

HHS Adult and Adolescent Antiretroviral Treatment Guidelines **September 2025 Updates**

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Last Updated: September 26, 2025

Disclosures

No conflicts of interest or relationships to disclose.

Disclaimer

Funding for this presentation was made possible by 5 TR7HA53202-02-00 from the Human Resources and Services Administration HIV/AIDS Bureau. The views expressed do not necessarily reflect the official policies of the Department of Health and Human Services nor does mention of trade names, commercial practices, or organizations imply endorsement by the U.S. Government. *Any trade/brand names for products mentioned during this presentation are for training and identification purposes only.*

Outline

What's New in the Guidelines

- New chapter on cardiovascular and metabolic complications
- Revisions to laboratory testing and monitoring
- New subsections on initial antiretroviral therapy
 - Acute/early infection, starting during a hospitalization, elite controllers
- Updates to suboptimal CD4 recovery section

New Chapter: Cardiovascular and Metabolic Complications in Persons with HIV

Updates on Cardiovascular Complications for PWH

Key Points

- Alternate ARVs should be considered in place of abacavir and lopinavir/ritonavir if high underlying ASCVD risk or known ASCVD (AII)
- For PWH aged 40-75 years, 10-year ASCVD risk score should be calculated using the Pooled Cohort Equations at least annually
- Women with HIV have higher relative ASCVD risk than men with HIV, plus greater underprediction of ASCVD by risk estimators

Updates on Cardiovascular Complications in PWH

Recommendations for When to Check a Serum Lipid Panel

Updates on Cardiovascular Complications for PWH

Recommendations for When to Check a Serum Lipid Panel

- At entry into care at the time of ART initiation
- 3-6 months after ART initiation once viral suppression has been achieved
- Annually for those aged ≥ 40 years or receiving statin therapy
- Every 1-3 years for those ≤ 40 years and not receiving statin therapy
- With changes in CVD risk factors

Statin Therapy

Recommendations Based on the REPRIEVE Trial

Persons with HIV and Low-Intermediate (<20%) ASCVD* Risk Estimate

Age 40-75 Years

ASCVD 10-Year Risk Score 5-20% **(AI)**

ASCVD 10-Year Risk Score <5% **(CI)**

Moderate-Intensity Statin

- Pitavastatin: 4 mg once daily **(AI)**
- Atorvastatin: 20 mg once daily **(AII)**
- Rosuvastatin: 10 mg once daily **(AII)**

Age <40 Years

Insufficient data for recommendation

*Abbreviations: ASCVD = atherosclerotic cardiovascular disease

Updates on Metabolic Complications for PWH

Weight Gain

- ART initiation should not be delayed due to concerns for weight gain (**AIII**) and ART should not be interrupted or discontinued due to weight gain (**AIII**)
- Specific ARVs should not be selected to prevent or reduce weight gain (**AII**), as available evidence suggests this strategy is ineffective
- Providers should include weight monitoring and counseling on strategies for weight control as part of comprehensive care for PWH

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New Statements About Initiation of ART

New and Revised Statements About Initiation of ART

Acute/early infection:
start as soon as
possible

New diagnosis in
hospital: start if
possible

Elite controller:
generally
recommend starting

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Recommended Frequency of CD4 Monitoring

Clinical Scenario	CD4 Monitoring
After initiating ART	3 months after initiation
During first 1-2 years of ART with viral suppression and CD4 count ≥ 300 cells/mm ³	Every 6 months
After 1-2 years of ART with viral suppression and CD4 count ≥ 300 cells/mm ³	Optional
If CD4 count < 300 cells/mm ³	Every 3-4 months
After modifying ART due to virologic failure	Every 3 to 6 months

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Other Lab Testing Updates

- When to include an integrase resistance test at baseline:
 - If transmitted integrase resistance suspected, history of cabotegravir PrEP, *or history of integrase inhibitor use for PEP*
- When to include an integrase resistance test at treatment failure:
 - If history of integrase inhibitor use for treatment *or prevention*
- HBV serology panel:
 - Entry into care *plus repeat if switching off TAF or TDF if non-immune*

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HIV-Hepatitis B Virus (HBV) Considerations

- HBV reactivation has been reported in PWH with prior exposure (positive core antibody, negative surface antigen) when HBV-active agents are withdrawn
- However, regardless of surface antibody level, PWH and prior HBV exposure are at low risk for reactivation and associated hepatitis/hepatocellular injury
- A monitoring strategy is considered a safe and effective way to assess for HBV reactivation in low-risk PWH (BIII)

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Suboptimal CD4 Recovery Despite Virologic Suppression

- Updated evidence on clinical consequences of suboptimal CD4 recovery, including increased risk of AIDS and non-AIDS events and mortality
- Early diagnosis and prompt ART initiation provides maximal CD4 recovery
- To date, there is no effective therapeutic intervention to improve CD4 count for people with suboptimal CD4 recovery
 - Interventions not recommended: changing ART, intensifying ART, interleukin-2
- Focus on preventive care, modifiable risk factors, management of comorbidities

Summary

Key Take-Home Messages of Guidelines Update

- Emphasizes importance of considering cardiovascular and metabolic comorbidities for PWH
- Underscores that managing comorbidities is more important than changing ART for most individuals with cardiometabolic complications
- Emphasizes that CD4 count monitoring is not necessary for individuals with a suppressed viral load and good CD4 recovery
- Highlights benefit of early ART, ART started during a hospitalization, and ART for elite controllers

Acknowledgment

This Mountain West AIDS Education and Training (MWAETC) program is supported by the Health Resources and Services Administration (HRSA) of the U.S. Department of Health and Human Services (HHS) as part of award [5 TR7HA53202-02-00](#) totaling \$2,820,772 with 0% financed with non-governmental sources.

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