Current Report - HIV

Treatment of AIDS and HIV-Related Conditions - 1996

Ronald H. Goldschmidt, MD, and Betty J. Dong, PharmD

Human immunodeficiency virus (HIV) care, including antiretroviral therapy, treatment of opportunistic infections and other complications of the acquired immunodeficiency syndrome (AIDS), and general care of HIV/AIDS patients and their families will continue to be important primary care problems. The dramatic shift toward primary care in the United States reinforces the importance of managing clinical problems rationally and efficiently without sacrificing quality. About 750,000 Americans are infected with HIV. AIDS is the leading cause of death for men 25 to 44 years old.

HIV/AIDS treatment guidelines can be helpful, especially for primary care clinicians caring for small numbers of HIV-infected persons. This Current Report – HIV article, based on our clinical experience at San Francisco General Hospital and a review of the medical literature, updates our annual treatment recommendations for adults and adolescents.¹

Antiretroviral Therapy Initiating and Changing Therapy

All patients with symptomatic HIV disease and AIDS, including persons with 200 or fewer CD4+ (T-helper) lymphocytes per microliter, should be encouraged to take antiretroviral therapy. Antiretroviral therapy for asymptomatic patients with CD4+ cell counts greater than 500/µL is not recommended unless patients indicate such a preference. A recent study showed that zidovudine (AZT, ZDV) monotherapy for asymptomatic pa-

Supported in part by the Pacific AIDS Education and Training Center, Grant No. 2 U69 PE00118-03, with the Bureau of Health Professions, Health Resources and Services Administration, Department of Health and Human Services. tients with CD4+ cell counts of more than $500/\mu$ L offered no advantage to waiting until CD4+ cell counts decreased to fewer than $500/\mu$ L.² Some experts argue, however, that combination drug therapy at these high CD4+ levels might be beneficial. The optimal time to initiate antiretroviral therapy for asymptomatic patients with CD4+ cell counts of $200-500/\mu$ L is controversial. A cell count near the midrange ($350/\mu$ L) is as good a threshold as any and is our personal choice when patients have no strong opinions.

Benefits of antiretroviral therapy are time limited.³⁻⁷ Changing therapy after a period of time should be expected, as no drug or combination of drugs has been shown to halt the progression of HIV disease. Antiretroviral therapy should be changed when drug toxicities occur or new clinical manifestations (opportunistic infections, malignancies, and other clinical signs of advancing disease) develop. Decreasing CD4+ cell counts, such as a decrease of 100/µL or a 50 percent decline during treatment with a particular antiretroviral regimen, also indicate the need to change drugs.

Antiretroviral Drugs

Six antiretroviral drugs are now approved by the Food and Drug Administration for use against HIV.⁸⁻¹² These drugs include the reverse transcriptase inhibitors zidovudine, didanosine (ddI), zalcitabine (ddC), stavudine (d4T), and lamivudine (3TC), and the protease inhibitor saquinavir. Two other protease inhibitors, indinavir and ritonavir, will be probably approved early this year.

Lamivudine and saquinavir have been approved on the basis of surrogate marker studies. No clinical end point (ie, opportunistic infections, AIDS, death, etc) studies have been reported for either drug. Neither lamivudine nor saquinavir should be used alone, as viral resistance can occur rapidly. The lamivudine studies showed modest increases in CD4+ cell counts (30-50/µL) and decreases in viral load. Saquinavir and other protease inhibitors

Submitted, revised, 11 January 1996.

From the Family Practice Residency Program, San Francisco General Hospital, and the Departments of Family and Community Medicine and Clinical Pharmacy, University of California, San Francisco. Address reprint requests to Ronald H. Goldschmidt, MD, Family Practice Inpatient Service, San Francisco General Hospital, San Francisco, CA 94110.

are associated with increases in CD4+ cell counts and decreases in viral load.

During the past year the findings of two important studies have been presented at conferences and in the news media. Although the results have not been published in the peer-reviewed medical literature, these reports have been influential in generating trends in HIV care. These studies, the AIDS Clinical Trials Group 175 (ACTG 175)¹³ trial and the Delta trial,¹⁴ compared zidovudine monotherapy with other therapies.

The ACTG 175 trial compared four separate treatments: zidovudine alone, didanosine alone, zidovudine plus didanosine, and zidovudine plus zalcitabine. The study included persons who had no prior zidovudine treatment (zidovudine-naive) and those who had received zidovudine (zidovudine-experienced). A total of 2500 persons with mean CD4+ cells counts of about 350/µL (range 200-500/ μ L) were enrolled for a median of 3 years. Among zidovudine-experienced subjects, changing from zidovudine monotherapy to didanosine alone or to didanosine plus zidovudine lowered the death rate from 10 percent to 5 percent and 6 percent, respectively. For zidovudinenaive subjects, zidovudine plus zalcitabine was associated with a statistically significant lower rate of developing the combined end point of new AIDS-defined conditions or death when compared with zidovudine monotherapy. In addition, there was a trend toward more deaths among the zidovudine monotherapy subjects (7 percent) than the other groups (3 to 4 percent), but this difference was not statistically significant.

The Delta trial compared zidovudine alone with zidovudine plus didanosine or zidovudine plus zalcitabine among more than 3000 subjects with a CD4+ cell count of about 200/µL for a median of 26 months. The Delta trial showed a statistically significant improvement in survival for combination therapy for zidovudine-naive persons. Death rates were 16.5 percent for zidovudine alone, 9.6 percent for zidovudine plus didanosine, and 11.6 percent for zidovudine plus zalcitabine.

These two studies provide some support for the concept of changing therapies after a period of time and suggest that the rationale for combination therapy has clinical validity. The ACTG 175 trial shows that adding didanosine to zidovudine therapy or changing to didanosine is benefi-

cial for zidovudine-experienced persons with CD4+ cell counts around 350/µL. For zidovudine-naive persons with CD4+ cell counts around 200/uL, the Delta trial indicates that combination therapy with zidovudine plus didanosine or zidovudine plus zalcitabine is superior to zidovudine monotherapy. These and other studies have prompted many AIDS experts to conclude that combination therapy should always be used, preferably early in the course of HIV disease (at or near CD4+ cell counts of 500/µL) when the viral burden is smaller, rather than later in the course of HIV disease. Although theoretically sound, clinical studies have not yet established that this approach is the best strategy. Combination therapy certainly appears more promising than monotherapy, but monotherapy might also be appropriate for those in whom drug toxicities or interactions are of concern or who are reluctant to take drugs.

Antiretroviral Drug Selection

An entire spectrum of approaches to antiretroviral therapy^{15,16} remains acceptable. Patients and their primary care clinicians need to discuss and individualize strategies of care. For example, some patients will prefer early intervention with multiple agents, whereas others will prefer to withhold medications as long as possible.

Most combination regimens should include the best studied drug, zidovudine, if tolerated. Preliminary reports of prolonged and substantial decreases in viral loads with ritonavir and indinavir therapies support adding a protease inhibitor to antiretroviral drug regimens. Because of potential interactions, zidovudine and stavudine should not be used together. The toxicity profiles of didanosine and zalcitabine prevent their concurrent use. Neurologic toxicity might limit concurrent use of stavudine with either didanosine or zalcitabine.

Depending upon individual patient characteristics and individual drug toxicities, the following reverse transcriptase-inhibitor regimens are reasonable: zidovudine in combination with didanosine, lamivudine, or zalcitabine; didanosine or stavudine alone; possibly lamivudine in combination with stavudine, didanosine, or zalcitabine. Zidovudine or zalcitabine alone could also be considered if other regimens are not acceptable. A protease inhibitor can be added to any of these above regimens.

Table 1. Treatment Rea	gimens for HIV D	lisease	
Company 1/E-vatornia n 1/		Onbthalmalogian 13/	Controintesting n 127
Skin/Mucocutaneous p. 132		Opininaniologic p. 154	uastronnestinar p. 157
		Oral Cavity p. 136	Pulmonary p. 139
Hematologic n 13/	•	Franhageal n 137	Central Nervous System n 142
			Contra Nervous System p. 112
System, Problem, and Drug Regimen	Duration	Adverse Effects/Drug Interactions	Comments
GENERAL/SYSTEMI Antiretroviral (Anti-H	iC IV)		
Combination or monotherapy with the	Indefinitely, unless toxicities	See individual agents	See text. Therapy indicated for all patients with AIDS (including CD4+ lymphocytes
tonowing at ugs	benefits		HIV disease. Initiation of therapy for asymptomatic persons with CD4+ cell counts of 200–500/µL at discretion of patient and primary care clinician
Zidovudine (AZT,	Indefinitely	Malaise, headache, nausea,	Zidovudine is the common first-choice
Retrovir) 200 mg po tid; lower dosages (eg,		insomnia, seizures, myalgias. Anemia, granulocytopenia,	agent, usually in combination with other retroviral drugs
for patients unable to		is an expected effect of zidovudine	Monitor for signs of zidovudine toxicity
and patients with renal		Toxic myopathy (with elevated	fusions or erythropoietin (if endogenous
failure or cirmosis		with long-term use. Lactic acidosis.	can be used if anemia (eg, hemoglobin
		transferase elevations (alanine trans-	require zidovudine therapy. Decrease
		[AST]). Blue to black discoloration	hilcount (ANC) < 500/µL; consider
		of halls and skin in pigmented races	granulocyte colony-stimulating factor (G-CSF). Transfusions and erythropoietin
		Drug interactions Careful monitoring required when	and G-CSF therapies are expensive; changing to alternate agent preferred
		used with other myelosuppressive drugs (ie, trimethoprim-sulfameth-	High-dosage (1200 mg po qd) zidovudine
		oxazole, ganciclovir). Probenecid can increase levels of zidovudine.	therapy can be considered for HIV dementia and thrombocytopenia.
		Acetaminophen (Jylenol) administration does not increase zidovudine toxicity	loxicity of high-dosage zidovudine can be substantial
Didanosine (ddI, Videx)	Indefinitely	Pancreatitis; painful peripheral	Monitor for signs of neuropathy.
250-mg powder bid		sible); nausea, abdominal cramps, diarrhea related to antacid in	to provide adequate buffer for absorption.
125-mg tablet or 167-mg powder po bid for	ţ	formulation; rash; hyperglycemia; hyperuricemia; aminotransferase	dissolve readily in water, can be crushed manually
reduction (ie, 200 mg/d)		elevations; headache, insomnia, seizures; elevated triglyceride	Administer didanosine on empty stomach
in renal failure		and amylase levels; thrombo- cytopenia; retinal atrophy	antagonists, and drugs (eg, ketoconazole,
		Drug interactions Concomitant administration of U	antibiotics) whose absorption is impaired
		antagonists, antacids, and omeprazole (Prilosec) can increase didenosing	by buildred products
		absorption, resulting in toxicity. Avoid	
		(eg, systemic pentamidine). Avoid concomitant neurotoxic drugs (eg.	
		zalcitabine, stavudine, vinca alkaloids, oral ganciclovir). Decreases absorption	1. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1
		of drugs whose absorption is impaired by buffered products (eg, ketoconazole	a da anti-arresta da anti-arresta da anti- arresta da arresta da a Arresta da arresta da ar
		itraconazole, tetracyclines, quinolone antibiotics). Oral and intravenous gan-	
		ciclovir increase didanosine toxicity	
			Continued

System, Problem, and	
Drug Regimen	I

Duration

Indefinitely

Adverse Effects/Drug Interactions

Comments

GENERAL/SYSTEMIC Antiretroviral (Anti-HIV) (cont.)

