

Humber and North Yorkshire Integrated Pharmacy and Medicines Optimisation Committee

Yorkshire and the Humber

A Guide to Symptom Management in Palliative Care

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Introduction

These symptom management guidelines have been produced by the regional Palliative and End of Life Care Group of Yorkshire and the Humber. They were updated in September 2023 and reflect a consensus of opinion from specialists working in the field of palliative medicine in hospitals, hospices and in the community.

This guidance covers some of the commonest symptoms in cancer and advanced progressive disease. For the management of other symptoms not included here, including fatigue, cough, sweating, anorexia and cachexia please see an introductory palliative care text and refer to the other useful resources listed in the introduction.

Disclaimer

These guidelines are the property of the Yorkshire and Humber Palliative and End of Life Care Group. It is intended that they be used by qualified medical and other healthcare professionals as an information resource, within the clinical context of each individual patient's needs. The group takes no responsibility for any consequences of any actions taken as a result of using these guidelines. Readers are strongly advised to ensure that they are acting in line with current accepted practice and legislation, as these may change. These include, but are not limited to, The National Institute for Health and Care Excellence (NICE), the NICE guidance on the prescription of opioids, the British National Formulary (BNF) and the Palliative Care Formulary (PCF). No legal liability is accepted for any errors in these guidelines, or for the misuse or misapplication of the advice presented here.

If unsure about what to do, please seek advice from your local specialist palliative care service.

Useful Resources

Details are given here of selected widely used medicines. See also British National Formulary (BNF) sections on "Controlled Drugs" and "Prescribing in Palliative Care." Check the BNF for formulations, dose recommendations, side effects and contra-indications.

Other useful resources are:

- Local intranet guidelines
- The latest version of 'The Palliative Care Formulary (PCF)' Twycross R and Wilcock A
- <u>Medicines Complete</u>
- Evidence.nhs.uk
- <u>NICE</u>
- <u>GMC guidelines</u> 'Treatment and care towards the end of life: good practice in decision making'
- <u>Hospice UK</u> publications and resources
- <u>SIGN</u>
- Symptom Management in EoLC for People with Dementia
- This is Me document helps staff better understand care needs of person with dementia
- SPICT tool

The Use of Medicines Beyond (Off-Label) and Without Marketing Authorisation

The use of medicines for off-label purposes is necessary when the clinical need cannot be met by the specifications of its marketing authorisations (MA), e.g. for an unauthorised indication, or in doses, preparations, patient population or route not covered by the MA. The recommendations within this symptom management guide include the off-label use of authorised medicines. These recommendations are based on current accepted palliative care practice in the UK. In practice, approximately 25% of medicines prescribed for palliative care patients are used off-license (e.g. when given by subcutaneous injection when only licensed for IM or IV use for the treatment of nausea and vomiting when only licensed as an antipsychotic); or unlicensed (e.g. when mixing medicines in a syringe before administration by continuous infusion).

Further information regarding off-label use of individual palliative care medicines can be found in the current version of the Palliative Care Formulary (PCF).

Prescribing medicines that are not licensed in the UK, or authorised medicines for off-label indications, may carry significant risks. The prescriber signing the prescription takes full responsibility for intended or unintended consequences of prescribing that medicine. When prescribing medicines off-label, it has been suggested that the prescriber should: discuss with the patient and document in the patient's records the reason why they are using the medicine off-label; where appropriate, gain informed consent from the patient; inform nurses and pharmacists to avoid misunderstandings where necessary; and give the patient a written leaflet where appropriate.

An Example Leaflet

In practice, recording every unlicensed use may be impractical and gaining informed consent in every instance may lead to unnecessary anxiety for the patient or carer. Practitioners must follow their clinical judgment on the balance of potential burden and benefit and their own organisation's policy on the use of authorised medicines for off-label purposes.

Principles of Symptom Management

- 1. Remember to consider the 'whole patient.' Symptoms are rarely purely physical or purely psychological and all symptoms and treatments impact on the patient, their family and friends.
- 2. Evaluate symptoms thoroughly. Consider all potential causes and remember to consider causes other than the underlying condition. Consider the impact of the symptom on the patient's quality of life.
- 3. Effective communication is essential. Explain in simple terms and avoid medical jargon. Discuss treatment options with patients and the people important to them and involve them in the management plan.
- 4. Correct the correctable, as long as the treatment is acceptable to the person/compatible with pre-existing priorities, practical and not overly burdensome. Remember non-drug treatments, e.g. palliative radiotherapy for metastatic bone pain.
- 5. Remember to consider non-pharmacological strategies to help relieve symptoms e.g. simple repositioning, or the use of a TENS machine may help pain; complementary therapies may help psychological distress. Although the evidence base for such treatments is not robust, some patients find them helpful.
- 6. When using drug treatments for persistent symptoms, give regularly and also 'as needed' (p.r.n.). Keep drug treatment as simple as possible.
- 7. Where the person has dementia, learning disabilities or other cognitive impairment, use oral medication first wherever possible and check medication adherence. Other formulations e.g. transdermal patches may also be useful.
- 8. Review regularly and adjust treatment accordingly.
- Plan in advance. Good communication is essential in establishing patients' wishes for their future care and treatment. Patients may want to document their wishes — the <u>Preferred</u> <u>Priorities for Care document</u> or an Advance Decision to Refuse Treatment may be helpful. Other helpful resources: <u>https://www.hospiceuk.org/our-campaigns/dyingmatters/dying-matters-resources</u>
- 10. Keep other staff informed.
- Ask for help. Refer to local guidelines or speak to the local Specialist Palliative Care Team (SPCT). Refer to General Medical Council (GMC) guidelines (see Useful Resources, page 4).

The following additional considerations are particularly important when assessing a person with dementia or severe learning disabilities:

- If the patient cannot self-report symptoms including pain, involve other people who know them, e.g. family members, professional carers, other clinicians, in order to better understand their 'normal state' and usual distress behaviours. Changes in behaviour can then be understood in context as they may indicate unrelieved symptoms.
- Consider using "<u>This Is Me</u>" or similar to ensure a patient's history and preferences are recorded and shared with staff.
- Use supportive communication strategies: ask short questions; allow additional response time; use gestures; minimise distractions/external noise; address sensory impairments; seek confirmation of assumptions made.
- Does the patient have the mental capacity to consent, with support, to examination/ investigation/taking medications? Will investigation alter management or can you treat on a presumed diagnosis?

Pain Management

Section A: Principles of Pain Management

- 1. Pain is common in advanced cancer and non-malignant conditions and its management can be difficult.
- 2. Pain is a total, personal experience with physical, psychological, social and spiritual dimensions. Optimal pain management will be compromised if any of these aspects are neglected. Management requires a multidisciplinary approach.
- 3. Regular review of the pain, effect and side effect of analgesics and how the pain is affecting the patient/family is vital for good pain control.
- 4. Not all pain experienced by a patient with cancer is caused by the cancer itself. Often several pains coexist and an accurate diagnosis of the cause as well as the type of pain and severity of each pain is necessary to enable effective pain management. Analgesic options will be determined by the specific cause, type and severity of pain. Previous exposure to analgesics (efficacy and side effects) will also determine the type and strength of analgesics used.
- 5. Principles of use of analgesia
 - a) Analgesics can be divided into three classes:
 - Non-opioid (simple analgesics), e.g. paracetamol or non-steroidal anti-inflammatory drugs (NSAIDs)
 - Opioids (weak and strong)
 - Adjuvants (co-analgesics)
 - b) Medicines from different classes are used alone or in combination according to the type of pain and response to treatment.

Two medicines of the same class (e.g. NSAIDs) should generally not be given concurrently; however, immediate release and sustained release opioids may be prescribed together.

Morphine sulfate orally (or morphine subcutaneously regarding parenteral use) is the most commonly used opioid in advanced cancer and other end-stage conditions, although non-opioids (e.g. paracetamol), a weak opioid and/or an adjuvant may suffice. Alternative opioids may be required in patients with renal and liver impairment or in those who develop side effects (seek specialist advice).

Opioids are safe and effective drugs to use in cancer pain management and to relieve other symptoms associated with cancer and non-malignant diseases.

