Systemic reatment	Structure of biologic or target of oral systemic	Dosing for plaque psoriasis ^a	Efficacy at primary end point ^b	Safety considerations
Oral systemics				
Methotrexate	Dihydrofolate reductase inhibitor	Dose: 15-20 mg once weekly with folic acid supplementation	Adults: 36% achieve PASI 75 at week 16 (placebo, 19%) ⁶⁵	Increased risk of hepatic, pulmonary, hematologic, and renal toxicity
			Subcutaneous methotrexate may confer greater efficacy and bioavailability ⁸⁵	Check complete blood counts, liver and kidney function, and hepatitis serology
Apremilast	Phosphodiesterase-4 inhibitor	Day 1: 10 mg in the morning	Adults: 33% achieve PASI 75 at week 16 (placebo, 5%) ⁸⁶	Gastrointestinal disturbances in 15%-20% and weight loss
		Day 2: 10 mg twice per day		Renal adjustment for creatinine clearance <30 mL/min/1.73m ²
		Day 3: 10 mg in the morning; 20 mg in the evening		
		Day 4: 20 mg twice per day		
		Day 5: 20 mg in the morning; 30 mg in the evening		
		Day 6 onward (maintenance dose): 30 mg twice per day		
Acitretin	Retinoid (vitamin-A derivative)	Dose: 25 mg-50 mg daily	Adults: 47% achieve PASI 75 at week 12 ⁸⁷	Hair loss and xerosis (worse at higher doses), hypertriglycer- idemia, hepatotoxicity, and teratogenicity
				Check liver function and lipids
Cyclosporine	Calcineurin inhibitor	Dose: 5 mg/kg as a twice-daily divided dose; taper when psoriasis is controlled	Adults: 65% achieve IGA 0/1 at week 8 (placebo, 0%) ⁸⁸ Two formulations are available: lipid or microemulsified (the latter is better absorbed)	Nephrotoxicity with long-term use
				Hypertension and hyperlipidemia
				nonmelanoma skin cancer risk in patients with prior PUVA
				Check renal function levels, electrolytes (magnesium and potassium), and blood pressure
	. 0	ment scale; PASI, Psoriasis Area Severity Index; PUVA, pso		ns to assess psoriasis disease severity. It combines the assessment of
	mor necrosis factor.			nd scale) and the affected areas into a single score between 0 (no cores of 75, 90, and 100 indicate a 75%, 90%, and 100% reduction i
All doses are for	adults unless otherwise specif	fied.	PASI score compared with baseline.	cores of 75, 90, and 100 indicate a 75%, 90%, and 100% reduction i