Zalcitabine (ddC, Hivid) 0.75 mg po tid; 0.375 mg po tid for patients < 30 kg. Dosage reduction in renal failure

Stavudine (d4T, Zerit) Indefinitely 20 mg po bid for patients > 60 kg; 15 mg po bid for patients 40–60 kg; reduce dosage for patients < 40 kg and for patients with renal failure

Lamivudine (3TC, Indefinitely Epivir) 150 mg po bid; 2 mg/kg po bid for patients < 50 kg Saquinavir (Invirase) Indefinitely

Saquinavir (Invirase) 600 mg po tid

Indinavir (Crixivan) 800 mg po tid

Ritonavir (Norvir) 600 mg po bid

Postexposure prophylaxis Zidovudine 200 mg 4 weeks po tid plus lamivudine 150 mg po bid Painful peripheral neuropathy (dosage related, reversible); rash; stomatitis, aphthous ulcers; pancreatitis; esophageal ulceration; seizures; aminotransferase elevations; cardiomyopathy

Drug interactions

Avoid alcohol and other pancreatic toxins (eg, systemic pentamidine). Avoid concomitant neurotoxic drugs (eg, didanosine, stavudine, isoniazid, vinca alkaloids, oral ganciclovir)

Painful peripheral neuropathy. Aminotransferase elevations. Anemia, macrocytosis. Psychological disturbances: insomnia, anxiety, panic attacks

Drug interactions Avoid concomitant use of drugs that can cause neurotoxicity (including didanosine and zalcitabine) or pancreatic toxicity. See didanosine

Headache, fatigue, insomnia; peripheral neuropathy, muscle aches; rash; aphthous ulcers

Headache, confusion; nausea; fever; abdominal pain

Drug interactions

Ketoconazole and grapefruit juice increase saquinavir serum concentration. Avoid concomitant use of saquinavir with rifampin or rifabutin

Nausea, vomiting, diarrhea, asymptomatic hyperbilirubinemia, aminotransferase elevations. Rash, dry skin; nephrolithiasis; insomnia

Drug interactions Avoid concomitant use of indinavir with rifampin or rifabutin

Nausea, vomiting, diarrhea, aminotransferase elevations; hypercholesterolemia, hypertriglyceridemia; paresthesias

Drug interactions Avoid concomitant use of ritonavir with rifampin or rifabutin

See above

Not as effective as zidovudine, didanosine, or stavudine for monotherapy. Neurotoxicity can improve with zalcitabine "rest periods"

Consider for patients intolerant to zidovudine, didanosine, and zalcitabine. Dosages listed in this table are lower than standard dosages (30–40 mg po bid), as studies suggest these lower dosages are associated with equivalent efficacy and a lower incidence of peripheral neuropathy

Not to be used as monotherapy; approved for use with zidovudine. Some evidence that combination therapy with lamivudine plus zidovudine is most effective therapy for zidovudine-naive patients

Decreases viral load. Resistance to saquinavir develops with time

Decreases viral load

Decreases viral load. Preliminary results of one study of ritonavir in patient with advanced disease suggests improvement in disease progression

Administration within 1–2 hours of needlestick or other injury advised. Zidovudine appears safe in pregnancy. Some experts recommend treatment with lamiduvine or didanosine in combination with zidovudine. Counseling required

Continued

Indefinitely

Indefinitely

System, Problem, and Drug Regimen	Duration	Adverse Effects/Drug Interactions	Comments
GENERAL/SYSTEMI Antiretroviral (Anti-HI (cont.)	C V)		
Pregnancy Zidovudine 100 mg po 5 times daily followed by intrapartum zidovudine 2 mg/kg	Until end of pregnancy	See above. Serious adverse effects on fetus not demonstrated in studies to date	Zidovudine therapy, initiated at 14 to 34 weeks' gestation, combined with intrapartum intravenous zidovudine plus oral zidovudine 2 mg/kg qid to infants
V for 1 hour, then mg/kg/h until delivery			for first 6 weeks of life, decreases transmission to infants
Wasting Syndrome Wegestrol (Megace) suspension (40 mg/mL) 300 mg po qd	Indefinitely	Nausea, vomiting; edema; depression. Progestin side effects (hyperglycemia, decreased testosterone levels)	Megestrol can increase appetite and cause fat accumulation with weight gain. Uncertain whether this weight gain improves health. Usually well tolerated. Available also as tablets, but large numbers of tablets are required for administration and are more expensive
Dronabinol (Tetra- ydrocannabinol THC] Marinol) 2.5 mg oo bid 30 minutes- hour before meals. Maximum 20 mg qd	Indefinitely	Restlessness, irritability, insomnia, dizziness, loss of coordination, psychotomimetic effects; fatigue; tachycardia	Increases appetite and can cause weight gain. Uncertain whether this weight gain improves health. Antinauseant. Not recommended for persons sensitive to marijuana effects
Human growth formone. Preparation, losage, and indications not established	Unknown	Arthralgias, joint stiffness, carpal tunnel syndrome; hyperglycemia; hypertriglyceridemia	Studies of a recombinant human growth hormone (r-hGH, Serostim) 0.1 mg/kg/d SQ (average dosage 6 mg/d) demon- strated increased exercise endurance and
			weight gain characterized by increased lean body mass and decreased fat. Experi- mental. Not approved by Food and Drug Administration (FDA)
Anabolic steroids. Preparation, dosage, and indications not established	Unknown	Edema; jaundice	No satisfactory studies to date. Not indi- cated for patients with normal testosterone levels. Treatment must be accompanied by exercise. Unknown whether anabolic steroid therapy improves health
Mycobacterium avium complex (MAC)			
Deserve for signs and symptoms of MAC disease	Indefinitely		
OR			
Rifabutin (Mycobutin) 800 mg po qd or 150 mg po bid	Indefinitely	Nausea (can be reduced by administering 150 mg po bid). Rash. Uveitis with dosages greater than 300 mg po qd and	Survival benefits of MAC prophylaxis not demonstrated. Prophylaxis can be offered for patients with advanced immuno- deficiency (eg, CD4+ cell count < 50 or
or Clarithromycin (Biaxin) 500 mg po qd-bid	Indefinitely	in patients receiving concomitant clarithromycin therapy. Red-orange discoloration of body fluids. Rare neutropenia, thrombocytopenia, anemia; flu-like syndrome; elevated	/5/µL). Azithromycin 1200 mg po q wk appeared effective in preliminary results from one study
or		bilirubin and alkaline phosphatase levels, hepatitis	
			Continued

Table 1. Continued System, Problem, and Drug Regimen Duration Adverse Effects/Drug Interactions Comments **GENERAL/SYSTEMIC** Mycobacterium avium complex (MAC) (cont.) Azithromycin Drug interactions Indefinitely Exclude Mycobacterium tuberculosis (Zithromax) 500 mg Rifabutin increases metabolism of infection before initiating MAC methadone, zidovudine, and po qd prophylaxis clarithromycin; higher dosage of these drugs might be required. Clarithromycin increases rifabutin blood levels and can lead to rifabutin toxicity Acute Ethambutol Optic neuritis (if > 25 mg/kg/d); Indefinitely, if (Myambutol) 15 mg/kg tolerated hyperuricemia; nausea, vomiting po qd (1 g po qd maxi-(minimum of mum); dosage reduc-12 weeks) tion in renal failure plus either Clarithromycin Clarithromycin and azithromycin Treatment indicated for patients with (Biaxin) 500 mg po bid. side effects include nausea, vomiting, progressive signs, symptoms, and Higher dosages (maxidyspepsia, diarrhea, hearing loss, laboratory abnormalities consistent with MAC disease. Evaluate benefits mum 1 g po bid) might aminotransferase elevations be necessary and risks of multidrug regimen before Drug interactions treating. Clinical improvement might take 2-4 weeks Clarithromycin increases serum levels or of rifabutin and can lead to rifabutin toxicity, including severe anterior At least two drugs (preferably ethambutol Azithromycin uveitis. Clarithromycin and azithroplus clarithromycin or azithromycin) 500 mg po qd mycin increase levels of carbamazeshould be used pine and theophylline. Avoid terfenadine (Seldane), astemizole (Hismanal), When both M tuberculosis and MAC infections are suspected, add isoniazid, or loratadine (Claritin) in combinarifampin, and pyrazinamide to ethambutol tion with azole antibiotics because of increased risk of torsades de pointes and clarithromycin. See M tuberculosis and ventricular tachyarrhythmias

For serious illness or failure to respond within 1 month, can add one or two of the following:

Clofazimine Indefinitely (Lamprene) 100 mg po qd

Ciprofloxacin (Cipro) 500–750 mg po qd-bid Indefinitely

Reversible pink to brown-black discoloration of skin, eyes, body secretions; rash. Hyperglycemia. Retinal degeneration

Nausea, vomiting, diarrhea.

Nausea, vomiting, abdominal pain. Anxiety, insomnía, euphoria; tremor; hallucinations; seizures

Drug interactions Ciprofloxacin binds to cations, resulting in decreased absorption.