- Side-effects are a major concern for patients, but these can usually be well managed
- Opioids are not routinely recommended for use in chronic pain in non-malignant diseases due to the risk of long-term side- effects
- Opioid abuse and addiction are thought to be rare in people with advanced illness when opioids are prescribed under close medical supervision. Risk assessment on prescribing opioids is advisable
- Therapeutic use of opioids for palliative reasons in patients with a history of substance misuse should involve specialist palliative care
- Correct opioid use, at the end of life, does not shorten life
- Some pains are only partially opioid- responsive. These include treatment related pains such as chemotherapy-induced neuropathy and pains unrelated to the underlying illness, such as tension headache, post-herpetic pain, muscle spasms, nerve damage/compression, bone pain, visceral distension/spasms, tenesmoid pain and activity provoked pain. These may require other measures including adjuvants, nerve blockade or oncological treatments if cancer related

In general, successful relief of pain in palliative care patients requires:

- Careful assessment and re-assessment
- Consider using the "modified WHO (World Health Organisation) analgesic ladder for the pharmacological management of cancer pain in adults and adolescents" which summarises the principles of analgesic use (Figure 1)

Figure 1: Modified WHO "ladder" for cancer pain relief in adults

Step 1 Non-opioid (Paracetamol and/or NSAID) ± adjuvant
Step 2/3 Opioid* ± non-opioid (Paracetamol and/or NSAID) ± adjuvant *(e.g.: morphine, oxycodone)

- Appropriate use and titration of opioids
- Early consideration of adjuvants and non-pharmacological approaches
- Regular review and assessment of pain control and use of as required (p.r.n.) medication. If patients are requiring several p.r.n. doses a day, assess the effect and side effects of these (if these fit with the drug's pharmacokinetics), and whether this is due to inadequately controlled background pain or the presence of breakthrough (episodic) pain
- Appropriate communication with the patient: holistic approach and explanation, asking about concerns and providing both verbal and written information where necessary on drug treatment (see section C, Communicating about opioids)
- Discussions with acute/chronic pain services, referral for anaesthetic interventions, consideration of radiotherapy or surgery as necessary

Distress and Pain in Dementia

(Some of this guidance might be applicable to some people with severe learning disabilities)

- People with dementia are less likely to ask for and receive pain relief. Pain assessment should include self-report, physical examination, caregiver reports and behavioural observations
- Pain may present as distress or delirium in people living with dementia. Analgesics, esp. opioids can cause Delirium so review pain and analgesic response (effects and side effects)
- Several tools are available to support pain assessment in people with cognitive impairment, including Pain Assessment in Advanced Dementia Scale (PAIN-AD), Abbey Pain Scale and Disability Distress Assessment Tool (DisDAT). When using these tools, watch for over-identification of pain when the person is actually distressed by other causes. Signs of distress may include agitation, walking with purpose withdrawal, nighttime waking, not eating/drinking
- People with dementia may be unable to communicate their experience of pain because of impaired memory or lack of expressive language. Ask about their pain using descriptive words such as aching, hurting, sore. Focus on current pain and ensure assessment is made during periods of activity and at rest. Consider using different representations of pain to help people self-report, e.g. visual, numerical and verbal rating scales. Select an appropriate tool for the individual and teach them how to use it when their pain is least severe

Section B: Assessment and Review

To manage pain effectively it is important to assess the:

- 1. Cause of pain.
 - a) Consider other causes/exacerbators of pain (e.g. psychological, spiritual and social) and perform a full holistic assessment.
 - Investigate appropriately, e.g. X-ray for pathological fracture; ultrasound or CT scan for deep soft tissue tumours; urgent MRI for suspected spinal cord compression.
 - c) Remember common non-malignant causes, e.g. arthritis, tension headache and infections, including oral candidiasis. In advanced, progressive disease there are usually multiple causes of pain and a management plan will be needed for each of these.
- 2. Type of pain: acute vs. chronic; nociceptive vs. neuropathic vs. inflammatory vs. visceral pain; episodic; breakthrough and incident pain.
- 3. Severity of pain. Use a combination of clinical assessments, e.g. facial expressions, groaning, the ability to move, the timing of pain, the number of sites and the patient's own perception, e.g. pain-rating scales.
- 4. Remember: Ask what helps? What makes it worse? Explore the effect of pain and analgesics (including side effects). Review and re-review as pain/analgesic requirements may change.
- 5. Consider atypical presentations in people living with Dementia or Severe Learning Disabilities, consider other signs of distress as signs of untreated pain. Use an appropriate assessment tool.
- 6. Which analgesic?

The principles governing analgesic use are summarised in the WHO guidelines for the pharmacological and radiotherapeutic management of cancer pain in adults and adolescents. See below for consideration of the dose, especially in people with renal function and weight. Monitor for effect and side effects.

- By mouth, where possible
- By the clock: Regular, as well as p.r.n. dose
- By the ladder: (Figure 1)
- After assessing the severity of pain, the modified analgesic ladder can be used to identify appropriate analgesics for the level of pain.

- Weak opioids are less likely to be indicated as low dose morphine has greater and more rapid analgesic effect (Step 2 and Step 3 combined in the WHO ladder)
- The patient should be reassessed, and analgesia administered in a step-wise manner working up the ladder until the patient's pain is managed
- Similarly, if the severity of pain is reduced, a patient's level of analgesics should move back down the ladder
- Alternative analgesia and adjuvants or non-pharmacological interventions should be considered at each level of the analgesic ladder
- Individual dose titration: Titrate dose against effect, buprenorphine, codeine and tramadol (has an upper dose limit). However, please consider specialist referral in high dose opioids (>120mg oral morphine equivalent/ 24hours) and inadequate pain control

Section C: Recommended Medicines

Opioid Analgesics

Weak opioids (less likely to be used)

- These include codeine, dihydrocodeine and tramadol
- Co-codamol is available in three strengths containing paracetamol and either 8mg, 15mg or 30mg of codeine. In elderly or frail patients, a lower strength may be required
- Codeine is a pro-drug of morphine. Its analgesic effect is via conversion to morphine, which varies between patients and there is a proportion of the population in whom codeine is ineffective

Strong opioids

• Strong opioids include morphine, diamorphine, oxycodone, fentanyl, alfentanil buprenorphine. Methadone and hydromorphone should be for specialist use only

Opioid side effects

These include:

- Constipation (very common, often persistent; always prescribe a regular laxative)
- Nausea and vomiting (often temporary; always prescribe a p.r.n. antiemetic)
- Drowsiness (dose-related and often temporary)
- Myoclonus, confusion, hallucinations and delirium (may need a dose reduction, if pain free, or change in opioid)

- Respiratory depression (rarely a problem if titrated correctly and regularly monitored). A decreased respiratory rate, associated with hypoxia and/or reduced consciousness, should raise concern for opioid-induced toxicity
- Neither tolerance nor addiction are significant problems in patients approaching end of life

General Principles

- a) Immediate and modified (slow) release preparations are available.
- b) All patients taking regular analgesics should also have analgesics prescribed for 'breakthrough pain' to take p.r.n.
- c) p.r.n. immediate release opioids should be individually titrated. Commonly this is 1/6 of the total daily dose but for some patients 1/10 (or less) may be sufficient. For a patient on 30mg slow-release morphine b.d. 1/6 of their total 24-hour morphine dose would be 10mg immediate release morphine. It is not necessary to make changes to the p.r.n. dose if this is effective for the patient, irrespective of the background dose.
- d) Maximum frequency and dose of p.r.n. opioids in 24 hours should be clearly stated.
- e) Prescribe opioids in mg rather than mL as differing strengths are available.
- f) Patients should be counselled on the difference in strength and therefore volume needed for administration (and returning old/unused strength supplies to community pharmacies to avoid administration errors).
- g) Prescribe regular laxatives and p.r.n. anti-emetics.
- h) Discuss side effects of opioids with the patient, including the potential impact on driving see <u>DVLA Guidance</u>
- i) When using opioid analgesics, if the pain is inadequately controlled and opioid responsive, then the background dose of opioids should be increased, usually by a maximum of 30-50% from current dose, taking the p.r.n. requirements into account, and after assessment of the effect and side effects of these.
- j) Halve the usual starting doses if the patient is elderly or frail.
- k) Some analgesics (including morphine) may accumulate in renal or hepatic impairment and specialist advice may be required. Careful individual tailoring of opioid dose is also required in patients with respiratory failure.
- It is advisable to seek specialist palliative care advice regarding patients receiving higher doses of opioids (>120mg/day oral morphine equivalent), especially when undertaking conversions to alternative drugs or routes of administration.

Communication About Opioids (NICE CG140)

When offering strong opioids to a patient, explore their concerns such as:

- addiction
- tolerance
- side effects
- fears that treatment implies dying

Provide verbal and written information on strong opioid treatment to patients and carers, including:

- When and why strong opioids are used to treat pain
- That opioids are not usually addictive

- How effective they are likely to be
- How, when and how often to take strong opioids (for background and breakthrough pain)
- How long pain relief should last
- Side effects and signs of toxicity
- Safe storage
- Follow-up and further prescribing
- Information on who to contact out of hours
- Offer patients access to frequent review of pain control and side effects

1.1 Oral Preparations

1.1.1 Morphine Sulfate

Formulations Available

Immediate release tablets and liquids are effective after 20-30 minutes and to last up to 4 hours. Examples include: Oramorph[®] solution (10mg/5ml, 20mg/1ml), Sevredol[®] tablets (immediate release morphine tablets — 10mg, 20mg, 50mg). Actimorph Orodispersible tablets ® (1mg, 2.5mg, 5mg, 10mg 20mg, 30mg) are available in some areas; seek advice from the specialist palliative care team.

Modified/slow-release tablets and capsules would be expected to be effective after 4 hours and to last for 12 hours. E.g. Zomorph[®] capsules.

Starting Regimen

- It is acceptable to start low dose strong opioids in opioid-naïve patients
- If the patient is already on an optimal dose of weak opioid with or without paracetamol and/or an adjuvant drug does not control the pain, the patient should be changed to morphine at a dose which is equivalent to the dose of the weak opioid that they are taking (see Appendix 1 for conversions). The weak opioid must then be stopped. For example, a patient taking 60mg of codeine phosphate q.d.s. regularly, could be commenced on modified release morphine (e.g. MST[®], Zomorph[®]) 15-20mg b.d. with immediate release morphine for breakthrough pain
- If the pain is intermittent or there are concerns about opioid sensitivity, commence immediate release morphine:
 - 2mg to 5mg 4 hourly p.r.n.
 - 1mg to 2.5mg 4 hourly p.r.n. in the frail or elderly

Reduce dose in renal failure (eGFR <30mL/min) or consider an alternative opioid (seek specialist advice).

