021.

Sharma V, Tun W, Sarna A, et al. Prevalence and determinants of unprotected sex in intimate partnerships of men who inject drugs: findings from a prospective intervention study. *Int J STD AIDS*. 2019;30(4):386-395.

Care Continuum





Policies and Issues: HIV Care Continuum. HIV. gov

HIV Screening (Recommendation)

USPSTF:

- Routine, voluntary HIV screening
- All people aged 15 to 65 years (including all pregnant persons)
- Insufficient evidence to determine optional intervals

CDC:

- Persons at increased risk: at least annually
 - PWID, Persons who exchange sex for money or drugs
 - MSM, heterosexual persons with multiple sexual partners



USPSTF. HIV Infection: Screening. Published June 11, 2019.

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Branson BM, et al. Revised Recommendations for HIV Testing of Adults, Adolescents, and Pregnant Women in Health-Care Settings. MMWR. September 22, 2006.

Consider Repeat HIV Screening

Anyone who has been sexually active or is sharing needles

- Anyone with sexually transmitted infections
- Anyone with certain medical conditions
 - Pneumococcal pneumonia, tuberculosis
 - Abnormal PAP smear, thrush, recent vaginal candidiasis
 - New onset of psoriasis and seborrheic dermatitis
 - Immune thrombocytopenia, pancytopenia, lymphoma
 - HBcAb+, HCAb+



Rational for HIV Screening

75% of pts newly diagnosed w/ AIDS: 4 visits in prior 5 years
Time from HIV infection to AIDS: > 5 years

60% of pts diagnosed with HIV: no identified risk/encouterDx
By risk (MSM, IVDU) only 34% could have been identified



Centers for Disease Control and Prevention (CDC). Missed opportunities for earlier diagnosis of HIV infection--South Carolina, 1997-2005. MMWR Morb Mortal Wkly Rep. 2006;55(47):1269-1272.

Case 1

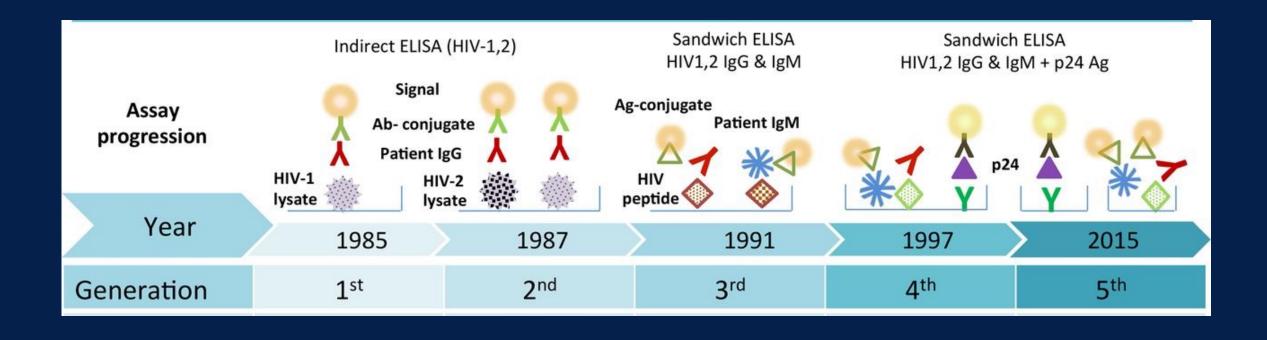
A 30-year-old male is here for follow-up. He was evaluated for mild fever, sore throat, myalgia, and fatigue a week ago. HIV 5th generation test: p24 (+) and HIV 1/2 Ab (-).

• Which of the following is the most appropriate next step?

- 1. HIV Viral Load and Treat
- 2. No further testing
- **3.** T-cell subset testing
- 4. Western Blot HIV-1 Ab testing
- 5. POC HIV testing