Administer 2–4 hours after antacids, sucralfate, dairy products, and didanosine

GENERAL/SYSTEMIC

Continued

Am Board Fam Pract: first published as 10.3122/jabfm.9.2.125 on 1 March 1996. Downloaded from http://www.jabfm.org/ on 5 May 2025 by guest. Protected by copyright

System, Problem, and Drug Regimen

GENERAL/SYSTEMIC

Mycobacterium avium complex (MAC) (cont.) Rifampin (Rimactane, Indefinitely Rifadin) 450–600 mg po qd or rifabutin 300 mg po qd

Duration

Amikacin (Amikin) 2–8 weeks 7.5–10.0 mg/kg IM/IV qd

Mycobacterium tuberculosis

Prophylaxis Isoniazid (INH) 300 12 months mg po qd

Active tuberculosis Isoniazid 300 mg po qd Begin with

plus

Rifampin 600 mg po qd

plus

Pyrazinamide (PZA) 15-30 mg/kg po qd (2 g po qd maximum)

plus either

Ethambutol 15 mg/kg po qd (2.5 g po qd maximum)

or

Streptomycin 15 mg/kg IM qd (1 g IM qd maximum)

Histoplasmosis and coccidioidomycosis Acute Amphotericin B (Fungizone) 1.0 mg/ kg IV qd. Decrease to 0.7–0.8 mg/kg qd

if not tolerated

Until 15 mg/kg total dosage has been administered Adverse Effects/Drug Interactions

Rifampin causes red-orange discoloration of body secretions and fluids; elevated bilirubin and alkaline phosphatase levels, hepatitis; anorexia; flu-like syndrome; thrombocytopenia

Drug interactions

Rifampin induces hepatic P-450 enzyme; higher dosages of dapsone, methadone, zidovudine, clarithromycin, fluconazole, ketoconazole, itraconazole, warfarin, protease inhibitors and estrogens might be required

Nephrotoxicity, ototoxicity

Aminotransferase elevations and

hepatitis; administer with pyridoxine to prevent peripheral neuropathy

Drug interactions

Increases metabolism of ketoconazole; larger dosages of ketoconazole might be required. Increased phenytoin and carbamazepine toxicity; monitor levels

See individual drug adverse effects and drug interactions

Not clear whether rifampin or rifabutin provides better activity in multidrug therapy against MAC

Comments

Monitor drug levels in patients with renal failure

INH prophylaxis for all HIV-infected persons with \geq 5-mm intermediatestrength tuberculin skin test induration and those with strong history of tuberculosis exposure regardless of skin test reactivity

Directly observed therapy can permit more flexible (eg, 3 times a week) treatment schedules. Consultation with tuberculosis experts and coordination with tuberculosis control agencies often required

See CENTRAL NERVOUS SYSTEM, Cryptococcus neoformans Amphotericin B recommended initially; oral therapy does not appear as effective. Itraconazole 200 mg po bid or fluconazole 400 mg po bid might be effective. Ketoconazole not indicated *Continued*

Treatment of AIDS 131

4 drugs. After 2 months can continue INH and rifampin only, depending upon susceptibility testing results. Total treatment: at least 9 months, and 6 months beyond culture conversion

Table 1. Continued			tan an a
System, Problem, and Drug Regimen	Duration	Adverse Effects/Drug Interactions	Comments
GENERAL/SYSTEM Histoplasmosis and coccidioidomycosis	IC		and a straight of the Star of the Star Star of the Star
(cont.) Maintenance			
Itraconazole (Sporanox) 200 mg po qd	Indefinitely	Nausea, vomiting. Hypokalemia; hypertension; aminotransferase elevations; adrenal insufficiency; rhabdomyolysis. Teratogenic	Fluconazole 400 mg po qd might be effective
OR		Drug interactions Potent hepatic enzyme inducers, such as rifampin and phenytoin, increase metabolism of intraconazole; higher itraconazole dosages can be required	
Amphotericin B 50 mg IV each week, 2 times a week, or every other we	eek		Optimum frequency of administration not determined
Cryptococcosis		See CENTRAL NERVOUS SYSTEM Cryptococcus neoformans	I ,
SKIN/MUCOCUTAN Kaposi sarcoma	NEOUS		
Observation	Indefinitely		Treatment not required unless lesions are symptomatic or cosmetically bothersome
OR			
Local treatment (radiation therapy, cryotherapy, excision, or intralesional vinblastine)	Until lesions and symptoms are resolved or controlled	Mucositis in head and neck regions from radiation therapy	Treatment effective for cosmetic purposes, for relief of symptoms, and to help reduce edema caused by lymphatic obstruction
OR			
Systemic chemo- therapy with vinblas- tine and vincristine, vincristine alone, or combination of doxo- rubicin, bleomycin, and vincristine	Same	Usual chemotherapeutic agent side effects	Multidrug therapy can help control disease but does not alter prognosis. Consultation by oncologist or AIDS specialist usually required
OR			
Interferon-alpha 5 mU/d SQ, increase by 5 mU/d every 2 weeks as tolerated to a maximum of 35 mU/d	Indefinitely	Fatigue, myalgia, asthenia; neutropenia, thrombocytopenia; aminotransferase elevations	Hematologic toxicities increased in patients taking zidovudine. Dosages greater than 10 mU/d necessary for efficacy
Seborrheic dermatitis			
Accute Hydrocortisone (HC) cream 2.5% plus ketoconazole cream 2% bid; severe cases can require ketocon- azole 200-400 mg no od for 3-4 weaks	Until resolved	See ORAL CAVITY, Candida albicans, ketoconazole	Commonly involves face, eyebrows, retroauricular areas, nasolabial folds, and scalp. Addition of antifungal cream en- hances therapeutic response and reduces the frequency of steroid application
po qu ioi o weeks			
			Continued

J Am Board Fam Pract: first published as 10.3122/jabfm.9.2.125 on 1 March 1996. Downloaded from http://www.jabfm.org/ on 5 May 2025 by guest. Protected by copyright.

Jrug Regimen	Duration	Adverse Effects/Drug Interactions	Comments
KIN/MUCOCUTAN Seborrheic dermatitis	VEOUS (cont.)		
Maintenance	Indofinitohr		
retoconazole cream	muenimieiy		
% bid			
			х. Х
Iucocutaneous ernes simplex			
cute			a an
cyclovir (Zovirax) 00–400 mg po times a day	7–10 days	Oral: nausea, vomiting, diarrhea,	Topical acyclovir ineffective for most episodes
times a day		· · ·	
<i>laintenance</i>			
cyclovir 200–400 mg o 2-3 times a day	Indefinitely		Chronic maintenance therapy can be necessary for repeated episodes
Disseminated, extensiv	ve,	,	
r persistent herpes			
implex	1		•
cwelovir	7–14 days or	Intravenous: lethargy, tremors.	Severe herpes infections (eg. esophagitis.
mg/kg/dose IV q 8 h;	until lesions	confusion, hallucinations; phlebitis;	colitis, encephalitis) require intravenous
osage reduction	resolve	increased serum creatinine,	acyclovir. Maintain good urine output
i ichai lanuic		техетью стузание перигоралу	crystallization
laintenance	T 1 C 1 T		•
cyclovir 200–400 mg	Indefinitely		
52+5 Unics a day			and the second
lerpes zoster			
hingles, disseminated	d,		
cyclovir 10 mg/kg/	7-10 days or		Intravenous therapy preferred. Alternate
ose IV q 8 h; or	until lesions		drugs are foscarnet, vidarabine, and
cyclovir 800 mg po	resolve		cidotovir (available via compassionate
eduction in renal			skin covered with polymyxin B-baci-
ilure for intra-			tracin (Polysporin) ointment q 8 h.
enous acyclovir			Keratoconjunctivitis requires more fre-
or		·	Arear (A & 1) trummine abhiesnon
amciclovir (Famvir)	Same	Headache, nausea, fatigue	Approved only for herpes zoster
00 mg po tid; dosage			intection. Appears as effective as
ilure			compromised patients. Better bioavail-
			ability than acyclovir
avaloria resistant			•
erpes infections			
oscarnet 40 mg/kg/	10-14 days or	See OPHTHALMOLOGIC,	See OPHTHALMOLOGIC, CMV.
ose IV q 8 h; dosage	until lesions	CMV	Trifluridine might be effective. See SKIN/
ilure	cicai		Cidofovir might be effective. See CMV
O r			
hifuriding (Virontic)	Same	Rare hypersensitivity reactions	Apply to affected areas and course with and
% solution q 8 h	Jaine	Rate hypersensitivity reactions	biotic ointment such as bacitracin or poly-
4	÷		myxin B. Keratoconjunctivitis requires
			more trequent (as often as 2 hours, maxi-
			man > drops a day) u nuritune appication
			Constinue J
			Continuea

SKIN/MUCOCUTANEOUS (cont.) **Bacillary angiomatosis** Erythromycin 2 months See GENERAL/SYSTEMIC. 500 mg po qid MAC, clarithromycin, azithromycin. Jarisch-Herxheimer reaction with systemic disease or Doxycycline 2 months 100 mg po bid is available **Eosinophilic folliculitis** Itraconazole 200 mg po once daily with Indefinitely High-potency fluorinated corticosteroid cream bid plus Antihistamine Indefinitely Avoid terfenadine, astemizole, or (eg, diphenhydramine [Benadryl], hydroxyzine [Atarax, Vistaril], torsades de pointes and ventricular doxepin [Sinequan])

HEMATOLOGIC Thrombocytopenia

Observation

OR

Prednisone 60 mg po qd Discontinue as soon as possible

14 days for

infection: 14-21 days

infection

acute retinal

usually required

for extraocular

Long-term corticosteroid therapy increases immunodeficiency; discontinue as soon as possible

Discontinue drugs that can cause

thrombocytopenia

OPHTHALMOLOGIC Cytomegalovirus (CMV)

Prophylaxis Gancyclovir (Cytovene) Indefinitely 1 g po tid

See OPHTHALMOLOGIC, CMV, maintenance

Neutropenia, leukopenia, anemia, thrombocytopenia (avoid if platelet count < 20,000/µL; aminotransferase elevations; renal failure; phlebitis, rash; nausea. Discontinue zidovudine during induction to minimize additive hematologic toxicity (neutropenia). To avoid hematologic toxicity, substitute didanosine, zalcitabine, or stavudine for zidovudine, or change to foscarnet plus zidovudine

Oral gancyclovir primary prophylaxis can be considered but is not currently recommended

Intravitreal ganciclovir by injection or implant appears effective if IV causes unacceptable toxicity. Does not provide systemic therapeutic effect or protection of contralateral eye

Start G-CSF (filgastim, Neupogen) 300 µg SQ qd to 3 times a week for ganciclovir-induced neutropenia (ANC < 500/uL) on two consecutive measurements

Continued

or

Induction Ganciclovir

(Cytovene)

renal failure

5 mg/kg IV q 12 h;

dosage reduction in

Comments

Skin lesions can resolve in 1-3 weeks, but 2 months' treatment needed. Systemic disease (ie, hepatic, splenic, central nervous system, bone, or other organ involvement) or cutaneous recurrences require treatment for 4 months or indefinitely. Azithromycin g po qd and possibly clarithyromycin 500 mg-1 g po qd can be used as alternatives, but less information about efficacy

food might be effective. If no response in 2 weeks, increase dosage to 200 mg po bid for 2 additional weeks. If no response after 4 weeks, discontinue itraconazole. See GENERAL/SYSTEMIC, histoplasmosis. Topical metronidazole might be helpful

loratadine in combination with azole antibiotics because of increased risk of tachyarrhythmias

Treatment not required in absence of bleeding. Consider platelet transfusions prior to invasive procedures. Splenectomy, high-dosage zidovudine, intra-venous gammaglobulin, and interferonalpha can raise platelet count

Am Board Fam Pract: first published as 10.3122/jabfm.9.2.125 on 1 March 1996. Downloaded from http://www.jabfm.org/ on 5 May 2025 by guest. Protected by copyright.

Table 1. Continued

System, Problem, and Drug Regimen

Duration Adverse Effects/Drug Interactions

System, Problem, and Drug Regimen Duration

OPHTHALMOLOGIC Cytomegalovirus (CMV)

14-day

induction

Indefinitely

Indefinitely

(cont.)