Once a stable dose is achieved it is usual to convert to modified release preparations, e.g. a patient on 5mg oral morphine immediate release (e.g. Oramorph[®]) 4-6 hourly receives a total of 20mg morphine in 24 hours. This is equivalent to 10mg every 12 hours of morphine (modified release) tablets/capsules, e.g. MST[®], Zomorph[®]

Patients being initiated on morphine by either method should have their doses reviewed every 24 hours.

If the pain is inadequately controlled and is thought to be responsive to opioids then the background dose of opioids should be increased, taking the p.r.n. requirements into account. It is not advisable to increase the 24-hour dose by greater than 30-50%. All patients being titrated on morphine should be monitored for side effects and signs of CNS toxicity (confusion, drowsiness, hallucinations, myoclonic jerks).

If the patient has moderate to severe renal impairment (generally an eGFR <30mL/min), morphine and its metabolites will accumulate and specialist advice may be required regarding dosing/alternative opioids.

Breakthrough Pain

All patients on modified release morphine should have immediate release morphine available p.r.n. for breakthrough pain.

1.1.2 Oxycodone

Oxycodone is a strong opioid with a similar dosing schedule to morphine. It is a useful second line strong opioid for patients who have not tolerated morphine and may be first line in moderate to severe renal impairment. Caution is needed in moderate to severe hepatic impairment. Oral oxycodone is 1.5 to 2 times more potent than oral morphine. Consult a dose conversion chart when converting to oxycodone or ask advice from your local palliative care team or pharmacy.

Formulations Available

Oxycodone is available as immediate release oral solution (e.g. $OxyNorm^{(R)}$), capsules (e.g. Shortec (R)) and tablets (Oxyact (R)), all with a duration of action of 4-6 hours or modified/slow release (e.g. $OxyContin^{(R)}$) with a duration of action of 12 hours. Most cost-effective product should be used - in line with local formulary.

Breakthrough Pain

Immediate release oxycodone should be available p.r.n., at a dose which is usually about 1/6 of the 24-hour oxycodone dose.

1.2 Parenteral Preparations

This section contains information needed for prescribing continuous subcutaneous infusions (CSCI) via syringe pump. Syringe pumps are indicated if the patient is unable to take oral medication or there are concerns about absorption.

1.2.1 Morphine and Diamorphine Injections

Both morphine and diamorphine (rarely used and generally reserved for patients that require very high doses of opioids due to its solubility in small diluent volumes) can be given p.r.n. subcutaneously (SC) with a duration of action of up to 4 hours. Alternatively, they can be given as a CSCI via a portable syringe pump.

In an opioid naïve patient, start with morphine, between 1mg to 2.5mg SC p.r.n.

If a CSCI is indicated, then start between 5mg and 10mg morphine over 24 hours as a CSCI. Compared to oral morphine, parenteral morphine is about 2-3 times more potent and parenteral diamorphine is about 3 times more potent, although there is variability between individuals (refer to your local conversion policy).

For example, to change a patient from oral morphine to a CSCI of morphine, divide the total 24-hour dose of oral morphine by two, e.g. if a patient is on slow-release morphine 30mg b.d., they will require 30mg subcutaneous morphine over 24 hours. As there is inter-individual variability, reassessment of effect and side effects is recommended after 24 hours.

It is extremely important that p.r.n. analgesia is prescribed. Usually this is 1/6 of the total 24 hour opioid dose, e.g. in the above example, the SC p.r.n. dose would be between 2.5mg and 5mg morphine.

1.2.2 Oxycodone Injection

Patients on oral oxycodone can be converted to a subcutaneous infusion of parenteral oxycodone. To convert to subcutaneous oxycodone from oral oxycodone, divide the total daily dose of oral oxycodone by two. As there is inter-individual variability, reassessment of effect and side effects is recommended after 24 hours. As discussed above, it is extremely important that p.r.n. analgesia is prescribed.

Note: There are two concentrations of parenteral oxycodone available: 10mg/ml and 50mg/ml.

1.3 Transdermal Preparations

Transdermal preparations are suitable for patients with pain already stabilised on other opioids. They may be useful in patients with poor compliance with oral opioids or swallowing/absorption problems. They are generally less constipating. They should not be started in unstable pain or in the last days of life due to their long titration period and duration

of action. For patients in their last days of life who are on transdermal patches, do not remove the patch. Add supplementary opioid via CSCI if required. Some patients may experience fewer adverse effects than with oral morphine. Both fentanyl and buprenorphine are safer than morphine in patients with renal failure.

N.B. All patients using transdermal patches should also be prescribed an immediate release preparation for breakthrough pain (generally of morphine or oxycodone), the dose of which is dependent on the patch strength (see conversion tables or local guidance for details).

Transdermal fentanyl patches are available in 12, 25, 37.5, 50, 75 and 100 micrograms per hour strengths and are applied every 72 hours.

Transdermal buprenorphine patches are available as:

- Low dose patches in 5, 10, 20 micrograms per hour strengths and are applied every 7 days. These may be helpful in patients with poor compliance who require a low dose opioid
- Higher strength patches in 35, 52.5, 70 micrograms per hour are usually applied every 96 hours (check product)

As different brands are becoming available, check recommended patch application interval. Consult a dose conversion chart (see local guidelines and Appendix 1) when starting transdermal opioids or ask for advice from your local palliative care team or pharmacist.

1.4 Other Routes of Administration of Strong Opioids

Formulations of sublingual, buccal and nasal fentanyl are available and may be advised in specific situations by specialist teams.

1.5 What if Opioids do not Work?

a) Are opioids the analgesic of choice?

Not all pain is opioid responsive. Consider its aetiology. Palliative radiotherapy is helpful for bone metastasis and can be given as a single treatment. In certain patients a nerve block will help, e.g. coeliac plexus block in pancreatic pain. Discuss with palliative care or chronic pain specialists.

b) Is the dose high enough?

If there is a partial response or inadequate duration of pain relief, i.e. if pain returns less than 4 hours after immediate release oral morphine or less than 12 hours after modified release morphine and there are no side-effects, increase the dose by 30% increments rather than shortening the interval between doses. Remember to check that the p.r.n. dose prescribed is adequate for the background dose.

c) Is the drug being absorbed?

If there is uncontrolled vomiting, dysphagia or high stoma output, consider alternative routes of delivery, e.g. subcutaneous, transdermal.

- d) Is pain breaking through with movement or painful procedures?
 Identify and minimise provoking factors. Consider pre-emptive doses of immediate release opioid; consider NSAIDs. Discuss with palliative care team.
- e) Are adjuvants required? Please see next section for indications.
- f) Who might be able to help?
 Do not be afraid to ask a more experienced colleague for help. Your hospital palliative care team, local hospice or community palliative care team will gladly offer advice.

Adjuvants

Choice of adjuvant analgesic will be determined by the aetiology of the pain.

Medicines for Cancer Induced Bone Pain

Consider palliative radiotherapy, bisphosphonates, NSAIDs and corticosteroids, as well as opioids and medicines for neuropathic pain (see section 2.2).

Medicines for Neuropathic Pain

The decision to use an antidepressant or an anticonvulsant depends on a patient's symptoms and the adverse effect profile (anti-cholinergic side- effects with amitriptyline include dry mouth, urinary hesitancy, postural hypotension and constipation. Sedation, dizziness, peripheral oedema and gastro-intestinal effects occur with gabapentin or pregabalin). Remember to consider the patient's comorbidities when prescribing.

Refer to NICE guidelines CG173: Neuropathic pain — pharmacological management: the pharmacological management of neuropathic pain in adults in non-specialist settings.

2.3 Antidepressants for Pain Control

Start with low dose, e.g. amitriptyline 10mg at night, titrating gradually every 2-5 days, if adverse effects allow, to 75mg at night (lower than usual antidepressant doses).

2.4 Anticonvulsants for Pain Control

Titration is often slower than stated in the BNF, with particular caution needed in frail and elderly patients. Start with the lowest dose. Use with caution and dose reduce in renal impairment and seek advice if necessary.

Gabapentin Start 100mg – 300mg o.d.-t.d.s. slowly titrate assessing efficacy/adverse effects; usual maximum dose 1.2g t.d.s.

Pregabalin Start 25mg to 75mg b.d.; slowly titrate assessing efficacy/adverse effects; maximum dose 300mg b.d.

In addition, for nerve root compression consider a short course of 4mg to 8mg dexamethasone, NSAIDs, palliative radiotherapy, and pain team interventions.

2.5 Medicines for Pain Due to Raised Intra-Cranial Pressure

Dexamethasone is the corticosteroid of choice with high anti-inflammatory potency, high solubility and low mineralocorticoid effect (less salt and fluid retention).

Use 4mg to 8mg unless severe symptoms or risk of coning, in which case use 16mg o.d. Titrate down to lowest effective dose and use for shortest possible time. Taper the dose slowly when stopping (not usually necessary if duration of treatment less than two weeks). Prescribe doses to be given in the morning to avoid causing insomnia. Remember the risk of hyperglycaemia.

Note: 1mg Dexamethasone ~ 7mg Prednisolone

5mg Prednisolone \approx 750 micrograms dexamethasone. (Approximation of equivalent anti-inflammatory effects).

2.6 Medicines for Other Pain

machine.

Management depends on the aetiology of the pain; thorough assessment is vital.

Painful skeletal muscle spasmsDiazepam2mg to 5mg, once at night (or b.d./t.d.s.)Baclofen5mg t.d.s.

<u>Liver capsule pain</u> Consider a short trial of NSAIDs or dexamethasone between 4mg and 8mg o.d.