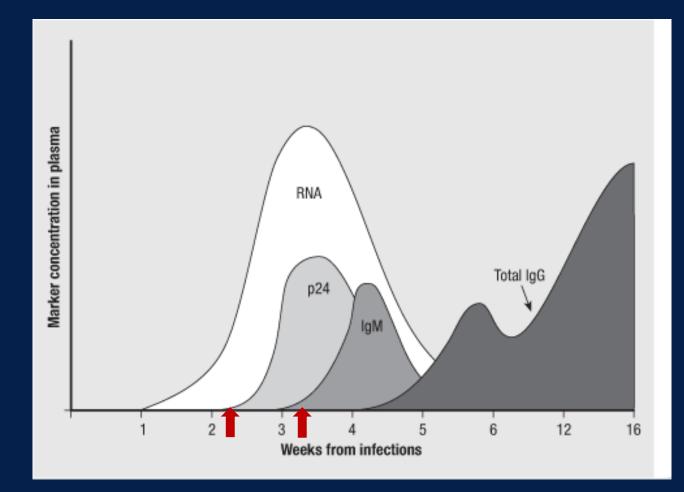
HIV Serologic Tests





Alexander TS. Human Immunodeficiency Virus Diagnostic Testing: 30 Years of Evolution. *Clin Vaccine Immunol*. 2016;23(4):249-253. Published 2016 Apr 4.

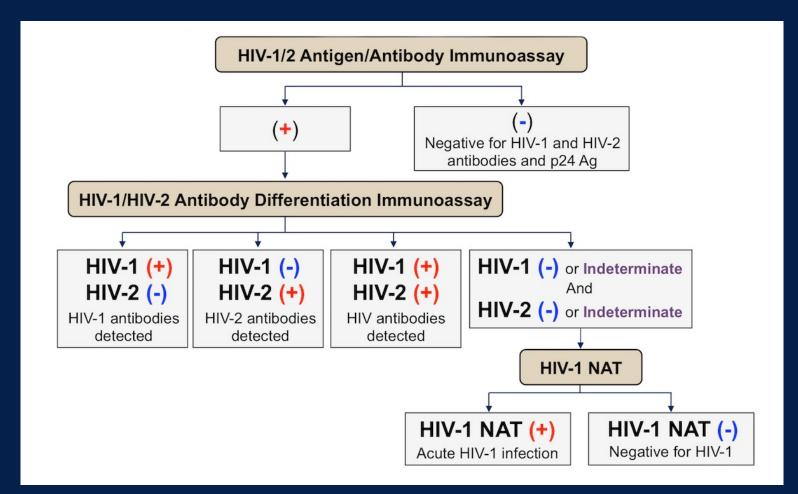
Markers in Acute HIV Infection





Murphy G, Parry JV. Eurosurveillance. Vol. 13. Issue 7-9. Jul-Sep 2008.

CDC Algorithm (4th Generation)





Spach DH. HIV Diagnostic Testing. National HIV Curriculum. CDC. 2018 Quick reference guide: recommended laboratory HIV testing algorithm for serum or plasma specimens.

Interpretation and Plan (5th Generation)

Acute infection (window 2 weeks) and chronic infection

Result	Interpretation	Plan
p24(-) HIV1/2 Ab(-)	HIV(-)	No further testing
p24(+) HIV1/2 Ab(-)	Acute infection	Viral load, Tx
p24(-) HIV1 Ab(+)	Chronic infection	Viral load, CD4, Tx
p24(-) HIV2 Ab(+)	Chronic infection	Refer to ID



Discussion

You saw a 28-year-old male through telemedicine 2 days ago.
 You ordered HIV 5th generation screening test, which showed HIV-1 infection.

How do you deliver the news of HIV diagnosis?



Delivering Test Results

In a private area & In a direct, neutral tone

If negative, provide HIV prevention counseling (ie PrEP)

- If positive,
 - Patient education
 - Linkage to Care HIV (scheduling a follow-up appointment)

- Partner notification requirement: depends on states (sexual partner, needle sharing partner)





- Jared is a 28 yo man with hx of opioid and methamphetamine use disorders. He uses via both injection and smoke/oral routes. He presents to clinic today to get started on buprenorphine-naloxone for his OUD.
- As part of your routine initial exam you order a CMP, HIV, HCV, and STI testing.
- 24 hours later you review his results and his HIV test is positive (5th generation test, p24 negative, HIV 1 antibody positive)



What is your next step?

A) Repeat HIV screening test
B) Check HIV viral load
C) Refer to an HIV/infectious disease specialist
D) Start patient on antiretroviral medication (ART) now





There are no infectious disease specialists in your area.

You call Jared and ask him to come in to discuss lab results.
 You call the lab and add on a HIV viral load and CD4 count.

Viral load comes back with 10,000 copies/ml
CD4 comes back at 550 cells/dl



When do we start ART?