Foscarnet (Foscavir) 90 mg/kg/dose IV q 12 h as 2-hour infusion; discontinuation or dosage reduction in renal failure

Alternative to ganciclovir or foscarnet Cidofovir (Vistide) Same 5 mg/kg IV with probenecid each week for 2 weeks, then every 2 weeks; dosage reduction in renal failure

Maintenance

Ganciclovir 5 mg/kg IV as 1-hour infusion. 7 times a week or 6 mg/kg IV 5 times a week; dosage reduction in renal failure

or

Ganciclovir 1 g po tid

OR

Foscarnet 90 mg/kg IV Indefinitely qd as 2-hour infusion 7 times a week; discontinuation or dosage reduction in renal failure

OR

Foscarnet

plus

Ganciclovir

Nephrotoxicity common; tremors, headaches, occasional seizures, muscle spasms; hypocalcemia, hypercalcemia, hypokalemia, hypophosphatemia, hyperphosphatemia; anemia, granulocytopenia; aminotransferase elevations; phlebitis, penile ulcerations

Adverse Effects/Drug Interactions

Drug interactions Avoid concurrent use of nephrotoxic agents when possible

Nephrotoxicity; fever; nausea; rash; proteinuria. Persons allergic to sulfa compounds can be allergic to probenecid

Anemia, leukopenia; nephrotoxicity; neuropathy

Drug interactions

Oral ganciclovir therapy causes 50% increase in didanosine blood levels; reduce didanosine dosage by 50%

Administered by infusion pump via central line. Infusion of 500 mL-1 L normal saline before each foscarnet administration can minimize nephrotoxicity. Twenty-four-hour creatinine clearance should be measured in cachectic patients and in patients with renal insufficiency to ensure proper use of administration nomogram

Comments

Not known whether cidofovir is as effective as ganciclovir or foscarnet. Available by compassionate use

Lifelong suppressive therapy required to prevent recurrence of retinitis. Daily administration is optimal. Administer G-CSF or change to foscarnet if ANC consistently < 500/µL

Oral ganciclovir might be as effective for maintenance therapy as intravenous regimens. Oral absorption is erratic when diarrhea is present. Administer on empty stomach to improve absorption

Maintenance with 120 mg/kg/d might be more effective but also more toxic

Combination therapy not routinely recommended. Can be used after resistance to both drugs demonstrated. Continue maintenance dosage of current drug; induce alternate drug, followed by maintenance with both drugs. Reinduction with ganciclovir or foscarnet might be helpful for recurrences when alternative drug cannot be administered

Continued

Am Board Fam Pract: first published as 10.3122/jabfm.9.2.125 on 1 March 1996. Downloaded from http://www.jabfm.org/ on 5 May 2025 by guest. Protected by copyright.

. .

System, Problem, and Drug Regimen	Duration	Adverse Effects/Drug Interactions	Comments
ORAL CAVITY Candida albicans Clotrimazole (Mycelex) troches 10 mg 5 times a day or vaginal suppositories 100 mg ad-bid. Dissolve	1–2 weeks or until resolved; maintenance (with lowest effective dosage) might be required	Minimal toxicity. Unpleasant taste, nausea, vomiting; amino- transferase elevations	Troches have high sugar content and often require frequent administration. Suppositories can be more convenient
slowly in mouth OR	for severe or fre- quent recurrences		
Nystatin (Mycostatin) 100,000 U/mL, swish and swallow 5 mL po q 6 h or one 500,000-U tablet dissolved slowly in mouth q 6 h	Same	Large oral doses can produce diarrhea, nausea, vomiting	Generally less effective than ketocona- zole, fluconazole, and clotrimazole. Can be effective in fluconazole-resistant candidal infection
OR			
Fluconazole (Diflucan) 100–200 mg po qd followed by maintenance therapy 50–100 mg po qd; 100–200 mg po once weekly less effective	Same	See CENTRAL NERVOUS SYSTEM, Cryptococcus neoformans	More expensive than other agents. Effective in oral candidiasis unresponsive to above oral agents. Higher dosages might be required. Itraconazole 200 mg po qd might be effective against fluconazole-resistant <i>Candida albicans</i>
OR			
Ketoconazole (Nizoral) 400 mg po qd followed by maintenance therapy 200 mg og d bid	Same	Nausea, vomiting; aminotransferase elevations; anaphylaxis, urticaria. Higher dosages can suppress testerone levels; gynecomastia; adrenal suppression	
for 7 consecutive days per month		Drug interactions Need gastric acidity to be effective; avoid antacids, H2 antagonists; administer 2 hours apart from didanosine. Higher dosages might be necessary if taking rifampin	
Amphotericin B mouthwash 0.1 mg/mL, swish and swallow 5 mL qid	Same	Unpalatable; nausea, vomiting	Not absorbed. No systemic effects. Must be prepared from IV solution. Intravenous amphotericin B might be necessary for severe disease
Periodontal disease Hydrogen peroxide gargles for 30 sec bid	Indefinitely		Less expensive than chlorhexidine. Listerine gargles can be effective
OR			Oral hygiene measures with manual removal of plaque are essential. Severe periodontal disease can require antibiotic therapy with metronidazole 250 mg po tid for 7–10 days (alternatives: clindamycin or amoxicillin/clavulanate [Augmentin])
Chlorhexidine gluconate (Peridex) oral rinse 15 mL swished in mouth	Indefinitely	Staining of teeth	
for 30 sec bid			Continued

Table 1. Continued			and the second secon
System, Problem, and Drug Regimen	Duration	Adverse Effects/Drug Interactions	Comments
ESOPHAGEAL <i>Candida albicans</i> Fluconazole 200– 400 mg po qd; higher dosages might be required	14–21 days; maintenance with lowest effective dosage	See CENTRAL NERVOUS SYSTEM, Cryptococcus neoformans	Empiric treatment for patients with dysphagia or odynophagia who have oral thrush. Endoscopy with biopsy and cul- tures appropriate for patients who fail to
OR			respond within I week. Ketoconazole less expensive than fluconazole and effective in most patients. Fluconazole
Ketoconazole 200 mg po bid; amphotericin B; see ORAL CAVITY, <i>Candida albicans</i>		de la deservação de la consecuencia de la consecuencia de la consecuencia de la consecuencia de la consecuencia Alterna que de la consecuencia de la Alterna de la consecuencia de la co	effective in more patients than ketocona- zole; can be reserved for ketoconazole- resistant esophageal candidiasis
OR			
Amphotericin B 0.3–0.4 mg/kg IV qd	10 days or until resolution		Candidal esophagitis unresponsive to oral agents requires low-dose IV amphotericin B
Cytomegalovirus		1	
Ganciclovir; foscarnet see OPHTHALMO- LOGIC, CMV	14–21 days	See OPHTHALMOLOGIC, CMV	Diagnose by endoscopic appearance plus biopsy showing CMV inclusion bodies and positive culture. Long-term suppressive therapy not routinely indicated. Consider only after multiple recurrences. Beware of drug resistance
Herpes simplex IV acyclovir; see SKIN/MUCO- CUTANEOUS, disseminated, exten- sive, or persistent	10–14 days; maintenance required	See SKIN/MUCOCUTANEOUS, disseminated, extensive, or persistent herpes simplex	Diagnose by endoscopic appearance plus positive culture
herpes simplex			· · · · · · · · · · · · · · · · · · ·
GASTROINTESTINA	AL.		
Prochlorperazine (Compazine) 2.5–10.0 mg IV or 5–10 mg po or IM q 6 h or 25 mg pr q 12 h	As needed	Fatigue, drowsiness, dizziness, depression; extrapyramidal reac- tions; dystonic reactions; amino- transferase elevations; constipation	Combinations of these agents often necessary
	Acneeded	Samo as above	Company of the second sec
(Reglan) 10 mg po qid qid or 1 mg/kg IV q 3 h or 10 mg IM q 4-6 h. Dosage reduction in renal failure	As needed		Same as above
Lorazepam (Ativan) 0.5-2.0 mg po or SL tid-qid	As needed	Similar to benzodiazepines; antegrade amnesia	Effective for anticipatory nausea
Ondansetron (Zofran) 0.15 mg/kg IV infusion for 15 min q 6 h or 4–10 mg po q 6 h or granisetron (Kytril) I mg po bid	As needed	Constipation, diarrhea, abdominal pain; fever, chills; headache; sedation	Reserved for intractable nausea and vomiting unresponsive to other agents. Ondansetron, metoclopramide, and dexa- methasone (4–10 mg po qd) combination helpful for intractable nausea and vomiting
Dronabinol 2.5–10.0 po q 8–12 h	As needed	See GENERAL/SYSTEMIC, wasting syndrome	Effective in drug-induced nausea
Droperidol (Inapsine) 2.5 mg IM/IV q 4–6 h	As needed	Similar to prochlorperazine	
			Continued

System, Problem, and Drug Regimen	Duration	Adverse Effects/Drug Interactions	Comments
GASTROINTESTIN Diarrhea	AL		
Symptomatic treatment Loperamide (Imodium) 4 mg po initially then 2 mg q 6 h around the clock and pri (maximum 16 mg qd)	As needed	Abdominal cramps, nausea, abdominal distention, vomiting; dizziness, drowsiness	Around-the-clock regimen more effective than prn. Treat to 2–3 bowel movements per day
Diphenoxylate- atropine (Lomotil) 2.5–5.0 mg po 3–6 times daily for 24–48 hours; then 2.5–5.0 mg tid and prn to control diarrhea (maximum 20 mg qd)	As needed	Ileus; nausea, vomiting, abdominal discomfort; anticholinergic side effects secondary to atropine	Same as above. 2.5 mg diphenoxylate- atropine is equivalent to 2 mg morphine sulfate
Paregoric 0.4 mg morphine/mL, 5-10 mL qd-qid, tincture of opium 10 mg morphine/mL, 0.3-1.0 mL po qid and prn (maximum 1 mL/dose or 6 mL/d), or equivalent	As needed	Ileus. Altered mental status, halluci- nations. Adverse effects common to narcotic analgesics	Same as above. 5 mL paregoric and 0.2 mL tincture of opium are equivalent to 2 mg morphine sulfate
Octreotide (Sandostatin) 100 µg SQ tid, increase by 100–200 µg q 1–2 wk until maximum of 500 µg SQ tid	Indefinitely	Nausea, steatorrhea; hyperglycemia; pain at injection site	Not approved by FDA. Efficacy not demonstrated. Long-term safety unknown. Octreotide does not improve malabsorption
Cryptosporidium See Diarrhea, symptomatic treatment	Indefinitely	See Diarrhea, symptomatic treatment	No drug effectively eradicates <i>Cryptosporidium</i> . Azithromycin, clarithromycin, atovaquone, and bovine colostrum (investigational) might be effective
Paromomycin (Humatin) 750 mg po tid	10-14 days or indefinitely	Nausea, vomiting, diarrhea; rare ototoxicity and nephrotoxicity (similar to other aminoglycosides) only if absorbed through ulcerative bowel lesions	Nonabsorbable aminoglycoside. Effective in some patients
Isospora belli Trimethoprim- sulfamethoxazole (TMP-SMX, Septra, Bactrim) 1 DS (double-	21 days	See PULMONARY, PCP	Usually effective
strength) tablet po qid			
Cytomegalovirus Ganciclovir; foscarnet; see OPHTHALOMO- LOGIC, CMV	14-21 days	See OPHTHALMOLOGIC, CMV	Diagnose by endoscopic appearance plus biopsy showing CMV inclusion bodies and positive culture. Recurrences should be re-treated as acute disease. Long- term suppressive therapy not routinely indicated. Consider only after multiple recurrences. Beware of drug resistance