<u>Musculoskeletal pain</u> Consider NSAIDs (oral or topical), or transcutaneous electrical nerve stimulation (TENS)

Intestinal colic Anti-spasmodics: hyoscine butylbromide 20mg SC.

*Also see Intestinal Obstruction section, page 25.

<u>Pelvic pain</u> Consider NSAIDS or corticosteroids, and antispasmodics for colic.

2.7 Non-Steroidal Anti-Inflammatory Drugs (NSAIDs)

Most patients with cancer/advanced progressive disease have risk factors for significant gastrointestinal adverse effects, therefore consider use of a proton-pump inhibitor (or a H2-receptor antagonist) alongside NSAIDs. Use NSAIDs with caution in patients with renal

impairment, (eGFR <30mL/min) uncontrolled hypertension or heart failure. Balance the short and long-term risks and benefits. All can cause renal impairment; choice is determined by side effect profile (cardiovascular vs GI) and route of admin s/c vs oral; if in doubt seek advice. There are a range of drug interactions, including with anti-coagulants.

Ibuprofen tablets400mg t.d.sNaproxen tablets250mg to 500mg b.d.Celecoxib capsules100 to 200mg b.d

2.8 Other Medicines

A wide variety of other medicines can be used as analgesics, e.g. ketamine, methadone, and tapentadol. Their use should be guided by specialists and local availability.

OPIOID ANALGESIC EQUIVALENCES

General Principles

- a) LOCAL ORGANISATIONAL OPIOID CONVERSION CHARTS for opioid use in palliative care MUST be used in preference to these conversion tables below.
- b) If there is any uncertainty regarding the safe prescribing of opioids seek specialist advice before doing so.
- c) It is advisable to double check calculations and document the method used in the patient record, including for appropriate p.r.n. opioid.
- d) Clinical judgement must also be applied, considering: underlying clinical situation; comorbidity and concomitant medicines; nature of pain and its opioid responsiveness; toxicity of current opioid; previous opioid doses and adherence; rapidity of opioid escalation; reason for switching — if pain is controlled, switching due to adverse effects or convenience is usually less problematic than switching if the pain is uncontrolled (seek specialist advice).
- e) In view of incomplete cross tolerance, caution is needed when converting between opioids. Start with the most conservative dose.
- f) Larger doses of opioid often require an empirical decrease in the dose of the replacement opioid and re-titration. For doses greater than 120mg oral morphine equivalence a day it is advisable to seek advice.
- g) Analgesic review and monitoring for adverse effects, including consideration of patient's place of care, need to be in place, documented in the patient record and communicated.

These tables should only be used in the context of these guidelines as a whole and used alongside the summary key principles outlined above.

Weak opioids – dose conversion to oral morphine N.B. Some people cannot efficiently metabolise codeine or tramadol to the active metabolite and therefore they may require a lower dose of morphine

Oral "Weak" opioid	Total MAX daily dose	Conversion factor	Approximate 24- hour oral morphine dose equivalence
Oral codeine phosphate	240mg/day	÷ 10	24mg/day
Oral dihydrocodeine	240mg/day	÷ 10	24mg/day
Tramadol hydrochloride	400mg/day	÷ 10	40mg/day

TRANSDERMAL FENTANYL

Comparative doses based on dose conversion ratio of between 100 to 150:1 (approximated). Changed every 72 hours (occasionally need to change every 48h if breakthrough pain on day 3). If possible, patients should not be switched between brands when on stable dose.

Start with the most conservative dose conversion, reassess and titrate if needed. Use local organisational opioid conversion charts in preference to these tables. Due to pharmacokinetics and incomplete cross tolerance, seek specialist advice for higher doses or if you are uncertain of the conversion.

Although unusual, if converting from fentanyl to oral opioids, seek specialist advice. In general, the most conservative dose conversion should be used (e.g. from 25 micrograms/h fentanyl patch to 30mg morphine b.d.). Please note that patches should be continued at the end of life.

Fentanyl patches micrograms/hr	24-hourly oral morphine dose	4-hourly and breakthrough oral morphine dose (rounded to practical amounts)
12	30 to 45mg	5 to 10mg
25	60 to 90mg	10 to 15mg
37	90 to 135mg	15 to 20mg
50	120 to 180mg	20 to 25mg
62	150 to 225mg	25 to 30mg
75	180 to 270mg	30 to 35mg

These tables should only be used in the context of these guidelines as a whole and used alongside the summary key principles outlined above and local guidelines.

TRANSDERMAL BUPRENORPHINE

Comparative doses are based on a dose conversion ratio of between 75 to 115:1 (approximated). Start with the most conservative dose conversion, reassess and titrate if needed. Use local organisational opioid conversion charts in preference to these tables. Seek specialist advice for higher doses or if you are uncertain of the conversion.

Although unusual, if converting from Buprenorphine to oral opioids, seek specialist advice. In general, the most conservative dose conversion should be used (e.g. from 35 micrograms/h Buprenorphine patch to 30mg morphine b.d.). Please note that patches should be continued at the end of life.

Buprenorphine patches micrograms/hr	Approximate 24-hourly oral morphine dose equivalence	Approximate 4-hourly and breakthrough oral morphine dose
Change every 7 days		
5	10 to 15mg	1 to 2mg
10	20 to 30mg	3 to 5mg
20	35 to 55mg	5 to 10mg
Change every 96 hours		
35	60 to 95mg	10 to 15mg
52.5	95 to 145mg	15 to 20mg
70	125 to 190mg	20 to 30mg

These tables should only be used in the context of these guidelines as a whole and used alongside the summary key principles outlined above and local guidelines.

Nausea and Vomiting

- It is important to fully assess and consider all possible causes, including those which may require specific treatments rather than an antiemetic alone (e.g. hypercalcaemia, gastritis, oral candidiasis)
- Causes can be multifactorial.
- Are environmental factors contributing? Can they be minimised, e.g. by reducing strong food smells, offering smaller portion sizes?
- Rationalise medications and reduce tablet burden
- Prescribe drugs regularly as well as p.r.n.
- Consider longer acting anti-emetics to reduce tablet burden
- If there is significant nausea and/or vomiting, the oral route may be temporarily ineffective and parenteral anti-emetics may be required. The subcutaneous route is preferred unless the intravenous route is required for another purpose
- Consider disease specific cautions when prescribing:
 - Cyclizine may worsen heart failure.
 - Centrally acting anti-dopaminergic drugs such as metoclopramide, haloperidol and levomepromazine may worsen Parkinsonian symptoms. Consider using domperidone or a 5HT3 antagonist (or cyclizine, with monitoring) in Parkinson's disease.

- Cyclizine and other antimuscarinic drugs block the final common pro-kinetic pathway through which metoclopramide acts, therefore concurrent administration should be avoided.
- Cyclizine and hyoscine butylbromide may crystallize when mixed in a syringe pump (especially in doses over 60mg Buscopan). Cyclizine is incompatible with 0.9% normal saline, use water for injections.

If initial advice in the Drug Management Table (pages 52-53) is not effective, contact your local palliative care team.

The European Medicines Agency's Committee on Medicinal Products for Human Use recommended key changes to the licensed use of metoclopramide and domperidone (because of the risk of prolonged QT syndrome). They concluded that the benefit of these drugs outweighs the risk only when used short- term for nausea and vomiting and not as a prokinetic. If the use of metoclopramide or domperidone is being considered above the MHRA (Medicines and Healthcare products Regulatory Agency) recommended dose of 30mg daily or for more than one week for domperidone and 5 days for metoclopramide, seek specialist advice. Avoid/seek advice before using domperidone for anyone known to have heart disease/ conduction defects — ECG monitoring may be advised for some patients. In palliative care, off-label use is recognised as standard practice and recommended cautions to the licensed use of the prokinetics should not necessarily change practice.

Cause	First-line Drug	Stat Dose (PO or SC)	24 Hour Range
Gastric stasis and irritation	Domperidone or	10mg PO	30mg PO
	Metoclopramide	10mg PO or SC	30mg to 60mg PO or SC
	+/- proton pump inhibitor/H2		
Bowel obstruction -	receptor antagonist Metoclopramide	10mg SC	30mg to 60mg SC
incomplete WITHOUT	Metoclopramide	(Only use SC)	(Only use SC)
Bowel obstruction WITH colic	Haloperidol +/-	1mg SC	1mg to 5mg
(or if develops colic while on Metoclopramide)	cyclizine +/-	50mg SC	SC 100mg to 150mg SC
, ,	hyoscine butylbromide (Buscopan [®])	10mg to 20mg SC	60 to 120mg SC
	NB: cyclizine and buscopan may be incompatible	(Only use SC)	(Only use SC)
Chemical e.g. • drugs • hypercalcaemia • uraemia	Haloperidol	0.5mg to 1mg PO or SC	1mg to 5mg PO or SC
Raised intracranial pressure	Dexamethasone	8mg to 16mg PO	8mg to 16mg PO
	+/- cyclizine	50mg PO or SC	100mg to 150mg PO or SC
Motion /Vestibular	Cyclizine or	50mg PO or SC	100 to 150mg PO or SC
	Hyoscine hydrobromide	300 micrograms sublingual 400 micrograms SC	300 micrograms sublingual q.d.s. 800 to 1,200 micrograms SC
			Patch behind ear change every 72 hours
2nd line or Multifactorial	Levomepromazine	3 to 6.25mg PO or 2 to 6.25mg SC	6.25 to 12.5mg PO or SC

Intestinal Obstruction in Advanced Cancer

Introduction

Intestinal obstruction in advanced cancer is the interrupted passage of food and fluids through the gastrointestinal tract due to mechanical or functional occlusion. It could be partial, complete, or intermittent and at single or multiple sites. The incidence of intestinal obstruction in advanced cancer is between 3% and 6%. It occurs more often in ovarian and bowel cancers.

