

in collaboration with

Lifting The Burden

European principles of management of headache disorders in primary care (2nd edition)

7. Acute or symptomatic management of episodic migraine

General principles

- All **adults** with episodic migraine should have access to acute medication.
- **Children** with short-lasting attacks may respond well to bed-rest without medical treatment.
- In adults and children, *regular* use of acute medication at high frequency (on >2 days/week) risks the development of **medication-overuse** headache.
- Many patients seek help in identifying triggers (see below). The importance of trigger factors in migraine is nonetheless often overemphasised.

Trigger and predisposing factors

- Correctly identified triggers offer the possibility of **avoidance** (perhaps by life-style change) as a sometimes major contribution to management.
- When triggers are relevant to individual patients, they are usually selfevident.
- **Cyclical hormonal fluctuations** may be an obvious factor in menstruating women.
- **Irregular lifestyle**, poor sleep pattern and "stress" are important predisposing factors in anybody with migraine. Missing meals is a potent trigger factor.
- Triggers may be less readily identified when they are **cumulative** in their effect, jointly lowering the threshold above which attacks are initiated.
- Even when they are correctly identified, triggers are **not always avoidable**.
- Contrary to popular belief, there is **no "migraine diet"**. The only dietary triggers with good evidential support are certain alcoholic drinks (especially red wine).

Drug intervention

All patients should climb a **treatment ladder** (stepped management), usually treating three attacks at each step before proceeding to the next. This strategy, when followed correctly, reliably achieves the most effective and cost-effective individualised care.

Step one: symptomatic therapy

- non-opioid analgesic
- plus, when needed, an antiemetic.

Recommended drugs and doses are shown in Table 1.

Table 1. Recommended drugs and doses for acute migraine therapy, stepone

Analgesics	Antiemetics
Adults	
 Non-steroidal anti-inflammatory drugs: acetylsalicylic acid 900-1000 mg or ibuprofen 400-800 mg or diclofenac 50-100 mg or (where these are contraindicated): 	 domperidone 10 mg (supportive evidence of efficacy is for 20 mg, but the European Medicines Agency recommends restriction to 10 mg orally [up to three times daily] or 30 mg by suppository [up to twice daily]), or
 paracetamol 1000 mg* 	metoclopramide 10 mg
 or (possibly benefiting from the different mechanisms of action): combinations of paracetamol with acetylsalicylic acid or ibuprofen 	(the European Medicines Agency restricts dosing to 10 mg [up to three times daily])
Children (when needed)	
Ibuprofen 200-400 mg according to age and weight	 domperidone (dosage according to age and weight)

*Paracetamol on its own has lower efficacy and is **not** first-line treatment.

Drugs to avoid

- **Opioids** (including codeine and dihydrocodeine) are ineffective for migraine, associated with multiple adverse effects, potentially addictive and commonly implicated in medication-overuse headache;
- **Barbiturates** have no place in the treatment of migraine.

Principles of step one

- Use **soluble analgesics** (or mouth-dispersible formulations with water) when available.
- Take **early** in the attack.
- Use **adequate dosage** (see table 1: in most cases, adequate doses require more than a single tablet).
- A **prokinetic antiemetic counters gastric stasis**, an early feature of migraine, which impairs bioavailability of oral medication.
- Rectal formulations (where available) may be preferable in the presence of vomiting.
- Proceed to step two after three attacks without success (local guidelines may recommend trying more than one analgesic in step one before proceeding to step two).

Step two: specific therapy

- Where available, and unless contraindicated, specific therapy (Table 2) should be **offered to all patients failing step one**.
- Availability of drugs varies from country to country.