Continued

System, Problem, and Drug Regimen	Duration	Adverse Effects/Drug Interactions	Comments
· · · · · · · · · · · · · · · · · · ·			
PULMONARY			
'neumocystis carinii			
neumonia (PCP)			
rophylaxis or suppression	of	and the second	
CP for patients with AIL	DŠ v v v v v v		
ncluding CD4+ cell coun	t		
200/µL), unexplained			
ver, or oral candidiasis	•		
MP-SMX	Indefinitely	See TMP-SMX, below	TMP-SMX considered most effective for
DS tablet po qd	(1,2,2,2,2,2,2,2,2,2,2,2,2,2,2,2,2,2,2,2		prophylaxis or suppression. Once-daily
qod or 3 times a			administration is easiest to remember.
eek (eg, M-W-F)			Three-days-per-week regimen might be
• 1 tablet po bid			best tolerated. Multiple TMP-SMX
	A		regimens have been used and all appear
en et al estadour de la composición de			effective. 1MP-SMX provides additional
			prophylaxis against toxoplasmosis
			· *
ternatives to TMP-SM	X		
· prophylaxis or suppress	ion		
apsone 50 mg po	Indefinitely	See dapsone plus TMP. Patients	Probably less effective than TMP-SMX;
d or 100 mg po qd		allergic to sulfa might tolerate	might be less toxic. Check glucose-6-
ith or without TMP		dapsone; some cross-sensitivity	phosphate dehydrogenase (G6PD) before
(rimpex) 15 mg/kg/d			starting dapsone. Lower dosages (eg, 100
pyrimethamine (Dara	1-		mg po 2 times a week) might be effective
im) 25-75 mg po q wl	k .		
R			
haled	Indefinitelv	Adverse systemic effects are mini-	Effective for prophylaxis against primary
ontamidine	,	mal because of low pentamidine	PCP when CD4+ cell count > $150/nL_{\odot}$
eropent) 300 mg		serum concentrations. Broncho-	Does not prevent extrapulmonary
4 wk using		spasm and coughing are common.	disease. Upper lobe recurrences from
espirgard II	· · · · · · · · · · · · · · · · · · ·	especially in smokers. Pretreatment	poor drug distribution when inhaled in
-bulizer		with inhaled bronchodilator (eg.	upright position. Do not use in patients
Jourison		albuterol) can help. Rare pancre-	with possible M tuberculosis infection
		atitis, hypoglycemia; rare nephro-	because of risk of M tuberculosis spread
		toxicity. Increased risk of	by aerosolization
R		spontaneous pneumothorax	 A second sec second second sec
	de provinción de la	•	
lindamycin 450–	Indefinitely	See above	Efficacy and proper dosages for PCP
Mmg no hid_tid	,		prophylavie unknown
o mg po oiu-nu	e dat ser		propinyians unknown
nlua	All the second second		
prus			
		See above	
imaquine 15 mg		See above	
o da	. •		
n	1. S		
K	•		
	To J. Code 1	See al and	
tovaquone	indefinitely	See above	Emcacy and proper dosages for PCP
lepron) suspen-			prophylaxis unknown
m (750 mg/5 mL)			
0 mg po bid		- ⁶ 90	and the second
th or without			
rimethamine	1 I I I I I I I I I I I I I I I I I I I		
–75 mg po q wk			
R		•	
vrimethamine	Indefinitely	Stevens-Johnson syndrome, toxic	No studies clearly demonstrate efficacy
mg-sulfadoxine	· · · · · · · · · · · · · · · · · · ·	epidermal necrolysis: bone mar-	,
10 mg (Fansidar)	a de la composición de	row suppression; gastrointestinal.	
no a 2 wk		central nervous system toxicity	S
poqzwrk		concern nor conservation contenty	

Continued

J Am Board Fam Pract: first published as 10.3122/jabfm.9.2.125 on 1 March 1996. Downloaded from http://www.jabfm.org/ on 5 May 2025 by guest. Protected by copyright.

System, Problem, and Drug Regimen Duration

Adverse Effects/Drug Interactions

Comments

PULMONARY, PCP

Acute PCP TMP-SMX. TMP 21 days 15 mg/kg/d given in 3 divided doses either po or as 1-hour to 2-hour IV infusions; lower dosages (TMP 12 mg/ kg/d) can be effective and less toxic

Adverse effects commonly appear between 7 and 14 days in more than 50% of patients

Rashes: maculopapular, exfoliative, Stevens-Johnson syndrome

Hematologic: neutropenia, leukopenia, thrombocytopenia, anemia

Drug interactions Concurrent leucovorin calcium therapy associated with increased rate of therapeutic failure

Gastrointestinal: nausea, vomiting, aminotransferase elevations

Renal: increased blood urea nitrogen (BUN) and creatinine; hyperkalemia secondary to hypoaldosterone effects of TMP

Hyponatremia

Drug fever. Sepsis-like syndrome, especially upon rechallenge

Adverse effects commonly appear between 7 and 14 days

Orthostatic hypotension can be severe and occur with initial infusion

Pancreatitis; early or delayed hypoglycemia (can occur after discontinuation of therapy); hyperglycemia

Drug interactions Avoid concomitant pancreatic toxins such as didanosine, zalcitabine, and alcohol TMP-SMX is the drug of choice and should be used unless severe reactions (eg, anaphylaxis, severe rashes, Stevens-Johnson syndrome) are of concern. Oral and intravenous routes equally effective. Can provide prophylaxis against toxoplasmosis

Mild rash does not necessitate stopping or changing treatment: institute antihistamine or consider oral desensitization

If ANC < $500/\mu$ L or if platelet count < 30×10^{9} /L and bleeding occurs, consider alternative treatment

Pretreatment with lorazepam, prochlorperazine, metoclopramide, or dronabinol to reduce nausea. See GASTRO-INTESTINAL, nausea and vomiting. Nausea can be less with oral TMP-SMX. Aminotransferase elevations 4–5 times normal require treatment change

TMP decreases creatinine tubular secretion and can falsely elevate serum creatinine levels. Discontinue TMP-SMX if serum creatinine > 3.0 mg/dL

Can be caused by large volume of 5% dextrose in water (D5W) needed for IV administration; can dilute each 80 mg TMP in 75 mL D5W or change to oral TMP-SMX. For severe hyponatremia (Na+ < 115 mEq/dL) can dilute in normal saline; administer within 1 hour of preparation to avoid TMP-SMX precipitation

Drug fever can herald onset of neutropenia, rash, hepatitis, and bone marrow toxicity

Slow IV infusion for 2 hours can prevent hypotension. Check blood pressure at end of infusion

Hypoglycemia can be profound and prolonged, requiring immediate IV D50W followed by D10W glucose infusions. Permanent diabetes can occur

Continued

Alternatives to TMP-SMX for acute PCP Pentamidine 21 days isethionate (Pentam) 4 mg/kg/d as 1-hour to 2-hour IV infusion once a day; 3 mg/kg/d might also be effective

System, Problem, and Drug Regimen Duration

PULMONARY, PCP Alternatives to TMP-SMX for acute PCP (cont.)

OR

Clindamycin (Cleocin) 600 mg IV or po tid 21 days

21 days

21 days

24 days

21 days

plus

Primaquine 30-mg base po qd

OR

Dapsone 100 mg 21 days po qd plus either TMP 15 mg/kg/d po in 3-4 divided doses or pyrimethamine 50-75 mg po qd

Trimetrexate (Neutrexin) 45 mg/m² IV qd

plus

Dapsone 100 mg po qd

plus

Leucovorin calcium (folinic acid) 20 mg/m² IV or po q 6 h

OR

Atovaquone suspension (750 mg/5mL) 750 mg po bid with food

plus

Pyrimethamine 50–75 mg po qd

Renal: increased BUN and creatinine; hyperkalemia. Concomitant nephrotoxic agents (eg, nonsteroidal anti-inflammatory agents) and dehydration increase risk of nephrotoxicity

Adverse Effects/Drug Interactions

Rare: neutropenia, thrombocytopenia; hypocalcemia, hypomagnesemia; aminotransferase elevations; cardiac arrhythmias (rare) with prolongation of QT interval and T wave flattening

Maculopapular rash (day 10–12 most common, usually selflimiting), fever; diarrhea, nausea, vomiting, abdominal cramps, *Clostridium difficile* colitis, aminotransferase elevations

Methemoglobinemia from primaquine, hemolysis in G6PDdeficient patients, leukopenia

See toxicities for TMP-SMX. Patients allergic to sulfa might tolerate dapsone. Methemoglobinemia, dose-related hemolysis, bone marrow suppression; rash; fever; nausea, abdominal pain; hyperkalemia; proteinuria papillary necrosis

Drug interactions Drug interactions with rifampin and rifabutin can render dapsone ineffective

Granulocytopenia, fever, rash; aminotransferase elevations

See above

Rash, drug fever; headaches; nausea, diarrhea, aminotransferase elevations; neutropenia, anemia; transient conjunctivitis; erythema multiforme Obtain accurate patient weight every 2–3 days to adjust pentamidine dosage. Discontinue pentamidine if creatinine > 3.0 mg/dL. Can resume administration if creatinine < 2 mg/dL

Comments

Consider in patients with mild-tomoderate PCP, intolerant of or unresponsive to TMP-SMX

Check G6PD before initiating primaquine therapy. Check methemoglobin levels when clinically indicated (see Dapsone). Vitamin C 1 g po tid might prevent methemoglobinemia. Lower dosage of primaquine (15 mg po qd) can be effective

Effective in mild-to-moderate PCP only. Check G6PD before starting dapsone. Check methemoglobin levels if symptomatic or discrepancy between oxygen saturation and simultaneous arterial PaO₂. Treat methemoglobinemia > 20% (15% if anemic or respiratory compromise) with methylene blue 1% solution 2 mg/kg IV once; methylene blue contraindicated in G6PD deficiency. Vitamin C 1 g po tid might prevent methemoglobinemia

Can be effective in some patients intolerant to or refractory to TMP-SMX therapy

Must be administered for 72 hours after the last dose of trimetrexate. Large IV fluid load with leucovorin administration can result in volume overload

Higher therapeutic failure rate than TMP-SMX. For patients who fail or are intolerant to TMP-SMX, pentamidine, dapsone-TMP, or clindamycinprimaquine. Take with high-fat diet to increase drug absorption. Patients with enteropathy might not absorb a sufficient amount of atovaquone to treat adequately

Continued

System, Problem, and Drug Regimen

Adverse Effects/Drug Interactions

Comments

PULMONARY, PCP (cont.)