Clinical Features

Symptoms depend on the level, type and duration of intestinal obstruction and may include any or all of the following:

- Nausea: often postprandial, intermittent and relieved by vomiting undigested food
- Vomiting: may be faeculent (irrespective of the site of obstruction)
- Dull aching pain: may be due to tumour mass and/ or nerve infiltration
- Colicky pain and altered bowel sounds: often due to mechanical obstruction
- Abdominal distension: however, this may be absent in high obstruction
- Paradoxical diarrhoea and/or constipation
- Other signs and symptoms include anorexia, dry mouth and dehydration

Diagnosis

- History and physical examination are the most useful
- Contrast radiography may help define site and extent of obstruction. CT scan assists in choice of surgical intervention. Abdominal X-rays (supine and erect) may help but "normal appearances" do not exclude bowel obstruction
- An important differential diagnosis is constipation due to faecal impaction which may mimic, or co- exist with and complicate, intestinal obstruction
- Passage of flatus stops in complete obstruction and therefore passage of flatus or stools argues against this

Management

All patients will require symptom management that is specific to the individual and based on the aims of treatment as well as prognosis. Surgical intervention should also be considered early in selected cases.

Surgical Management

Palliative surgery is a reasonable option for some patients. However, selecting patients who are likely to benefit from a surgical procedure (e.g. bowel resection, by-pass +/- stoma formation or venting gastrostomy) is difficult. These decisions are best made with an experienced surgical colleague and careful discussion with the patient. Patients likely to benefit are those with no other life-threatening disease and single-site obstruction. Other factors to consider include patient performance and functional status, prognosis (ascites is a poor predictor of outcome), co-morbidity, nutritional status and options for further treatment such as chemotherapy.

Medical Symptom Management

When surgical approaches are inappropriate or not possible, symptomatic palliative treatment aimed at reducing symptoms and providing the highest possible quality of life becomes the main priority. Good symptom management can usually be achieved and greatly improves quality of life.

General measures include:

- Frequent and meticulous mouth care
- Small amounts of oral fluids and food as desired
- Large bore NG tube for decompression is indicated while surgery is being considered or as an intervention and after discussion with the patient may be an acceptable option to them as a venting mechanism to relieve vomiting in cases of bowel obstruction

Pharmacological Measures for Symptoms

Medication should generally be given by subcutaneous injection or continuous subcutaneous infusion (CSCI).

- a) Nausea and vomiting
 - Set realistic goals. Nausea can usually be reduced significantly but vomiting may continue once or twice daily
 - Give anti-emetics parenterally and regularly. Subcutaneous infusion is often helpful (see nausea and vomiting section, page 22)
 - Anti-secretory drugs. These include hyoscine butylbromide, and octreotide (used under the guidance of a specialist)
 - A proton pump inhibitor is also often used to reduce gastric secretions. Pantoprazole can be given subcutaneously as an infusion seek specialist advice
- b) Pain

Colicky pain

- Stop stimulant laxatives and prokinetic drugs, e.g. metoclopramide, in complete obstruction
- Use antispasmodics (hyoscine butylbromide, 60 to 120mg/24 hours by CSCI)

Dull aching pain

• Morphine subcutaneously (if helpful consider starting a CSCI)

Pain from tumour mass

- Consider dexamethasone/chemotherapy/ radiotherapy to reduce tumour/peri-tumour oedema (under specialist guidance)
- c) Constipation
 - Examine lower rectum or stoma for faecal impaction, if safe and appropriate to do so
 - For partial obstruction, use laxatives (gentle softeners e.g. docusate) with caution

- d) Ongoing nutrition and hydration
 - IV fluids and total parenteral nutrition (TPN) are rarely appropriate in advanced cancer. SC fluids may be used for thirst; usually give 1-2L/24 hours, preferably site this in the abdomen
 - Oral intake of food and drink can continue for the patient's enjoyment and is often surprisingly well tolerated the patient will decide if the risk of vomiting outweighs the pleasure of eating

Note: Patients with a high obstruction without other life- threatening complications require special consideration regarding symptom management, hydration and nutrition, e.g. venting gastrostomy, subcutaneous fluids. TPN may be considered in individual cases.

Constipation

Constipation is very common in palliative care patients due to a combination of factors including immobility, reduced food and fluid intake, medication, bowel pathology and sometimes hypercalcaemia. Diagnosis is usually made on the basis of a history of decreased frequency of bowel movements, the passage of small hard faeces and the need to strain. Constipation can present with overflow diarrhoea. Abdominal X-ray is rarely required. Consider patient education and information about the causes of constipation, increasing fluid intake and making appropriate dietary changes to help improve symptoms.

Guidelines on the Use of Laxatives in Constipation

- There is limited research for the management of constipation in palliative care patients and a lack of evidence to support a particular laxative regimen
- Patient preference regarding laxative formulation (tablet, liquid, volume required), palatability and drug tolerability (flatulence, colic) can impact greatly on adherence and therefore patient views should be sought
- Assess the cause and treat where possible. Most patients on regular opioids will require laxatives
- A combination of laxatives with different mechanism of action (e.g. stool softener and stimulant laxative) is usually required
- Examples of stool softeners include docusate, macrogol and lactulose
- Examples of stimulant laxatives include senna, bisacodyl, sodium picosulfate
- Local units may have their own guidelines on first line laxatives
- Review laxatives every 2 days and titrate as required
- Avoid stimulant laxatives if colic is present. If faecal leakage occurs, consider reducing the dose of the softener
- Lactulose may cause significant flatulence and bloating. It is not generally 1st line
- In complete bowel obstruction, do not prescribe laxatives without seeking advice
- If patients are managing well on their laxative regimen, there is no need to change laxatives
- Consider rectal intervention and follow local guidelines if bowels have not moved in 3 days despite oral laxatives, particularly in frail bedbound patients or those with paralysis
- Neurogenic bowel dysfunction can occur in patients with spinal cord compression / other central neurological conditions, and may require commencement of a tailored bowel regimen (see <u>https://www.mascip.co.uk/wp-content/uploads/2015/02/CV653N-</u><u>Neurogenic-Guidelines-Sept-2012.pdf</u>

Seek advice from your local SPCT regarding opioid-induced constipation resistant to optimal laxative regimens. Some opioids, e.g. fentanyl or buprenorphine are potentially less constipating. Also, peripherally acting opioid antagonists could be considered, e.g. Naloxegol or Naldemedine.

Breathlessness

Definition: an uncomfortable awareness of breathing. Breathlessness occurs very commonly in advanced cancer and in cardiorespiratory and neurological diseases. Look for reversible causes as listed below.

Sudden onset breathlessness		
Possible cause:	Consider:	
Asthma	Bronchodilators, corticosteroids	
Pulmonary oedema	Diuretics, morphine	
Pneumonia	Antibiotics, physiotherapy	
Pulmonary embolism	Anticoagulants, morphine	
Pneumothorax	Chest drainage	
Breathlessness arising over several	days	
Possible cause:	Consider:	
Exacerbation of COPD	Antibiotics, bronchodilators, corticosteroids, physiotherapy	
Pneumonia	Antibiotics, physiotherapy	
Bronchial obstruction by tumour	Dexamethasone 16mg o.d. early radiotherapy (RT), laser or stents	
Superior vena caval obstruction	Dexamethasone 16mg o.d.	
	Urgent stenting	
Breathlessness of more gradual on	set	
Possible cause:	Consider:	
Congestive cardiac failure	Diuretics, digoxin, ACE inhibitors	
Anaemia	If chronic, this is unlikely to be the major cause of	
	breathlessness. Transfusion may help if Hb<80g/l. Oral iron	
	is ineffective in chronic normochromic normocytic anaemia	
Pleural effusion	Pleural aspiration and follow with pleurodesis if	
	appropriate. These procedures may be distressing for frail	
	patients	
Pulmonary fibrosis	Possible if history of cytotoxics (esp. bleomycin), or lung	
	radiotherapy	
Ascites	Paracentesis if appropriate	
Primary/secondary carcinoma lung	Resection, RT or chemotherapy as appropriate	
Carcinomatous lymphangitis	Dexamethasone 8 to 12mg once in the morning. Stop if not effective within one week. Bronchodilators may help	

Palliative Management Where There are no Reversible Causes

Chronic Breathlessness is defined as "disabling breathlessness which persists despite optimum treatment of the underlying physiology."

- Breathlessness is frightening for the patient, family and staff
- Reassurance and explanation are vital parts of the treatment whatever the cause
- Modification of lifestyle, breathing retraining, relaxation and tailored exercise may be beneficial if instituted early enough and should be provided for all breathless patients as tolerated
- Consider referral to physiotherapist or occupational therapist
- A portable/table fan directed onto the face and hand-held fan
- Good oral care is important if the patient is mouth breathing
- Humidified oxygen may help acute breathlessness only in the presence of significant hypoxaemia. An exception to this rule is people with COPD where they may get benefit for breathlessness even with mild hypoxaemia. Use a trial of oxygen alongside other measures. Review regularly
- Most patients requiring palliation for breathlessness will not benefit from oxygen therapy (unless they are significantly hypoxaemic). Measurement of oxygen saturation levels using a pulse oximeter may aid decision making as to whether or not oxygen will be of benefit

Medicines to Consider

All medicines for symptomatic relief of breathlessness are respiratory sedatives. When prescribed, their use should be monitored carefully. In the context of distressing breathlessness in the terminal stages of illness the benefits usually outweigh the risks.

