



# Non-Opioid Symptom Management

Everything other than pain · match the drug to the mechanism, because the wrong antiemetic simply does not work

Pick by  
*mechanism*

## NAUSEA: MATCH THE RECEPTOR *the commonest reason an antiemetic fails is that it was aimed at the wrong pathway*

| Cause                                                          | First choice                                                                                | Why                                                                                                                                                                                       |
|----------------------------------------------------------------|---------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Drugs, metabolic, opioids<br><i>chemoreceptor trigger zone</i> | <b>Haloperidol</b> , low dose, or metoclopramide                                            | <b>The CTZ is dopamine driven</b> , so a dopamine antagonist is the rational choice. Haloperidol at antiemetic doses is far lower than antipsychotic dosing and is a palliative workhorse |
| Gastric stasis, early satiety                                  | <b>Metoclopramide</b> , a prokinetic                                                        | It moves the stomach as well as blocking dopamine. <b>Contraindicated in complete bowel obstruction</b> , where a prokinetic increases colic                                              |
| Bowel obstruction                                              | <b>Haloperidol plus an antisecretory</b> (glycopyrrolate or octreotide), with dexamethasone | <b>Avoid prokinetics</b> . The aim is to reduce secretions and colic rather than push contents through an obstruction. Consider a venting gastrostomy in the right patient                |
| Vestibular, movement-related                                   | <b>Antihistamine or anticholinergic</b> : meclizine, scopolamine                            | The vestibular pathway is histaminergic and cholinergic, so dopamine blockers underperform here                                                                                           |
| Raised intracranial pressure                                   | <b>Dexamethasone</b>                                                                        | Treats the cause by reducing peritumoral edema                                                                                                                                            |
| Chemotherapy                                                   | <b>Ondansetron</b> , with dexamethasone and an NK1 antagonist per protocol                  | Serotonin driven. <b>Note ondansetron constipates and prolongs the QTc</b> , and is a poor choice for opioid-induced nausea where a dopamine antagonist works better                      |

## BREATHLESSNESS *the most frightening symptom, and the most under-treated*

### OPIOIDS ARE THE TREATMENT

**Low-dose opioid is first-line for refractory breathlessness**, and it works independently of any effect on oxygenation. **Used properly it does not hasten death**, and fear of that is the main reason patients are left breathless. Start low, titrate, document the indication.

### NON-DRUG MEASURES WORK

**A fan directed at the face** stimulates the trigeminal nerve and genuinely reduces the sensation, with good evidence and no side effects. Also sitting upright, an open window, pacing activity, and a calm presence.

### OXYGEN ONLY IF HYPOXIC

**Oxygen does not help breathlessness in a non-hypoxic patient**, where airflow does just as much. Treat reversible causes when consistent with goals: effusion, anemia, bronchospasm, heart failure, pulmonary embolism. **Benzodiazepines are second-line**, for the anxiety component.

## THE OTHER SYMPTOMS *each has a drug worth knowing by name*

| Symptom                                             | Approach                                                                                                                                                                                                                                                                                                                                                                          |
|-----------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Delirium                                            | <b>Look for the reversible cause first</b> : opioids and other drugs, urinary retention, constipation, infection, hypercalcemia, hypoxia. <b>Non-drug measures are the mainstay</b> . Use <b>haloperidol</b> for distressing agitation, and note <b>benzodiazepines usually worsen delirium</b> except in alcohol withdrawal or terminal agitation                                |
| Constipation                                        | <b>A stimulant laxative with every opioid, from day one</b> , since tolerance to constipation never develops. Senna with or without a softener; escalate to an osmotic. <b>Exclude impaction and obstruction before escalating</b> , and consider a peripherally acting mu antagonist such as methylnaltrexone for genuinely refractory opioid-induced constipation               |
| Respiratory secretions<br><i>the "death rattle"</i> | <b>Reposition first</b> , and reduce parenteral fluids. Antimuscarinics such as <b>glycopyrrolate</b> (which does not cross the blood-brain barrier, so less sedating) or hyoscine. <b>They prevent new secretions rather than clearing existing ones, so give early. Explain to the family that it does not distress the patient</b> , since it usually distresses them far more |
| Anorexia and cachexia                               | <b>Dexamethasone</b> gives a short-lived appetite improvement. <b>Explain that artificial nutrition does not extend life or improve quality in advanced cancer or advanced dementia</b> , and can add burden. Focus on <b>food as comfort and pleasure</b> , and address the family's distress, which is often greater than the patient's                                         |
| Fatigue                                             | Treat the reversible: anemia, hypothyroidism, depression, hypercalcemia, drugs. <b>Energy conservation and graded activity</b> help. Corticosteroids give a short-term lift; methylphenidate is sometimes used with specialist input                                                                                                                                              |
| Mouth care                                          | <b>The single most appreciated intervention at the end of life</b> , and the most neglected. Regular oral care, saliva substitutes, treating candidiasis. <b>Dry mouth, not thirst, is what fluids fail to fix</b>                                                                                                                                                                |

## THE LESS COMMON ONES *each has a specific answer that is easy to forget*

| Symptom                     | Approach                                                                                                                                                                                                                                                                                                                                          |
|-----------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Hiccups                     | Exhausting and demoralizing when persistent. Look for <b>gastric distension</b> first, which is the commonest cause and responds to a prokinetic and a simethicone-type agent. Then <b>baclofen</b> , or chlorpromazine, gabapentin or metoclopramide. Consider phrenic irritation from tumor or subdiaphragmatic disease                         |
| Pruritus                    | Depends on the cause: <b>cholestatic</b> itch responds to cholestyramine, rifampin, naltrexone or sertraline rather than to antihistamines. <b>Uremic</b> itch responds to gabapentin. <b>Opioid-induced</b> itch is histamine release, so consider rotating the opioid. <b>Antihistamines mostly just sedate</b> in the non-histaminergic causes |
| Anxiety and insomnia        | Ask what the fear actually is: uncontrolled pain, dying alone, being a burden, or an unresolved practical matter. <b>Much of it is not treated by a drug</b> . Where medication is needed, a short-acting benzodiazepine, and treat any coexisting delirium first rather than deepening it                                                        |
| Seizures at the end of life | Anticipate them in brain tumors, brain metastases and severe metabolic derangement. <b>Prescribe a PRN by a route that will still work</b> , usually buccal or subcutaneous midazolam, and ensure the family knows what to do. Continue antiseizure cover when the oral route is lost rather than simply stopping it                              |

## PEARLS & PITFALLS

- **Constipation causes nausea, delirium, pain and agitation**. Before adding a fourth drug for any of those, **check when the patient last opened their bowels and do a rectal examination**. This is the highest-yield thing on this sheet.
- **Review the medication list and stop what no longer serves the goals**: statins, bisphosphonates, most antihypertensives, oral hypoglycemics, vitamins. **Deprescribing is an active intervention**, and each stopped drug is one fewer burden.
- **Anticipatory prescribing is what prevents a crisis at 2 a.m.** Write PRN orders in advance for **pain, breathlessness, nausea, agitation and secretions**, with a route that will still work when swallowing fails, usually subcutaneous.
- **Distinguish terminal agitation from reversible delirium and from unrelieved symptoms**. A patient thrashing may be in pain, in urinary retention, or hypoxic. **Examine before sedating**, since sedation for an untreated full bladder is a failure of assessment, not a failure of the drug.

**References:** Published palliative care symptom management guidance and consensus reviews on nausea by mechanism, refractory breathlessness, delirium and end-of-life secretions · standard palliative prescribing references.

*Educational aid; verify against current guidance, local formulary and patient factors.*