Duration

6-8 weeks for

acute therapy

Same

Same

Same

Adjunctive corticosteroid therapy for acute PCP with $PaO_2 \leq 70 \, mmHg$ Prednisone po or 21 days methylprednisolone (Solu-Medrol) IV: 40 mg bid for 5 days followed by 40 mg qd for 5 days, followed by 20 mg qd for 11 days (can be tapered to zero for last 11 days also)

CENTRAL NERVOUS SYSTEM

Toxoplasma gondii Prophylaxis Most PCP Indefinitely prophylaxis regimens provide some protection against toxoplasmosis

Hyperglycemia, sodium retention, potassium wasting. Psychiatric syndromes. Exacerbation of Kaposi sarcoma, thrush, herpes infections, and other opportunistic infections

Corticosteroids indicated in conjunction with antipneumocystis therapy in all patients with $PaO_2 \leq 70$ mmHg. Begin corticosteroids concurrent with PCP treatment or if PaO2 decreases to \leq 70 mmHg within 72 hours of initiating PCP treatment

See PULMONARY, PCP

Leukopenia, anemia,

thrombocytopenia

TMP-SMX, dapsone plus TMP or pyrimethamine, clindamycin plus primaquine, atovaquone plus pyrimethamine, and pyrimethamine-sulfadoxine provide some prophylaxis against toxoplasmosis. Other PCP regimens (eg, aerosolized pentamidine) not effective; adding another agent to provide toxoplasmosis prophylaxis not required. Clarithromycin and azithromycin provide some benefit

Clinical response or regression of lesions on imaging studies is usually noted within 2 weeks. Maintenance required indefinitely to prevent relapse

Sulfadiazine probably provides effective prophylaxis and suppression against PCP

Pyrimethamine 75–100 mg po qd (every other day if bone marrow suppression) plus leucovorin calcium (folinic acid) 10-25 mg po qd plus either

Acute

Sulfadiazine 1.0-1.5 g po q 6 h

or

Clindamycin 600-900 mg po or IV gid

Alternative when intolerant of sulfadiazine and clindamycin Pyrimethamine plus leucovorin as above

plus one of the following

Clarithromycin Same 1 g po bid or azithromycin 1200-1500 mg po qd

or

Rash, drug fever; bone marrow

See PULMONARY, PCP

suppression, leukopenia,

See above

See GENERAL/SYSTEMIC, MAC

Continued

Table 1. Continued System, Problem, and

Drug Regimen	Duration	Adverse Effects/Drug Interactions	Comments
CENTRAL NERVOU	S SYSTEM		a de la companya de Este de la companya de
Alternatives (cont.)			
Atovaquone	Same	See PULMONARY, PCP	Not proved effective
(750 mg/5 mL) 750 mg po qid with meals			
or			
Doxycycline 100 mg po tid-qid or mino-	Same	Tetracycline side effects	Not proved effective
cycline 200 mg po bid			
or	-		
Dapsone 100 mg po qd OR	Same	See PULMONARY, PCP	
Pyrimethamine alone 100–200 mg po qd	Same	See PULMONARY, PCP	Not as effective as above regimens
OR			
TMP/SMX as for acute PCP	Same	See PULMONARY, PCP	
<i>Maintenance</i> Pyrimethamine 25–50 mg po qd	Indefinitely		Add leucovorin calcium if evidence of leukopenia
plus either	4		
Sulfadiazine 1 g po q 12 h	Indefinitely		Other agents used for acute toxoplasmosis might be effective at lower dosage for maintenance
Or			
Clindamycin 300–450 mg po q 6 h	Indefinitely		
Cryptococcus neoforman Probhylaxis	IJ		
Fluconazole provides			Primary prophylaxis not routinely
limited prophylaxis		a shekara nga nga nga nga nga nga nga nga nga ng	recommended. Can be considered for
			No long term survival hopeft
			Fluconazole resistance reported
Acute meningitis or disseminated		an an an an an Araba an Araba. An an Araba an Araba an Araba an Araba	
Amphotericin B 0.7–1.0 mg/kg/d IV with or without 5-flucytosine	6–8 weeks; amphotericin total dosage not to exceed	Renal failure, hypokalemia, hypomagnesemia; fever, chills; anemia, thrombophlebitis	Pretreatment with diphenhydramine, acetaminophen, or IV meperidine can de- crease amphotericin-induced fevers, chills, and rigors. Administer for 4–6 h in D5W.
(Ancobon) 100 mg/kg po qd in 4 divided doses for first 2–4 weeks If clinically im-	2 g	Granulocytopenia; nausea, vomiting diarrhea, aminotransferase elevations; rash from flucytosine	Addition of heparin 500 U and hydrocor- tisone 50 mg to amphotericin IV solution can decrease phlebitis. Infusion of 500 mL- 1 L. normal saline before administration of
proved after 7.5 mg/kg total amphophotericin B administration, can		Flucytosine toxicities (rash, leukopenia), in absence of clear benefits, limit its use	amphotericin B can minimize renal toxicity. 5-Flucytosine not indicated if granulo- cytopenia or thrombocytopenia is present

Markedly increased intracranial pressure (> 300 mm) might require acetazolamide (Diamox) 250–500 mg po or IV qid, cerebrospinal fluid drainage (15 mL or more per day) or possibly corticosteroids or mannitol therapy

Continued

OR

po bid

change to fluconazole

400 mg po qd or itraconazole 200 mg

System, Problem, and Drug Regimen	Duration	Adverse Effects/Drug Interactions	Comments
CENTRAL NERVOU <i>Cryptococcus neoforman</i> Fluconazole 400 mg po qd	JS SYSTEM 15 (cont.) 8–12 weeks	Nausea, vomiting, diarrhea; dizziness; aminotransferase elevations; rare cutaneous reactions <i>Drug interactions</i> Increased phenytoin (Dilantin) and warfarin (Coumadin) levels; higher fluconazole dosages might be neces- sary for patients taking rifampin	As effective as amphotericin B against mild or moderate disease; unknown whether equally effective against severe disease. Higher dosages (eg, 800–1200 mg po qd) might be necessary in severe disease. Fluconazole penetrates central nervous system and most body tissues, including prostate
<i>Maintenance</i> Fluconazole 200–400 mg po qd	Indefinitely	Same	Higher dosages might be necessary for recurrent disease
OR		ter an	a de la companya de l A companya de la comp
Amphotericin B 0.5–0.8 mg/kg/d 3–5 tímes a week	Indefinitely	Same	• •
Syphilis Aqueous crystalline penicillin G 2–4 mU IV q 4 h (total 12–24 mU/d)	10–14 days	Usual penicillin adverse effects; Jarisch-Herxheimer reaction; seizures from high-dosage penicillin in renal failure	Continued serologic and clinical follow- up required to assess adequacy of treat- ment. Persons with ophthalmic, auditory or cranial nerve abnormalities or other syndromes consistent with neurosynhilis
OR Procaine penicillin	10–14 days	Same. Probenecid rash	should receive daily penicillin therapy for 10–14 days. Intravenous penicillin preferred for adequate central nervous
G 2.4 mU IM qd plus Probenecid 500 mg po qid			system penetration. For penicillin- allergic patients, consultation with an expert advised. Administer additional benzathine penicillin 2.4 mµ IM weekly after completion of neurosyphilis treat- ment to ensure 3 weeks total penicillin therapy
Peripheral neuropathy Amitriptyline (Elavil) or desipramine (Norpramin) 25-150 mg po hs	Indefinitely	Usual tricyclic side effects; drowsi- ness; orthostatic hypotension; anticholinergic symptoms	Pain relief occurs in 3–5 days. Desipramine causes less sedation and fewer anticholinergic effects. Other tricyclic drugs might be equally effective
Phenytoin (diphenylhydantoin, Dilantin) 100 mg po tid	Indefinitely	Usual side effects and drug-drug interactions	Generally ineffective
Carbamazepine (Tegretol) 100-300 mg po bid	Indefinitely	Leukopenia, bone marrow suppression, rare agranulocytosis; rash; drowsiness, dizziness; aminotransferase elevations	Less desirable because of bone marrow effects. Need to monitor carbamazepine levels to avoid toxicity
Mexiletine (Mexitil) 150 mg po bid-tid	Indefinitely	Nausea, vomiting, epigastric pain; dizziness, tremor; bradycardia; rare seizures, leukopenia, agranulocytosis	Less desirable because of side effects
Capsaicin (Axsain, Zostrix-HP) 0.075% topical cream qid	Indefinitely	Minor burning sensation, skin irritation, erythema	Pain relief delayed 2–4 weeks. No systemic effects

J Am Board Fam Pract: first published as 10.3122/jabfm.9.2.125 on 1 March 1996. Downloaded from http://www.jabfm.org/ on 5 May 2025 by guest. Protected by copyright.

Viral Load Measurements

HIV viral loads are correlated with disease progression. Many clinicians are using viral load measurements to monitor antiretroviral efficacy or development of drug resistance. Although reduction of viral load has been observed after starting or changing antiretroviral therapies, studies have not adequately correlated these changes with clinical outcomes. It is unclear whether the benefits of viral load measurements justify the routine use of this expensive (\$200) test. We believe that monitoring the patient's clinical course and CD4+ cell counts remains critical.

Prophylaxis and Treatment of Opportunistic Infections

New guidelines for prevention of opportunistic infections have been published by the Centers for Disease Control and Prevention (CDC).¹⁷ These guidelines are available from the CDC National AIDS Clearinghouse at 1-800-458-5231.

Pneumocystis carinii pneumonia (PCP) remains the single most important opportunistic infection in AIDS because of its frequency, its substantial morbidity and mortality, and its susceptibility to prophylaxis and treatment.^{18,19} PCP prophylaxis should be given to all persons with CDC-defined AIDS (including a CD4+ cell count of less than 200/µL on two occasions) and other manifestations of advanced immunodeficiency, such as recurrent oral candidiasis and persistent unexplained fever greater than 100°F for 2 or more weeks.

Toxoplasmosis prophylaxis, recommended by the CDC for patients with CD4+ cell counts less than 100/ μ L, is achieved with most PCP prophylaxis regimens except dapsone alone and aerosolized pentamidine.²⁰⁻²² For those receiving dapsone alone or pentamidine, we recommend treating toxoplasmosis in the small number of patients who develop it rather than administering specific prophylaxis.

Mycobacterium avium complex (MAC) prophylaxis is recommended by the CDC for persons with a CD4+ cell count of less than $75/\mu$ L.^{17,23-25} We do not consider MAC prophylaxis an essential part of HIV primary care.^{26,27} An alternative strategy is to treat clinical MAC disease if it occurs. Similarly, because of the expense and lack of clear benefit, prophylaxis against candidal and other fungal diseases, as well as prophylaxis against cytomegalovirus retinitis, is not recommended.

The Table

Table 1 gives our recommendations for treating most specific diseases and major symptoms of HIV/AIDS. The recommendations are principally in order of efficacy. Where more than one drug is effective for a specific condition, recommendations are usually in an order that favors the least expensive choice among equally effective options. The most common and clinically important adverse effects and drug interactions are listed.