Opioids

Opioids should only be considered for severe breathlessness persistent despite nonpharmacological techniques. Benefits are greater at steady state, so oral morphine modified release 5 to 10mg twice daily (with concurrent prescription of a laxative) is advised. For chronic breathlessness associated with advanced disease titrate by 5mg b.d. every 7 days according to function until side-effects or a maximum dose of 15mg b.d. is reached.

Alternatively, if fluctuating severe renal function is a concern, oral morphine (immediate release) may be given as regular 2.5mg up to a maximum daily dose of 20mg. Note the dosing interval may need to be more than 4-hourly. If tolerated, change to modified release.

If the patient is already taking a strong opioid for analgesia, contact palliative care team for advice.

Benzodiazepines

Lorazepam 500 micrograms to 1mg oral or sublingual (e.g. Genus brand, as not all brands suitable for sublingual use, recommend check with local pharmacist). Maximum 4mg/day may give relief during panic attacks if non-pharmacological techniques are ineffective.

If anxiety appears to be a significant driver for the breathlessness, then try an anxiolytic anti-depressant, e.g. mirtazapine 15 to 30mg nocte Midazolam 2.5mg SC may benefit patients who cannot tolerate the oral/sublingual route. These medicines can be continued in the terminal phase. See section on 'Last Days of Life' (page 44).

Sertraline (a selective serotonin reuptake inhibitor) does not provide greater benefit for chronic breathlessness over placebo.

Delirium

Delirium is extremely common in patients with advanced disease, including advanced dementia, and is exacerbated by dehydration. It is a source of increased morbidity and distress and interferes with the ability to communicate effectively.

Delirium is often unrecognised or treated inappropriately and can be misdiagnosed as dementia, depression, anxiety or psychosis.

Clinical Features

- a) Disturbance of consciousness (i.e., reduced clarity of awareness of the environment) with reduced ability to focus, sustain, or shift attention.
- b) A change in cognition (such as memory deficit, disorientation, language disturbance) or the development of a perceptual disturbance that is not better accounted for by a pre-existing, established, or evolving dementia.
- c) The disturbance develops over a short period of time (usually hours to days) and tends to fluctuate during the course of the day.
- d) There is evidence from the history, physical examination or laboratory findings that the disturbance is caused by the direct physiological consequences of a general medical condition.

Subtypes of delirium are based on the type of arousal disturbance:

- Hyperactive
- Hypoactive
- Mixed (both hyper and hypoactivity)

Assessment

Obtain a thorough history to determine the patient's pre-morbid level of functioning, their use of alcohol, nicotine and illicit substances and the chronology of the onset of the changes in their mental state.

Consider that people with dementia are particularly vulnerable to the side effects of medicines that exacerbate confusion, e.g. anticholinergics, amitriptyline. In Parkinson's dementia and Lewy Body dementia, dopamine antagonists can cause confusion, hallucinations and delusions.

Cognitive assessment tools such as the Abbreviated Mental Test Score should be used to gauge the patient's cognitive state but will not differentiate delirium from other causes of cognitive impairment. Use of a screening tool to identify those at risk and those with delirium is recommended. The Short-Confusion Assessment Method is the preferred assessment tool according to NICE guidance. Identify any reversible causes: medication, e.g. medicines with anticholinergic side effects such as cyclizine (<u>Anticholinergic Burden</u> <u>Score</u> to review medications); corticosteroids; infection; biochemical abnormalities; hypoxaemia; and alcohol or nicotine withdrawal.

See THINK DELIRIUM

Management

Non-pharmacological measures are the mainstay of treatment and include:

- Addressing reversible causes, including dehydration, constipation, urinary retention, uncontrolled pain, liver or renal impairment, electrolyte disturbance (sodium, glucose, hypercalcaemia), infection, hypoxia, medications (changes, cessation, withdrawal)
- Maintaining adequate hydration and nutrition
- Managing the patient's environment to reduce confusion and distress e.g.
 - Visible clock to aid orientation
 - Encourage family to visit and explain things fully to them
 - Consistent nursing
 - Good lighting during daytime
- Pharmacological interventions are usually indicated only in severe delirium with distressing hallucinations (use the lowest dose for the shortest possible time):
 - Oral haloperidol 500 micrograms to 1.5mg nocte or b.d. Additional doses every four hours as needed (usual maximum 5mg/day)
 - SC haloperidol 500 micrograms to 1mg; observe for 30-60 minutes; repeat if necessary
 - Review at least every 24 hours and seek further advice from Specialist Palliative Care if not effective
 - Seek advice in Parkinson's disease
 - Discontinue within 7 days if symptoms resolve
 - Benzodiazepines should be used with caution due to their tendency to sedate and increase confusion

Neuropsychiatric Symptoms

These are nearly universal in dementia and agitation is among the most distressing for patients and family carers. Exclude or treat specific causes, e.g. pain, and prioritise non-pharmacological approaches.

Obtain a thorough history to determine the patient's pre-morbid level of functioning, the onset of changes in their mental state and any potential cause, e.g. infection, medication. Whilst antipsychotics may be necessary, all psychotropics including benzodiazepines can increase confusion and should be used with extreme caution.

Occasionally in severe anxiety/agitation at the end of life a trial of a benzodiazepine or antipsychotic may be appropriate, weighing up the risks and benefits. Antipsychotics should be avoided, if possible, in Lewy body dementia as these can cause severe side effects.

Oncological Emergencies

Metastatic Spinal Cord Compression (MSCC)

Introduction

- Spinal cord compression is a well-recognised complication of metastatic cancer
- This can be a catastrophic event leading to paralysis below the level of the compression, urinary retention and faecal incontinence
- If treated early, these problems can usually be prevented or at least partially reversed
- MSCC and vertebral body metastases (VBM) occur more frequently in some tumour types when there is metastatic disease (especially bone) and in the later stages of a cancer trajectory. Lung, breast and prostate cancers account for over 50% of cases; lymphoma and myeloma account for 20%. Patients at high risk may have been identified by treating clinical teams and informed both of features to look out for and what to do if they suspect that they may be developing VBM or MSCC. Such patients should have been provided with an MSCC information booklet
- Many of the features of MSCC (back pain, weakness, bladder and bowel changes) are non-specific features of advanced cancer so the patient's symptoms and signs in the 'context' of their cancer must be considered
- MSCC and VBM can be suspected clinically but can only be proven by imaging (MRI is the gold standard)
- Patients should only be referred for MRI if they are fit enough to tolerate an MRI scan (40 minutes lying flat) and able to travel to the local radiotherapy or spinal surgical unit for treatment if MSCC is confirmed
- Patients with suspected MSCC should have an MRI within 24 hours
- Patients with suspected VBM should have an MRI within 7 days
- Initial MRI imaging will be performed at the patient's local hospital unit and is accessed via the local/regional cancer unit pathway, usually via their Acute Oncology contact point

Please refer to NICE Clinical Guideline CG75 (<u>https://www.nice.org.uk/guidance/cg75</u>); Metastatic spinal cord compression: diagnosis and management of adults at risk of and with metastatic spinal cord compression.

Symptoms

Symptoms suggestive of spinal metastases:

- 1. Pain in thoracic or cervical spine
- 2. Progressive lumbar spinal pain
- 3. Severe unremitting lower spinal pain
- 4. Spinal pain aggravated by straining
- 5. Localised spinal tenderness
- 6. Nocturnal spinal pain preventing sleep

Symptoms suggestive of MSCC:

- 1. Pain
 - a) Back pain or nerve root pain either unilateral or bilateral, particularly if associated with alteration in gait
 - b) May be aggravated by movement, coughing or lying flat
 - c) May precede other symptoms by up to 6 weeks
 - d) May be absent in approximately 10% of patients
- 2. Weakness

Motor weakness below the level of the lesion. This may be rapid or slow in onset and can be subtle in the early stages. Descriptions of perceived changes in strength are important.

- 3. Subjective sensory disturbance Often precedes objective physical signs, e.g. "I feel like I am walking on cotton wool." Proprioceptive changes may lead to gait dysfunction perceived as 'poor balance.'
- 4. Bladder/bowel dysfunction. Urinary retention often develops insidiously and generally occurs late.

Signs

The absence of signs does not exclude early spinal cord compression. Investigations should be considered on the basis of history alone in a patient who is at risk.

- Weakness/paraparesis/paraplegia
- Change in sensation below level of lesion (not always complete loss of sensation)
- Reflexes: absent at level of lesion, increased below the lesion
- Clonus
- Painless bladder distension
- Loss of anal tone
- Autonomic dysreflexia for high MSCC (above T8)

N.B. Sensory and reflex changes may occur secondary to other disease processes or previous neurotoxic chemotherapy.

Investigations and Management

The following general principles apply to the investigation and management of VBM and MSCC.

