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Managing Clozapine Side Effects				
Side Effect	Incidence	Monitoring	Treatment	
Bradycardia Orthostatic hypotension Syncope	9%-13%	PulseBlood pressureOrthostatic blood pressureDizziness	 Divided dose (2–3 times daily) Start with low dose Slow titration Adequate fluid intake 	 Behavioral changes (sit on edge of the bed with legs hanging down for a minute before getting up) Fludrocortisone, if necessary
Constipation	15%-25%	Clinical interview	 Drink fluids Increase fiber Exercise	 Minimize other anticholinergics (eg, benztropine, diphenhydramine) Consider bowel regimen, similar to opioids (eg, docusate 100 mg-250 mg BID)
Drowsiness	Up to 45%	Clinical interview	Bedtime dosing	Lowest dose possible
Hyperglycemia	27% increased to ≥ 126 mg/dL	Baseline and quarter- ly fasting glucose or HbA1c	Dietary intervention Exercise	• If A1c > 6.5/7%, add metformin (6.5 for younger people with long life expectancy)
	50% mean increase of ≥ 71 mg/dL	Baseline, week 12, and annually	Dietary intervention Exercise	Omega-3 fatty acidsFibrates (caution with statins)
Myocarditis Cardiomyopathy	0.002%	• Troponin & CRP weekly for first 4–8 weeks	• Discontinue if CRP > 100 mg/L	• Troponin ≥ 2 x ULN
Seizures	2.4%-4.2%	Clinical interview/ contact from ER or hospital	• Use lowest dose possible (< 600 mg ideally)	Concomitant valproate
Sialorrhea	15%-40%	Clinical interview	Glycopyrronium 2–4 mg at bed- time	Botulinum toxin
Tachycardia	15%-25%	• Pulse	• Lower dose	Beta blockers (eg, metoprolol)
Weight gain ≥ 7% from base- line	35%	• Baseline weight/ BMI at weeks 4, 8, 12 and quarterly	Dietary intervention Exercise	If gain > 5 lbs in first month or > 10 lbs from baseline, consider metformin