References

A selected bibliography highlights the most important management and therapeutic problems in HIV/AIDS. References including articles about pulmonary disease,^{18-22,28-35} herpesvirus infections,³⁶⁻⁴⁴ dermatologic problems,⁴⁵⁻⁴⁸ the AIDS wasting syndrome,⁴⁹⁻⁵³ diarrhea,⁵⁴⁻⁵⁷ neurologic disease,^{44,58-69} tuberculosis,⁷⁰⁻⁷² and other mycobacterial ^{23-27,73,74} and fungal diseases⁷⁵⁻⁸⁰ are included. Additional references are intended to assist providers with drug reactions^{33,34,81-83} and prevention^{84,85} including special considerations in pregnancy,^{86,87} and for health care workers sustaining percutaneous exposure to blood.⁸⁸

Other Sources of Information

To assist clinicians in providing HIV care, many local, regional, state, university, and national information services are available. Our national HIV Telephone Consultation Service (Warmline) in the University of California, San Francisco, Department of Family and Community Medicine at San Francisco General Hospital provides clinical consultation and education for health care providers; the Warmline is in operation on weekdays at 1-800-933-3413. Information about clinical trials is available through the AIDS Clinical Trials Information Service of the CDC and the National Institutes of Allergy and Infectious Diseases at 1-800-TRIALS A and through the AIDS Treatment Information Service (ATIS) at 1-800-HIV-8440, which also has printed guidelines and information about approved therapies and management protocols. The AIDS Education and Training Centers (AIDS ETCs) of the Health Resources and Services Administration (HRSA) at 1-301-443-6364 offers education, training, and consultation services to health care providers; HRSA also offers a bimonthly teleconference service.

Conclusion

For the family physician and other clinicians, key elements of HIV care are antiretroviral therapy, prophylaxis against opportunistic infections, and treatment of acute complications of AIDS. The importance of the provider-patient-family relationship in providing this care cannot be underestimated. The Current Report - HIV treatment guidelines are intended to help family physicians and other primary care clinicians provide interventions that will delay or prevent many of the complications of HIV/AIDS.

We gratefully acknowledge the staff of the HIV Telephone Consultation Service and the faculty and staff at San Francisco General Hospital for making this work possible, and Mary A. Hanville and Jill J. Legg, MD, for assistance in preparing this manuscript.

References

- 1. Goldschmidt RH, Dong BJ. Treatment of AIDS and HIV-related conditions - 1995. J Am Board Fam Pract 1995;8:139-62.
- Volberding PA, Lagakos SW, Grimes JM, Stein DS, Rooney J, Meng T-C, et al. A comparison of immediate with deferred zidovudine therapy for asymptomatic HIV-infected adults with CD4 cell counts of 500 or more per cubic millimeter. N Engl J Med 1995;333:401-7.
- Lundgren JD, Phillips AN, Pedersen C, Clumeck N, Gatell JM, Johnson AM, et al. Comparison of longterm prognosis of patients with AIDS treated and not treated with zidovudine. JAMA 1994;271:1088-92.
- Concorde: MRC/ANRS randomised double-blind controlled trial of immediate and deferred zidovudine in symptom-free HIV infection. Concorde Coordinating Committee. Lancet 1994;343:871-81.
- Volberding PA, Lagakos SW, Grimes JM, Stein DS, Balfour HH Jr, Reichman RC, et al. The duration of zidovudine benefit in persons with asymptomatic HIV infection. Prolonged evaluation of protocol 019 of the AIDS Clinical Trials Group. JAMA 1994;272: 437-42.
- Ioannidis JP, Cappelleri JC, Lau J, Skolnik PR, Melville B, Chalmers TC, et al. Early or deferred zidovudine therapy in HIV-infected patients without an AIDSdefining illness. Ann Intern Med 1995;122:856-66.
- 7. Moore RD, Chaisson RE, Hidalgo J. The efficacy of zidovudine is time limited. JAMA 1994;272:1001.
- Spooner KM, Lane HC, Masur H. Antiretroviral therapy: reference guide to major clinical trials in patients infected with human immunodeficiency virus. Clin Infect Dis 1995;20:1145-51.
- 9. Lipsky JJ. Zalcitabine and didanosine. Lancet 1993;341:30-2.
- 10. Montaner JS, Schechter MT, Rachlis A, Gill J, Beaulieu R, Tsoukas C, et al. Didanosine compared with continued zidovudine therapy for HIV-infected pa-

tients with 200 to 500 CD4 cells/mm³. A doubleblind, randomized, controlled trial. Ann Intern Med 1995;123:561-71.

- Eron JJ, Benoit SL, Jemsek J, MacArthur RD, Santana J, Quinn JB, et al. Treatment with lamivudine, zidovudine, or both in HIV-positive patients with 200 to 500 CD4+ cells per cubic millimeter. N Engl J. Med 1995;333:1662-9.
- 12. Lipsky JJ. The glimmer of HIV proteinase inhibitors. Lancet 1995;345:936-7.
- 13. Hammer S, Katzenstein D, Hughes M, Gundacker H, Hirsch M, and Merigan T for the ACTG 175 Study Team NIAID Sponsored AIDS Clinical Trials Group. Nucleoside monotherapy (MT) vs combination therapy (CT) in HIV infected adults: a randomized double-blind, placebo-controlled trial in persons with CD4 cell counts 200-500/mm³. In: Program and abstracts of the 35th Interscience Conference on Antimicrobial Agents and Chemotherapy, San Francisco, September 17-20, 1995. Washington, DC: American Society for Microbiology, 1995:8. Abstract.
- 14. Choo V. Combination superior to zidovudine in Delta trial. Lancet 1995;346:895.
- Sande MA, Carpenter CC, Cobbs CG, Holmes KK, Sanford JP. Antiretroviral therapy for adult HIV-infected patients. Recommendations from a state-ofthe-art conference. JAMA 1993;270:2583-9.
- Goldschmidt RH, Dong BJ, Legg JJ. Antiretroviral strategies revisited. J Am Board Fam Pract 1995;8: 62-9.
- Centers for Disease Control and Prevention. USPHS/IDSA guidelines for the prevention of opportunistic infections in persons infected with human immunodeficiency virus: a summary. MMWR 1995; 44 (RR-8):1-34.
- Centers for Disease Control. Recommendations for prophylaxis against *Pneumocystis carinii* pneumonia for adults and adolescents infected with human immunodeficiency virus. MMWR 1992;41(RR-4):1-11.
- 19. Masur H. Prevention and treatment of *Pneumocystis* pneumonia. N Engl J Med 1992;328:1853-60.
- Heald A, Flepp M, Chave JP, Malinverni R, Rüttimann S, Gabriel V, et al. Treatment for cerebral toxoplasmosis protects against *Pneumocystis carinii* pneumonia in patients with AIDS. The Swiss HIV Cohort Study. Ann Intern Med 1991;115:760-3.
- Podzamczer D, Salazar A, Jiménez J, Consiglio E, Santín M, Casanova A, et al. Intermittent trimethoprim-sulfamethoxazole compared with dapsonepyrimethamine for the simultaneous primary prophylaxis of *Pneumocystis* pneumonia and toxoplasmosis in patients infected with HIV. Ann Intern Med 1995; 122:755-61.
- 22. Carr A, Tindall B, Brew BJ, Marriott DJ, Harkness JL, Penny R, et al. Low-dose trimethoprim-sulfamethoxazole prophylaxis for toxoplasmic encephalitis in patients with AIDS. Ann Intern Med 1992; 117:106-11.
- 23. Masur H. Recommendations on prophylaxis and therapy for disseminated *Mycobacterium avium* complex disease in patients infected with the human immunodeficiency virus. Public Health Service Task Force on Prophylaxis and Therapy for *Mycobacterium avium* Complex. N Engl J Med 1993;329:898-904.

- 24. Centers for Disease Control and Prevention. Recommendations on prophylaxis and therapy for disseminated *Mycobacterium avium* complex for adults and adolescents infected with human immunodeficiency virus. US. Public Health Service Task Force on Prophylaxis and Therapy for *Mycobacterium Avium* Complex. MMWR 1993;42(RR-9):14-20.
- Nightingale SD, Cameron DW, Gordin FM, Sullam PM, Cohn DL, Chaisson RE, et al. Two controlled trials of rifabutin prophylaxis against *Mycobacterium* avium complex infection in AIDS. N Engl J Med 1993;329: 828-33.
- Goldschmidt RH, Dong BJ. Rifabutin prophylaxis against Mycobacterium avium complex disease. J Am Board Fam Pract 1994;7:58-61.
- 27. Goldschmidt RH, Hearst N, Chambers DB. Rifabutin prophylaxis against *Mycobacterium avium* complex infection. N Engl J Med 1994;330:436-7.
- Consensus statement on the use of corticosteroids as adjunctive therapy for *Pneumocystis* pneumonia in the acquired immunodeficiency syndrome. The National Institutes of Health–University of California Expert Panel for Corticosteroids as Adjunctive Therapy for *Pneumocystis* Pneumonia. N Engl J Med 1990; 323:1500-4.
- 29. Black JR, Feinberg J, Murphy RL, Fass RJ, Finkelstein D, Akil B, et al. Clindamycin and primaquine therapy for mild-to-moderate episodes of *Pneumocystis carinii* pneumonia in patients with AIDS: AIDS Clinical Trials Group 044. Clin Infect Dis 1994;18:905-13.
- Bozzette SA, Finkelstein DM, Spector SA, Frame P, Powderly WG, He W, et al. A randomized trial of three antipneumocystis agents in patients with advanced human immunodeficiency virus infection. N Engl J Med 1995;332:693-9.
- Hughes W, Leoung G, Kramer F, Bozzette SA, Safrin S, Frame P, et al. Comparison of atovaquone (566-C80) with trimethoprim-sulfamethoxazole to treat *Pneumocystis carinii* pneumonia in patients with AIDS. N Engl J Med 1993;328:1521-7.
- 32. Sattler FR, Frame P, Davis R, Nichols L, Shelton B, Akil B, et al. Trimetrexate with leucovorin versus trimethoprim-sulfamethoxazole for moderate to severe episodes of *Pneumocystis carinii* pneumonia in patients with AIDS: a prospective, controlled multicenter investigation of the AIDS Clinical Trials Group protocol 029/031. J Infect Dis 1994;170:165-72.
- Jung AC, Paauw DS. Management of adverse reactions to trimethoprim-sulfamethoxazole in human immunodeficiency virus-infected patients. Arch Intern Med 1994;154:2402-6.
- Gluckstein D, Ruskin J. Rapid oral desensitization to trimethoprim-sulfamethoxazole (TMP-SMZ): use in prophylaxis for *Pneumocystis carinii* pneumonia in patients with AIDS who were previously intolerant to TMP-SMZ. Clin Infect Dis 1995;20:849-53.
- Hirschtick RE, Glassroth J, Jordan MC, Wilcosky TC, Wallace JM, Kvale PA, et al. Bacterial pneumonia in persons infected with the human immunodeficiency virus. N Engl J Med 1995;333:845-51.
- 36. Safrin S, Crumpacker C, Chatis P, Davis R, Hafner R, Rush J, et al. A controlled trial comparing foscarnet with vidarabine for acyclovir-resistant mucocutaneous herpes simplex in the acquired immunodeficiency syndrome. N Engl J Med 1991;325:551-5.