1) Acute HIV
 2) CD4 < 200 cells/dl
 3) CD4 < 500 cells/dl
 4) CD4 > 500 cells/dl
 5) All of the Above



ART "Rapid Start"

Day 1 <u>New HIV Diagnosis</u> Days 1-7 <u>Follow RIA protocol</u> Obtain Baseline Bloodwork Focused Medical/Psychological Evaluation Prescribe ART First HIV Primary Care Visit



Rapid Initiation of ART (RIA) Frequently Asked Questions (ny.gov)

Current Recommendations for Same-Day ART Initiation

- Rapid start or initiating ART on same day as HIV is diagnosed is an emerging strategy to reduce loss to follow-up and decrease time to viral suppression
- Evidence base limited but growing, and outcomes favorable thus far

DHHS ^[1]	WHO ^[2]	IAS-USA ^[3]	
 Recommended at time of diagnosis (when possible) or soon afterward 	 Recommended for all PWH, including same day, if patient is ready* 	 Start ART as soon as possible, including immediately after diagnosis, if patient is 	
 Resource intensive US experience from observational trials 		ready	

*Rapid initiation defined as within 7 days of diagnosis. Priority should be given to patients with advanced disease.



Halperin J and Rockstroh 2019, DHHS Guidelines 2019, WHO Guidelines 2017, Saag 2018

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Why do we start ART early?

1. ART substantially reduces HIV transmission (by >95%)



2. Survival benefit with initiation of ART, even at CD4 count >500



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Roger JAMA 2016, Roger Lancet 2019, Opravil AIDS 2002

Why do we start ART early?

3. ART regimens are effective, safe, convenient (1 pill/day), and well tolerated

4. People with HIV have higher levels of inflammation and endothelial dysfunction which improves with ART



Initial Lab Work-Up

HIV Tests

- Repeat HIV screen (if first screen done outside system)
- HIV Viral Load
- CD4 Count
- HIV Genotype (integrase only if concern for resistance*)

Basic Labs
CBC, CMP, UA
A1c, Lipids

Co-occurring infections

- STI testing gonorrhea, chlamydia, syphilis, trichomonas (in women)
- Hepatitis serologies (A, B, C)
- Toxoplasmosis IgG
- TB (ppd or IGRA)
- Cryptococcus antigen



DHHS ART Guidelines, 2019

DHHS Opportunistic Infections Guidelines, 2019

Follow-Up or ART Modification

	2-8 Weeks After ART Initiation or Modification	Q 4 to 8 Weeks Until VL < 200	Q 3 to 4 months (First 2 years)	Q 6 months
Viral Load	Х	Х	Х	
CD 4 count			Х	
BMP	Х			Х
LFTs	Х			Х



Most common antiretroviral medications 2022 Nucleoside Reverse Transcriptase Inhibitors

Emtricitabine FTC Lamivudine 3TC Abacavir ABC TenofovirAF TenofovirDF Zidovudine

TAF TDF ZDV/AZT

Protease Inhibitors Darunavir DRV.cbc

Integrase Inhibitors

Dolutegravir DTG Bictegravir BTG Elvitegravir EVG.cbc Raltegravir RAL Cabotegravir CAB

Non-nucleoside Reverse Transcriptase Inhibitors

Efavirenz EFV Rilpivirine Ril Doravirine Dor



Recommended Regimens for Rapid ART

DHHS^[1]

Recommended Regimens

BIC/FTC/TAF DTG + (TAF or TDF) + (3TC or FTC) (DRV/RTV or DRV/COBI) + (TAF or TDF) + (3TC or FTC)

Regimens Not Recommended

NNRTI-based regimens or DTG/3TC due higher rate of transmitted NNRTI and NTRI drug resistance

Regimens requiring ABC until HLA-B*5701 test results received

IAS-USA^[2]

Recommended Regimens

DTG + (FTC or 3TC)/(TAF or TDF)

DRV/RTV + (FTC or 3TC)/(TAF or TDF)

Regimens Not Recommended

NNRTI-based regimens due to concerns over transmitted drug resistance (K103N)

Regimens requiring ABC until HLA-B*5701 test results received



Halperin J and Rockstroh 2019, DHHS Guidelines 2019, Saag 2018

Why integrase inhibitors?

