Table 3. Indications for Discontinuing and Restarting Opportunistic Infection Secondary Prophylaxis or Chronic Maintenance in Adults and Adolescents with HIV

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Opportunistic Infection	Indication for Discontinuing Primary Prophylaxis	Indication for Restarting Primary Prophylaxis	Indication for Discontinuing Secondary Prophylaxis/Chronic Maintenance Therapy	Indication for Restarting Secondary Prophylaxis/Chronic Maintenance
Bacterial Enteric Infections: Salmonellosis	Not applicable	Not applicable	Resolution of *Salmonella* infection and after response to ART with sustained viral suppression and CD4 counts >200 cells/mm^3 (CII)	No recommendation
Bartonellosis	Not applicable	Not applicable	• Received at least 3–4 months of treatment, *and* • CD4 count >200 cells/μL for ≥6 months (CIII) Some specialists would only discontinue therapy if *Bartonella* titers have also decreased by four-fold (CIII).	No recommendation
Candidiasis (Mucocutaneous)	Not applicable	Not applicable	If used, reasonable to discontinue when CD4 count >200 cells/mm^3 (AIII).	No recommendation
Coccidioidomycosis	CD4 count ≥250 cells/μL for ≥6 months (CIII)	Restart at CD4 count <250 cells/μL (BIII)	Only for patients with focal coccidioidal pneumonia (AII): • Clinically responded to ≥12 months antifungal therapy, with CD4 count >250 cells/mm^3, and receiving effective ART. • Should continue monitoring for recurrence with serial chest radiographs and coccidioidal serology. For patients with diffuse pulmonary (BIII), disseminated non-meningeal (BIII), or meningeal diseases (AII): • Suppressive therapy should be continued indefinitely, even with increase in CD4 count on ART.	No recommendation

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Cryptococcal Meningitis	Not applicable	Not applicable	If the following criteria are fulfilled (BII): • Completed initial (induction and consolidation) therapy, *and* • Received at least 1 year of antifungal therapy, *and* • Remain asymptomatic of cryptococcal infection, *and* • CD4 count ≥100 cells/mm^3 and with suppressed plasma HIV RNA in response to ART	CD4 count <100 cells/ mm^3 (AIII)
Cytomegalovirus Retinitis	Not applicable	Not applicable	• CMV treatment for at least 3 to 6 months; and with CD4 count >100 cells/mm^3 for >3 to 6 months in response to ART (AII). • Therapy should be discontinued only after consultation with an ophthalmologist, taking into account anatomic location of lesions, vision in the contralateral eye, and feasibility of regular ophthalmologic monitoring. • Routine (i.e., every 3 months) ophthalmologic follow-up is recommended after stopping therapy for early detection of relapse or immune restoration uveitis, and then periodically after sustained immune reconstitution (AIII).	CD4 count <100 cells/mm^3 (AIII)
Histoplasma capsulatum Infection	On ART, with CD4 count >150 cells/mm^3 and undetectable HIV-1 viral load for 6 months (BIII)	For patients at high risk of acquiring histoplasmosis, restart if CD4 count falls to <150 cells/mm^3 (CIII)	If the following criteria (AI) are fulfilled: • Received azole therapy for >1 year, *and* • Negative fungal blood cultures, *and* • Serum or urine *Histoplasma* antigen below the level of quantification, *and* • Undetectable HIV viral load, *and* • CD4 count ≥150 cells/mm^3 for ≥6 months in response to ART	CD4 count <150 cells/mm^3 (BIII)

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Isospora belli Infection	Not applicable	Not applicable	Sustained increase in CD4 count to >200 cells/mm^3 for >6 months in response to ART and without evidence of *I. belli* infection (BIII)	No recommendation
Leishmaniasis: Visceral (and possibly cutaneous leishmaniasis in immunocompromised patients with multiple relapses)	Not applicable	Not applicable	There is no consensus regarding when to stop secondary prophylaxis. Some investigators suggest that therapy can be stopped if CD4 count increases to >200 to 350 cells/mm^3 for 3 to 6 months in response to ART, but others suggest that therapy should be continued indefinitely.	No recommendation
Microsporidiosis	Not applicable	Not applicable	No signs and symptoms of non-ocular (BIII) or ocular (CIII) microsporidiosis and CD4 count >200 cells/mm^3 for >6 months in response to ART.	No recommendation
Mycobacterium avium Complex Disease	Initiation of effective ART (AI)	CD4 count <50 cells/mm^3: only if not on fully suppressive ART (AIII)	If the following criteria are fulfilled (AI): • Completed ≥12 months of therapy, *and* • No signs and symptoms of MAC disease, *and* • Have sustained (>6 months) CD4 count >100 cells/mm^3 in response to ART.	CD4 count <100 cells/mm^3 (AIII)
Pneumocystis Pneumonia	CD4 count increased from <200 to >200 cells/mm^3 for >3 months in response to ART (AI) Can consider when CD4 count is 100–200 cells/mm^3 if HIV RNA remains below limits of detection for ≥3 months to 6 months (BII).	CD4 count <100 cells/mm^3 (AIII) CD4 count 100–200 cells/mm^3 and HIV RNA above detection limit of the assay (AIII).	CD4 count increased from <200 cells/mm^3 to >200 cells/mm^3 for >3 months in response to ART (BII). Can consider when CD4 count is 100–200 cells/mm^3 if HIV RNA remains below limits of detection for ≥3 months–6 months (BII). If PCP occurs at a CD4 count >200 cells/mm^3 while not on ART, discontinuation of prophylaxis can be considered once HIV RNA levels are suppressed to below limits of detection for ≥3 months to 6 months (CIII). If PCP occurs at a CD4 count >200 cells/mm^3 while on ART, continue PCP prophylaxis for life, regardless of how high the CD4	CD4 count <100 cells/mm^3 (AIII) CD4 count 100–200 cells/mm^3 and with HIV RNA above detection limit of the assay (AIII).

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			cell count rises as a consequence of ART (BIII).	
Talaromycosis (Penicilliosis)	CD4 count >100 cells/mm^3 for >6 months in response to ART (BII) *or* If achieved sustained HIV viral suppression for >6 months (BIII)	CD4 count <100 cells/mm^3 (BIII)—if patient is unable to have ART, or has treatment failure without access to effective ART options, and still resides in or travels to the endemic area	CD4 count >100 cells/mm^3 for ≥6 months in response to ART (BII) *or* If achieved sustained HIV viral suppression for >6 months (BIII)	CD4 count <100 cells/mm^3 (BIII)
Toxoplasma gondii Encephalitis	CD4 count increased to >200 cells/mm^3 for >3 months in response to ART (AI) Can consider when CD4 count 100–200 cells/mm^3 if HIV RNA remain below limits of detection for at least 3-6 months (BII)	CD4 count <100 cells/mm^3, (AIII) CD4 count 100–200 cells/μL and with HIV RNA above detection limit of the assay (AIII).	Successfully completed initial therapy, receiving maintenance therapy and remain free of signs and symptoms of TE, and CD4 count >200 cells/mm^3 for >6 months in response to ART (BI).	CD4 count <200 cells/mm^3 (AIII)

For information regarding the evidence ratings, refer to the Rating System for Prevention and Treatment Recommendations in the Introduction section of the Guidelines for the Prevention and Treatment of Opportunistic Infections in Adults and Adolescents with HIV.

Key: ART = antiretroviral therapy; CD4 = CD4 T lymphocyte cell; CMV = cytomegalovirus; MAC = *Mycobacterium avium* complex; PCP = *Pneumocystis* pneumonia; TE = *Toxoplasma* encephalitis