- Balfour HH Jr, Benson C, Braun J, Cassens B, Erice A, Friedman-Kien A, et al. Management of acyclovir-resistant herpes simplex and varicella-zoster virus infections. J Acquir Immune Defic Syndr 1994;7:254-60.
- 38. Mortality in patients with the acquired immunodeficiency syndrome treated with either foscarnet or ganciclovir for cytomegalovirus retinitis. Studies of Ocular Complications of AIDS Research Group, in collaboration with the AIDS Clinical Trials Group. N Engl J Med 1992;326:213-20.
- Dieterich DT, Poles MA, Lew EA, Mendez PE, Murphy R, Addessi A, et al. Concurrent use of ganciclovir and foscarnet to treat cytomegalovirus infection in AIDS patients. J Infect Dis 1993;167:1184-8.
- 40. Intravenous versus oral ganciclovir: European/Australian comparative study of efficacy and safety in the prevention of cytomegalovirus retinitis recurrence in patients with AIDS. The Oral Ganciclovir European and Australian Cooperative Study Group. AIDS 1995;9:471-7.
- 41. Drew WL, Ives D, Lalezari JP, Crumpacker C, Follansbee SE, Spector SA, et al. Oral ganciclovir as maintenance treatment for cytomegalovirus retinitis in patients with AIDS. N Engl J Med 1995;333:615-20.
- Holland GN, Tufail A. New therapies for cytomegalovirus retinitis. N Engl J Med 1995;333:658-9.
- Ernest JT. Intraocular device for cytomegalovirus infection. Lancet 1995;346:983-4.
- 44. Kim YS, Hollander H. Polyradiculopathy due to cytomegalovirus: report of two cases in which improvement occurred after prolonged therapy and review of the literature. Clin Infect Dis 1993;17:32-7.
- Berger TG, Obuch ML, Goldschmidt RH. Dermatologic manifestations of HIV infection. Am Fam Physician 1990;41:1729-42.
- Cohen PR, Grossman ME. Recognizing skin lesions of systemic fungal infections in patients with AIDS. Am Fam Physician 1994;49:1627-34.
- Adal KA, Cockerell CJ, Petri WA Jr. Cat scratch disease, bacillary angiomatosis, and other infections due to *Rochalimaea*. N Engl J Med 1994;330:1509-15.
- Koehler JE, Tappero JW. Bacillary angiomatosis and bacillary peliosis in patients infected with human immunodeficiency virus. Clin Infect Dis 1993; 17:612-24.
- Macallan DC, Noble C, Baldwin C, Jebb SA, Prentice AM, Coward WA, et al. Energy expenditure and wasting in human immunodeficiency virus infection. N Engl J Med 1995;333:83-8.
- Von Roenn JH, Armstrong D, Kotler DP, Cohn DL, Klimas NG, Tchekmedyian NS, et al. Megestrol acetate in patients with AIDS-related cachexia. Ann Intern Med 1994;121:393-9.
- 51. Oster MH, Enders SR, Samuels SJ, Cone LA, Hooton TM, Browder HP, et al. Megestrol acetate in patients with AIDS and cachexia. Ann Intern Med 1994;121:400-8.
- 52. Grunfeld C, Feingold KR. Metabolic disturbances and wasting in the acquired immunodeficiency syndrome. N Engl J Med 1992;327:329-37.
- 53. Mulligan K, Grunfeld C, Hellerstein MK, Neese RA, Schambelan M. Anabolic effects of recombinant human growth hormone in patients with wasting associated with human immunodeficiency virus infection. J Clin Endocrinol Metab 1993;77:956-62.

Treatment of AIDS 147

J Am Board Fam Pract: first published as 10.3122/jabfm.9.2.125 on 1 March 1996. Downloaded from http://www.jabfm.org/ on 5 May 2025 by guest. Protected by copyright.

- 54. DuPont HL, Marshall GD. HIV-associated diarrhoea and wasting. Lancet 1995;346:352-6.
- 55. White AC Jr, Chappell CL, Hayat CS, Kimball KT, Flanigan TP, Goodgame RW. Paromomycin for cryptosporidiosis in AIDS: a prospective, doubleblind trial. J Infect Dis 1994;170:419-24.
- Simon DM, Cello JP, Valenzuela J, Levy R, Dickerson G, Goodgame R, et al. Multicenter trial of octreotide in patients with refractory acquired immunodeficiency syndrome-associated diarrhea. Gastroenterology 1995;108:1753-60.
- 57. Blanshard C, Benhamou Y, Dohin E, Lernestedt JO, Gazzard BG, Katlama C. Treatment of AIDS-associated gastrointestinal cytomegalovirus infection with foscarnet and ganciclovir: a randomized comparison. J Infect Dis 1995;172:622-8.
- Simpson DM, Tagliati M. Neurologic manifestations of HIV infection. Ann Intern Med 1994;121:769-85.
- Newton HB. Common neurologic complications of HIV-1 infection and AIDS. Am Fam Physician 1995; 51: 387-98.
- Luft BJ, Hafner R, Korzun AH, Leport C, Antoniskis D, Bosler EM, et al. Toxoplasmic encephalitis in patients with the acquired immunodeficiency syndrome. N Engl J Med 1993;329:995-1000.
- 61. Kovacs JA. Toxoplasmosis in AIDS: keeping the lid on. Ann Intern Med 1995;123:230-1.
- 62. Powderly WG. Cryptococcal meningitis and AIDS. Clin Infect Dis 1993;17:837-42.
- 63. Saag MS, Powderly WG, Cloud GA, Robinson P, Grieco MH, Sharkey PK, et al. Comparison of amphotericin B with fluconazole in the treatment of acute AIDS-associated cryptococcal meningitis. N Engl J Med 1992;326:83-9.
- 64. Powderly WG, Saag MS, Cloud GA, Robinson P, Meyer RD, Jacobson JM, et al. A controlled trial of fluconazole or amphotericin B to prevent relapse of cryptococcal meningitis in patients with the acquired immunodeficiency syndrome. N Engl J Med 1992; 326:793-8.
- 65. Haubrich RH, Haghighat D, Bozzette SA, Tilles J, Mc-Cutchan JA, and the California Collaborative Treatment Group. High-dose fluconazole for treatment of cryptococcal disease in patients with human immunodeficiency virus infection. J Infect Dis 1994; 170:238-42.
- Hook EW 3d, Marra CM. Acquired syphilis in adults. N Engl J Med 1992;326:1060-9.
- 67. Gordon SM, Eaton ME, George R, Larsen S, Lukehart SA, Kuypers J, et al. The response of symptomatic neurosyphilis to high-dose intravenous penicillin G in patients with human immunodeficiency virus infection. N Engl J Med 1994;331:1469-73.
- 68. Malone JL, Wallace MR, Hendrick BB, LaRocco A Jr, Tonon E, Brodine SK, et al. Syphilis and neurosyphilis in a human immunodeficiency virus type-1 seropositive population: evidence for frequent serologic relapse after therapy. Am J Med 1995;99:55-63.
- 69. Centers for Disease Control and Prevention. 1993 sexually transmitted diseases treatment guidelines. MMWR 1993;42(RR-14):1-102.
- 70. Centers for Disease Control. The use of preventive therapy for tuberculosis infection in the United States. Recommendations of the Advisory Committee for Elimination of Tuberculosis. MMWR 1990;39 (RR-8):9-12.

- Centers for Disease Control. Initial therapy for tuberculosis in the era of multidrug resistance. Recommendations of the Advisory Council for the Elimination of Tuberculosis. MMWR 1993;42 (RR-7):1-8.
- 72. Iseman MD. Treatment of multidrug-resistant tuberculosis. N Engl J Med 1993;329:784-91.
- Horsburgh CR Jr. Mycobacterium avium complex infection in the acquired immunodeficiency syndrome. N Engl J Med 1991;324:1332-8.
- 74. Wolinsky E. Mycobacterial diseases other than tuberculosis. Clin Infect Dis 1992;15:1-10.
- 75. Diamond RD. The growing problem of mycoses in patients infected with the human immunodeficiency virus. Rev Infect Dis 1991;13:480-6.
- 76. Fungal infection in HIV-infected persons. American Thoracic Society. Am J Respir Crit Care Med 1995;152:816-22.
- 77. Como JA, Dismukes WE. Oral azole drugs as systemic antifungal therapy. N Engl J Med 1994;330:263-72.
- 78. Powderly WG, Finkelstein DM, Feinberg J, Frame P, He W, van der Horst C, et al. A randomized trial comparing fluconazole with clotrimazole troches for the prevention of fungal infections in patients with advanced human immunodeficiency virus infection. N Engl J Med 1995;332:700-5.
- 79. Wheat J, Hafner R, Korzun AH, Limjoco MT, Spencer P, Larsen RA, et al. Itraconazole treatment of disseminated histoplasmosis in patients with the acquired immunodeficiency syndrome. Am J Med 1995; 98:336-42.
- 80. Wheat J, Hafner R, Wulfsohn M, Spencer P, Squires K, Powderly W, et al. Prevention of relapse of histoplasmosis with itraconazole in patients with the acquired immunodeficiency syndrome.
- Lee BL, Safrin S. Interactions and toxicities of drugs used in patients with AIDS. Clin Infect Dis 1992; 14:773-9.
- Coopman SA, Johnson RA, Platt R, Stern RS. Cutaneous disease and drug reactions in HIV infection. N Engl J Med 1993;328:1670-4.
- White MV, Haddad ZH, Brunner E, Sainz C. Desensitization to trimethoprim sulfamethoxazole in patients with acquired immune deficiency syndrome and *Pneu*mocystis carinii pneumonia. Ann Allergy 1989;62:177-9.
- Makadon HJ, Silin JG. Prevention of HIV infection in primary care: current practices, future possibilities. Ann Intern Med 1995;123:715-9.
- 85. Gabel LL, Crane R, Ostrow DC. HIV-related disease: family physicians' multiple opportunities for preventive intervention. J Am Board Fam Pract 1994:7:218-24.
- 86. Centers for Disease Control and Prevention. Recommendations of the US Public Health Service Task Force on the use of zidovudine to reduce perinatal transmission of human immunodeficiency virus. MMWR 1994;43(RR-11):1-20.
- Connor EM, Sperling RS, Gelber R, Kiselev P, Scott G, O'Sullivan MJ, et al. Reduction of maternal-infant transmission of human immunodeficiency virus type 1 with zidovudine treatment. N Engl J Med 1994; 331:1173-80.
- Centers for Disease Control and Prevention. Casecontrol study of HIV seroconversion in health-care workers after percutaneous exposure to HIV-infected blood—France, United Kingdom, and United States, January 1988–August 1994. MMWR 1995;44:929-33.