

Migraine Prophylaxis

Background

Migraine headache (MH) is a common, debilitating neurologic disorder that affects approximately 18% of women and 6% of men in the United States.^{1,2} Several medications have been developed with established efficacy as abortive agents in MH; however, they do not prevent future headaches and frequent or excessive use can result in medication-overuse headaches. To avoid the development of this condition most experts recommend limiting the use of acute medications to no more than 2 days per week.^{3,4}

Although their mechanism of action is incompletely understood and tolerability is sometimes an issue, a trial of prophylactic therapy is recommended for patients with severe and/or recurrent headaches as it has been shown to reduce MH frequency by as much as 50%. Despite this, it is estimated that only 30% of patients utilize prophylactic therapies who might benefit from them.^{5,6}

Key Points

- Prophylactic therapy should be considered in the following circumstances:
 - 1) Patients who suffer two or more attacks per month that produce disability lasting 3 or more days per month
 - 2) Patients with contraindications to, or who have failed acute therapies
 - 3) Use of acute medications more than twice per week
 - 4) The presence of uncommon migraine conditions, including hemiplegic migraine, migraine with prolonged aura, or migrainous infarction
- Drugs used as prophylactic agents include beta-blockers, tricyclic antidepressants, anti-seizure medications, NSAIDs, calcium channel blockers, and angiotensin blockade agents. Other agents in use include feverfew, magnesium, hormone therapy, SSRIs and riboflavin
- **Agents with proven efficacy and considered first-line include propranolol, timolol, amitriptyline, divalproex sodium, sodium valproate, and topiramate^{5,7}**
- Start with lower doses and titrate slowly; long-acting formulations are helpful for improving adherence; a trial of at least 2 months of consistent use is necessary to establish efficacy
- When possible, patient comorbidities should be taken into account when selecting a prophylactic agent

Pharmacologic Agent Comparison

DRUG	TYPICAL DOSE	ADVERSE EFFECTS	COST*
<i>First-line Agents</i>			
Amitriptyline (generic)	10-150 mg per day	Drowsiness, weight gain, anticholinergic symptoms	\$3.66 [†]
Divalproex Sodium (Depakote®, Depakote® ER)	500-1000 mg per day	Sedation, nausea, weight gain, hair loss, tremor, liver toxicity	\$148.99
Propranolol or propranolol LA (generic)	80-240 mg per day	Fatigue, exercise intolerance, weight gain, dizziness, insomnia, depression	\$13.99 [†]
Timolol (generic)	20-30 mg per day	Fatigue, exercise intolerance, weight gain, dizziness, insomnia, depression	\$28.60 [†]
Topiramate (Topamax®)	50 mg twice daily	Paresthesia, fatigue, nausea, anorexia	\$212.99
<i>Second-line Agents</i>			
Gabapentin (generic)	1200-2400 mg per day	Dizziness and somnolence	\$89.99 [†]
Naproxen (generic)	550 mg twice daily	Nausea, vomiting, gastritis	\$23.98 [†]
Verapamil (generic)	240-320 mg per day	Dizziness, edema, constipation	\$53.98 [†]
Lisinopril (generic)	20 mg per day	Dizziness, cough	\$10.99 [†]
Candesartan (Atacand®)	16 mg per day	Dizziness, fatigue	\$49.99
Vitamin B ₂ (Riboflavin)	400 mg per day	Diarrhea, polyuria	\$8.99
Feverfew	100-250 mg per day	GI upset	\$6.39

*Monthly cost per www.drugstore.com for higher end of dosing range

[†]Generic available (cost provided)

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