High barrier to resistance Well-tolerated, minimal side effects Minimal drug-drug reactions





Where to go for help

UCSF – national clinician conference center
 National Clinician Consultation Center (ucsf.edu)

- AETC AIDS Education and Training Center
- DHHS Department of Health and Human Services
- CDC Center for Disease Control



5-minute updates in HIV treatment/HIV Prevention



Primary Prophylaxis Guidelines

Prophylaxis against disseminated mycobacterium avium complex (MAC)

- No longer recommended for adults/adolescents who immediately initiate ART (AII)
- Only recommended in patients with HIV not on ART/viremic with CD4 <50

Two RCT, placebo-controlled trials + observational data demonstrates people with HIV on ART have minimal risk of developing MAC



Daley, IDSA, 2020 #ASAMAnnual2022

Clinical Decision Points

CD4 <200 Begin PJP prophylaxis -> (Bactrim DS QD or MWF) Risk for Candida (no prophylaxis) CD4 <100 Toxoplasmosis prophylaxis (if IgG+) -> (Bactrim DS QD) CD4 <50 Risk for MAC (no prophylaxis) Risk for CMV retinitis (no prophylaxis)



ART/PrEP updates

ART

 Long-acting injectable ART Cabotegravir/rilpivirine (integrase/NNRTI)

PrEP

 TDF/FTC (Truvada) once daily
 NEW: TAF/FTC (Descovy) once daily Not for women at risk through sex
 NEW: Long-acting injectable cabotegravir (integrase inhibitor)



Sexually Transmitted Infection Updates

- Uncomplicated gonococcal infection: ceftriaxone 500mg IM x1 (increased from 250mg)
- Chlamydia infection: doxycycline 100mg BID x 7 days (prior 1gm azithromycin x 1)
- Hepatitis C: Screening now include all adults 18-79 years of age



MMWR, Dec 2020, USPFTF, March 2020 #ASAMAnnual2022

Final Takeaways

Test everyone for HIV (opt-out)

 Repeat HIV screening annually and consider PrEP for anyone at high risk

All HIV+ patients should receive ART

- Decreased transmission, increased survival with rapid start
- First line Rapid Start ART: Integrase inhibitor or Darunavir/c with TAF/FTC



Any Questions?



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Acknowledgement

 Peter Veldkamp, MD, MSc
 Professor of Medicine, Division of Infectious Diseases, University of Pittsburgh School of Medicine



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Resources (Website)

- 1. National HIV Curriculum. Created by University of Washington. https://www.hiv.uw.edu
- 2. Clinical Info HIV gov. Clinical Guidelines. https://clinicalinfo.hiv.gov/en/guidelines
- 3. National Clinician Consultation Center. HIV/AIDS Management. UCSF. https://nccc.ucsf.edu/clinician-consultation/hiv-aids-management/
 - Submit your care online
 - Call for a Phone Consultation
- 4. HIV Drug Interactions. University of Liverpool. https://www.hiv-druginteractions.org
- 5. HIV Basics. Centers for Disease Control and Prevention. https://www.cdc.gov/hiv/basics/index.html



Resources (Article)

- Primary Care Guidance for Persons with HIV: 2020 Update by the HIV Medicine Association of the Infectious Diseases Society of America. <u>https://www.idsociety.org/practice-guideline/primary-care-management-of-people-with-hiv/</u>
- 2. HIV Infection in Adults: Initial Management. Am Fam Physician. 2021 Apr. https://www.aafp.org/afp/2021/0401/p407.html



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- **3.** Sullivan PS, Satcher Johnson A, Pembleton ES, et al. Epidemiology of HIV in the USA: epidemic burden, inequities, contexts, and responses. *Lancet*. 2021;397(10279):1095-1106.
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- **10.** Murphy G, Parry JV. Eurosurveillance. Vol. 13. Issue 7-9. Jul-Sep 2008.
- **11.** CDC. 2018 Quick reference guide: recommended laboratory HIV testing algorithm for serum or plasma specimens.
- **12.** Telling others. CDC. State HIV Laws. The Center for HIV Law and Policy.
- **13**. The New Standard: Treatment Initiation at Time of HIV Diagnosis: Provider FAQ. <u>https://health.ny.gov/diseases/aids/providers/treatment/docs/faqs.pdf</u>. June 2019.
- 14. Halperin J and Rockstroh J. Key Decisions in HIV Care: Rapid Initiation of ART. <u>https://www.clinicaloptions.com</u>. September 29, 2021.
- **15.** Guidelines for managing advanced HIV disease and rapid initiation of antiretroviral therapy, July 2017. Geneva: World Health Organization; 2017. Licence: CC BY-NC-SA 3.0 IGO



- 16. 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. Laboratory testing: laboratory testing for initial assessment and monitoring of patients with HIV receiving antiretroviral therapy. December 18, 2019.
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- 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.
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