Investigations Urgent

- Contact your local cancer unit/Acute Oncology
- team/spinal surgical team or pathway coordinator as per local policy via their dedicated number to discuss the case and the need for further assessment/evaluation
- Whole spine MRI investigation of choice and shows full extent of disease. This
should be done within 24 hours if MSCC is suspected

- Do not use plain radiographs to diagnose or exclude spinal metastases or MSCC
- If suspected VBM only (i.e., no neurology) then whole spine MRI within 7 days is indicated. Discuss with the patient's Consultant Clinical Oncologist (if they have one), the Consultant Clinical Oncologist linked to the appropriate site-specific MDT (if you do not know who this is contact the site-specific MDT coordinator) or, if all other avenues fail, the Clinical Oncology specialty registrar on-call. Symptomatic VBM is not an emergency in the same way as suspected MSCC

Management

Corticosteroids can be commenced if there is strong clinical suspicion of cord compression and no contraindications, pending definitive investigations. Give dexamethasone 16mg stat orally (with PPI cover), then continue on 16 mg/day until further review. This may give short term improvement while arrangements are being made for investigations and treatment. After surgery or radiotherapy, in consultation with the oncologist, dexamethasone can be reduced, usually over 7-10 days unless neurological function deteriorates. Monitor blood glucose levels while patient is on corticosteroids (peak levels would be expected in early evening after a morning dose).

Surgery can be appropriate in certain situations. The Clinical Oncology team will undertake appropriate assessment and triage of patients with proven MSCC on MRI. Direct approach to the surgical team for patients with pre-existing proven malignant disease and MSCC should not be undertaken. In most cases surgery should be followed by high dose radiotherapy. Adhere to local policy with regards to referral for radiotherapy or surgery (decompression and spinal stabilisation).

Palliative Radiotherapy

This forms the mainstay of treatment in most cases. The Clinical Oncology team will arrange delivery of this after MRI review and patient triage have been undertaken by them.

Chemotherapy

This is rarely used in the acute management of MSCC but may be indicated in future management as MSCC/ VBM reflects a background of progressive cancer.

Active Anti-Cancer Therapy

This may not be appropriate for patients in the late stages of their cancer trajectory, in those who are unfit for travel, MRI scanning or radiotherapy treatment, or who have established paraparesis and are pain free.

Supportive Care

Always make a full holistic care assessment. Pain relief should be offered to all patients.

Venous Thromboembolism Prophylaxis

This should be undertaken following assessment according to local policy.

Rehabilitation

Patient positioning and mobilisation should be undertaken according to patient ability/deficit. Flat bed rest is not routinely advocated, and patients may sit inclined as their pain and sitting balance permit. If safely ambulant, this should be encouraged. Patients should have a physiotherapy and/or occupational therapy assessment to agree an initial rehabilitation plan. Aggressive rehabilitation is often not appropriate (as it will be hampered by the progressive background malignancy with fatigue and limited ability to comply) but fitter patients with residual but reduced function, and at an earlier phase of their cancer trajectory, may benefit from onward referral to local rehabilitation teams (though availability of local services may vary).

Superior Vena Caval Obstruction (SVCO)

Introduction

- Most commonly seen in lung cancer (75-80%)
- Can be the presenting feature of lymphoma, particularly in younger patients
- Regard as emergency as patient's condition may deteriorate rapidly

Symptoms and Signs

- 1. Breathlessness is the most common symptom (>60%)
- 2. Swelling or discolouration of the face and neck 3. Feeling of fullness in the head
- 3. Bending forward or lying flat may aggravate signs and symptoms
- 4. Non-pulsatile raised jugular venous pulse
- 5. Dilated anterior chest wall and neck veins

Investigations

Discuss with radiologist regarding local policy:

- Chest X-ray may reveal widened mediastinum or mass
- Thoracic CT scan is the investigation of choice

Management

Vascular stenting is usual treatment of choice although radiotherapy or chemotherapy may be good alternatives. Chemotherapy may be the treatment of choice in lymphoma and small cell lung carcinoma (if diagnosis previously established). Radiotherapy is useful in patients with non-small cell lung carcinoma.

The evidence for the use of corticosteroids as a holding measure before definitive treatment is lacking. Where used this should be for a limited duration. Recommend discussion with respiratory/oncology.

Recurrent Superior Vena Caval Obstruction

Radiotherapy may be considered. Vascular stents may be replaced. Thrombolysis may be considered if a stent is blocked by thrombus.

Outcome

Treatment often gives useful symptomatic relief.

If the SVCO is untreatable, the patient has a prognosis of days.

Hypercalcaemia of Malignancy

Introduction

- Affects approximately 20-30% of patients with advanced cancer
- Most commonly seen in multiple myeloma, breast, and lung (squamous) carcinomas
- Can also occur in lymphoma, leukaemia, renal and prostate cancer
- Consider in unexplained nausea, vomiting, confusion or constipation
- More commonly due to tumour secretion of parathyroid hormone-related protein rather than bone metastases
- May develop insidiously
- Cancer is usually advanced if hypercalcaemia develops

Symptoms and Signs

Severity of symptoms is more related to the speed of rise in serum calcium rather than the absolute level.

- Non-specific early symptoms: lethargy, malaise, anorexia
- Common symptoms: nausea and confusion
- Other symptoms: constipation, thirst and dehydration
- Late features: drowsiness, fits, coma

Investigations

- Corrected serum calcium
- Urea and electrolytes

Management

Treat if serum calcium is elevated, the patient is symptomatic and it is clinically appropriate.

- Pre and post dose rehydration with 0.9% sodium chloride tailored to the patient's renal function, cardiovascular status and oral intake
- Intravenous bisphosphonate, e.g. zoledronic acid 4mg IV is the agent of choice
- Pamidronate 90mg, or ibandronate between 2 to 4mg are other options. Choice and dose depends on local guidelines and the patient's renal function. Review other medication like thiazides, lithium and Vitamin D supplements

Follow Up

Recheck calcium if symptoms have not improved after 3-4 days.

- Maximal response to bisphosphonates is seen after 6-11 days
- If appropriate, repeat or give a different bisphosphonate if calcium level has not decreased.
- Consider investigating for hyperparathyroidism in selected patients
- For recurrent hypercalcaemia consider intermittent intravenous bisphosphonates or denosumab (unlicensed; consider after discussion with specialist). Patients should have a dental assessment and their dental practice informed, to minimise the risk of osteonecrosis of the jaw

Outcome

- Average duration of response is 3-4 weeks
- Patients should be informed that hypercalcaemia may recur and to monitor for symptoms
- Prognosis depends on the underlying pathology, but refractory hypercalcaemia is a poor prognostic indicator

Lymphoedema

Introduction

Lymphoedema occurs due to the inability of the lymphatic system to maintain normal tissue homeostasis. This results in an accumulation of protein-rich fluid in the subcutaneous tissues. Lymphoedema is one form of chronic oedema. In patients with cancer, lymphoedema is often secondary to the underlying cancer or previous cancer treatment

Characteristic Features

- Oedema
- Chronic inflammation
- Skin changes, e.g. dry skin, thickened tissues (Stemmer's sign)
- · Heaviness and aching in the affected limb
- In early lymphoedema there may be diurnal changes and pitting of the tissues
- With time, and due to chronic inflammatory changes, the tissues may become more fibrosed

General Management

Where available, patients should be referred to specialist lymphoedema clinics.

The core treatment elements are:

- Skin care keep skin intact, clean and well hydrated with non-perfumed emollient (e.g. Diprobase[®], Doublebase[®] or Zerobase[®])
- Compression/support stocking
- Movement and exercise
- Simple lymph drainage, self-massage techniques. Avoid affected limb for any medical procedure where possible, e.g. injection, venepuncture, blood pressure measurement

Management of Cellulitis in Lymphoedema

Comprehensive advice is available in the consensus document from the Lymphoedema Support Network <u>www.lymphoedema.org/</u> and British Lymphology Society <u>www.thebls.com</u>

Treat early, monitor closely and continue antibiotics for at least 14 days after clinical improvement is observed.

Oral flucloxacillin 500mg q.d.s. (clarithromycin 500mg b.d. if penicillin allergic). Acute infection is usually painful; review analgesics. Avoid compression garments and NSAIDs in acute attack. If the patient develops systemic symptoms, IV antibiotics may be required; seek specialist advice.

Recurrent Cellulitis

Antibiotic prophylaxis should be considered if the patient has had 2 or more episodes of cellulitis per year. Penicillin V 500mg daily (erythromycin 500mg daily if penicillin allergic) first line. Consult and refer for specialist advice <u>https://www.thebls.com/</u> and refer for specialist advice.

Mouth Care

Essential Mouth Care

Ensure that the patient is asked about mouth problems and that the mouth is examined.

- If the patient is conscious, support them to brush their teeth with fluoride-containing toothpaste. Perform mouth care at least twice a day
- Dentures should be cleaned at least daily with a denture brush or toothbrush, and soap, or denture tablets. Dentures should be removed at night and stored in water
- If the person is unconscious, hourly care is recommended using a soft small toothbrush with water or non-foaming toothpaste to clean the teeth, gums, tongue and mouth
- Apply dry mouth gel (e.g. Biotene[®] oral balance gel) 2 4 hourly when required to the lips, tongue and oral cavities to help keep mouth moist

Dry Mouth

Regular essential mouth care is the most important part of management.