Table 2. Specific anti-migraine drugs, formulations and doses for step two(listed alphabetically)

Almotriptan	• tablets 12.5 mg
Eletriptan	tablets 20 mg and 40 mg
	 tablets 80 mg (not widely available)
	(for some people, 80 mg is effective when 40 mg is not)
Frovatriptan	• tablets 2.5 mg
Naratriptan	• tablets 2.5 mg
Rizatriptan	 tablets and mouth-dispersible wafers 10 mg
	 tablets 5 mg (to be used when propranolol is being taken concomitantly)
Sumatriptan	 tablets and rapidly dissolving tablets 50 mg and 100 mg
	 nasal spray 10 mg (licensed for adolescents) and 20 mg
	 subcutaneous injection 6 mg
Zolmitriptan	 tablets and mouth-dispersible tablets 2.5 mg and 5 mg
	 nasal spray 5 mg

Drugs to avoid

• **Ergotamine** is a poor substitute for triptans: it has very low and unpredictable bioavailability, which impairs its efficacy, and poor tolerability. It is no longer recommended for routine use.

Principles of step two

- Triptans are more effective when taken while headache is still mild (but not during aura) (this instruction should be given only to patients who can reliably distinguish migraine from tension-type headache).
- The initial dose of all oral triptans (except eletriptan in some cases) is one tablet.
- A **second dose** for non-response is not recommended by most triptan manufacturers but, taken not less than 2 hours after the first, may nonetheless be effective in some cases.
- Triptans should not be used regularly on ≥10 days/month to avoid the risk of medication-overuse headache.
- Triptans differ slightly, but there are large and unpredictable individual **variations in responses** to them:
 - one may work where another has not;
 - patients are best served if they can try several, in different formulations, and choose between them.
- When **nausea** is present, domperidone 10 mg may be added.
- When **vomiting** is present, zolmitriptan nasal spray (absorbed through the nasal mucosa) or sumatriptan subcutaneous injection may be preferred.
- Efficacy of sumatriptan may be increased by combination with naproxen 500-1000 mg (there are no data on combinations of other triptans and NSAIDs).
- When all other triptans are ineffective, sumatriptan by subcutaneous injection 6 mg should be considered.
- Triptans are associated with return of symptoms within 48 hours (relapse) in up to 40% of patients who have initially responded (see below).

Treatment of relapse

- A **repeat dose** of a triptan is usually effective.
- A further relapse may occur:
 - in a minority of patients, this happens repeatedly, a major management problem with high risk of developing medication-overuse headache;
 - a different triptan should be tried in future attacks;
 - concomitant use of a triptan and naproxen may reduce susceptibility to relapse.

Contraindications and special precautions in step two

- Triptans should not be taken **during aura** of migraine with aura, but at the onset of headache.
- All triptans should be **avoided** by people with:
 - uncontrolled hypertension (one reason for measuring blood pressure);
 - coronary heart disease, cerebrovascular disease or peripheral vascular disease;
 - multiple risk factors for coronary or cerebrovascular disease;
- In the **elderly**, all of these are more common, and triptans should therefore be used with **greater caution**.
- In **pregnancy**: limited safety data are available only for sumatriptan, which should be used with caution and **only under specialist supervision**.
- In addition, there are **specific precautions** attached to some triptans (see pharmacopoeia).

Step two for children and adolescents

- Failure of step one in children is an indication for specialist referral.
 - No specific anti-migraine drug has been shown to have efficacy in children (under 12 years old).
- For adolescents (12-17 years), the following have efficacy and are approved:
 - sumatriptan nasal spray 10 mg;
 - zolmitriptan nasal spray 2.5 mg and/or 5 mg (in some countries).

Follow-up

Every patient to whom treatment is offered, or whose treatment is changed, requires follow-up to ensure that optimum treatment has been established.

- Use of a **calendar** is recommended to monitor acute medication use or overuse. An example of a simple calendar is available as <u>Supplementary</u> <u>materials #17</u>.
- The use of **outcome measures** is recommended to guide follow-up. The following are included here among the management aids:
 - the HURT questionnaire (<u>Supplementary materials #20</u>) was developed expressly for primary care;
 - the HALT-30 Index (<u>Supplementary materials #19</u>) records lost productive time during the preceding month.
- Failure of acute therapy may be an indication for prophylaxis (see <u>Supplementary materials #8).</u>