Look for and address reversible causes:

- Review medication as many drugs cause dry mouth (e.g. anticholinergics, opioids, corticosteroids and antiemetics), consider dehydration, consider humidifying oxygen
- Recognise and support patients with anxiety
- Frequent sips of cold unsweetened drink if the person is able to swallow
- · Consider the use of crushed ice and ice pops
- Avoid acidic products that can lead to dental caries and oral candidiasis
- Consider sugar-free chewing gum to stimulate saliva production. Chewing gum is as effective as and preferred to artificial saliva.
- Artificial saliva substitutes: Biotene oral-balance[®], Oralieve gel/spray, AS Saliva Orthana[®] (contains pork)

Coated Tongue

- Often caused by inadequate salivary function so associated with dry mouth
- Manage as for dry mouth
- Increase frequency of mouth care. Hourly rinsing with water or saline can be helpful
- Gentle brushing of the tongue with a soft tooth brush several times a day can aid debridement/ removal of previously applied lubricants
- Reassess regularly

Sore or Ulcerated Mouth

- Regular essential mouth care is the most important part of management
- Identify and treat the cause where possible (e.g. trauma, aphthous ulcers, infection, malignancy, chemotherapy or radiotherapy induced, nutritional deficiencies, GI conditions, haematopoietic disorders and drug-induced)
- Increase frequency of mouth care. Hourly rinsing with water or saline can be helpful
- A coating agent such as Gelclair[®], alcohol free mouthwash can be useful and may help with eating when used 30-60 minutes before meals

- Topical analgesia options: paracetamol mouth rinse, benzydamine hydrochloride (Difflam[®]), choline salicylate (Bonjela[®])
- If not responsive to the above measures, consider use of topical anaesthetics and apply directly to painful area, e.g. lidocaine (Xylocaine[®]) 10% spray applied using cotton bud p.r.n. Avoid anaesthesia to pharynx before meals/drinks
- For severe oral pain, consider the combined use of topical and systemic preparations. Seek specialist advice
- Reassess regularly

Oral Candidiasis

Can present as a dry mouth, loss of taste, reddened tongue, soreness, dysphagia, angular cheilitis and also can be asymptomatic.

- Regular essential mouth care management reduces the chance of infection and should be continued
- Look for and treat reversible causes: immunosuppression, steroid use (oral/inhaled), dry mouth, dehydration, poor oral hygiene, mucosal damage
- Use a new toothbrush
- Consult local guidelines for use of antifungals/ check for drug interactions. Options include:
 - Nystatin oral suspension 100,000 units/ml, 5ml q.d.s. for 7 days; hold in mouth for 1 minute, then swallow. Note: Nystatin and chlorhexidine mouthwash should not be used at the same time as they will deactivate each other. Use an hour apart
 - Fluconazole (capsules or, if cannot take, suspension) 50mg daily for 7 days. (May need higher doses/longer courses for immunosuppressed patients)
- For patients with dentures ensure dentures are thoroughly cleaned and soaked in appropriate antiseptic (e.g. chlorhexidine) for 15 minutes then rinsed in water
- Dispose of toothbrush following completion of drug treatment

Taste Disturbance

- Regular essential mouth care is the most important part of management
- Address dry mouth and infection
- Maintain nutrition where possible and refer to a dietician

The Last Days of Life

Recognition of imminent death is important. It allows withdrawal of unnecessary treatments and preparation of the patient and family/carers for death. This phase is often heralded by a more rapid deterioration in the patient's general condition. It can be difficult to recognise. Consider if there are any potentially reversible causes for the patient's condition, e.g. infection, opioid toxicity, metabolic abnormalities (uraemia, hypercalcaemia) and if it is appropriate to actively manage these and if treatment is feasible and acceptable to the person/or consistent with their pre-existing goals and priorities if they lack capacity to engage in decision making. In a patient with an advanced illness the following symptoms and signs may indicate that the prognosis is short:

- Profound weakness
- Confined to bed for most of the day
- Drowsy for extended periods
- Disorientated
- Severely limited attention span
- Loss of interest in food and drink
- Too weak to swallow medication

Actions

- 1. Sensitively check the understanding of patient and family/carers and explain the plan of care.
- 2. Negotiate appropriate treatment and advance care plans with the patient if they have capacity. Record in patient records and on ReSPECT. Check if there is a valid Advance Decision to Refuse Treatment (ADRT) or if the patient has a Lasting Power of Attorney (LPA) for health and welfare.
- If the patient does not have capacity, clinical decisions must be made in the patient's best interests in line with the Mental Capacity Act. Family, carers and other healthcare professionals should be consulted. The role of the family is to advise on what the patient would have wanted for him/herself. Consider use of Deprivation of Liberty Safeguards (DOLS)/Liberty Protection Safeguards (LPS).
- 4. Establish the patient's preferred place of care and preferred place of death. This should take into account the needs and wishes of the patient and the family/carers.
- 5. Fast Track/NHS Continuing Care Funding Form or equivalent needs to be completed for patients wishing to be cared for in a home or care home setting.
- 6. CPR recommendation should be reviewed as part of broader discussion of treatment escalation in inpatient and community settings. In accordance with local policy, complete a transferable regional DNACPR or ReSPECT form if this has not already been completed. Before a DNACPR recommendation is formally implemented the person with capacity or their representatives (if they lack capacity) must be given the opportunity to discuss the approach to CPR. This applies even if CPR is not deemed to be of clinical benefit.
- 7. If at home/care home ensure out of hours Handover has been completed and the electronic record/notes updated as per local policy, e.g. EPaCCS/local register/single point of access.

- 8. Ensure that the patient and family/carers have the telephone numbers for NHS 111, out of hours palliative care line, locally available services, the community palliative care team, local hospice and their Primary Care Team.
- 9. Professional carers may need to acknowledge and share their own feelings. Mutual support and teamwork are important.
- 10. Ensure all anticipatory medications prescribed. If the patient is being discharged ensure they are sent with the patient. This ensures the patient does not have to wait for medication if they start to develop symptoms.
- 11. Ensure all individuals involved in the patient's care are aware of their condition, such as the Primary Care Team, local hospices and community palliative care teams.
- 12. Ensure regular review by nursing and medical staff, including night care, as agreed with the patient and their family.
- 13. When the patient is in the last days/hours of life, support with an individualised care plan as per local policy.

Physical Care

Care and support

- 1. If a patient is to be discharged home from hospital to die, ensure the Primary Care Team and where appropriate, the community palliative care team are aware.
- 2. Adequate day and night nursing support needs to be arranged. Consider night sitters.
- 3. Involve family/carers in practical care as much as they wish and discuss the plan of care.

Priorities of care include:

- Assess regularly for common symptoms at the end of life: pain, agitation, respiratory secretions, nausea and vomiting and breathlessness
- Treat dry mouth with meticulous and regular mouth care
- Immobility and pressure areas bed, mattress, positioning needs to be assessed
- Continence consider catheter, convene or pads and monitor for signs of retention
- Bowel care assess for bowel problems that may cause discomfort, such as constipation or diarrhoea
- Assess the psychological, religious, cultural and spiritual care needs of the patient and family

Hydration and Nutrition

A reduced need for food and fluids is part of the normal dying process and patients should be supported to take food and fluids by mouth for as long as tolerated. Symptoms of thirst/dry mouth are often due to mouth breathing or medication/oxygen therapy and good mouth care is essential. Adults in the last days of life need to have their hydration status assessed daily and have a discussion about the risks and benefits of hydration options.

(NICE: Care of the dying adult in last days of life; pathway QS144, 2017).

For many patients, the use of clinically assisted (artificial) hydration will not be of benefit

and decisions about their use should be made in a patient's best interests. If clinically assisted artificial hydration or nutritional support is in place, review the rate/volume/route according to individual need. The possible benefits of withdrawing or reducing clinically assisted hydration/nutrition include reduced vomiting, incontinence, and reduced painful venepuncture. If indicated, depending on setting, fluids can be administered subcutaneously; usually 1L/24 hours and preferably site in the abdomen. Monitor for uncomfortable fluid accumulation at the infusion site.

Medications

Reassess the indications and potential benefits in the context of the terminal phase for ALL medications. Only continue medication needed for symptom management (these might include disease-directed therapies such as insulin). If the oral route is not appropriate, the subcutaneous, transdermal or rectal routes can be used for many symptom management drugs.

Antibiotics are not commonly indicated during the last weeks of life. Shared decision making and goals of care agreement are essential. Non-pharmacological and symptom directed approaches should be used. The main indication is when symptomatic/pain relief from UTI is needed. See <u>SAPG guidance</u>.

For patients who are diabetic on insulin, see <u>guidance</u> and seek advice from diabetic team. When in the last hours/days of life refer to local symptom management guidelines where available. Ensure anticipatory medications are prescribed and available for the common symptoms which may develop in the last hours or days of life: pain, terminal restlessness, respiratory tract secretions, breathlessness, nausea and vomiting.

Terminal Restlessness

Assess the patient carefully. Restlessness can occur at the end of life but there may be a precipitant, therefore look for evidence of:

- Physical discomfort pain related to the underlying condition, urinary retention, faecal impaction or to a new event, e.g. haemorrhage, malfunctioning syringe pump
- Respiratory distress breathlessness, cough, tracheal obstruction
- Neurological problems fits, hallucinations, myoclonic jerks, motor restlessness. Remember, these may be caused by medicines (including opioids and anti-emetics)
- Psychological distress (see below)
- Delirium

If there are no reversible precipitating factors or psychosis/delirium, midazolam is the drug of choice (see syringe pump section, page 50).

After non-pharmacological measures, haloperidol is indicated for delirium. If midazolam alone is not effective, consider adding haloperidol or levomepromazine.

Respiratory Tract Secretions or 'Death Rattle'

This is a rattling noise produced by the movement of secretions in the upper airways in

patients who are too weak to expectorate effectively. Relatives and carers may find this distressing. It is important to explain to the relatives/carers that this is unlikely to be causing distress to the patient.

- Repositioning of the patient and postural drainage may help
- Anti-secretory drugs can be used (see syringe pump section or local symptom management guidelines)
- Prompt drug treatment is required

Distressing Terminal Events

Events such as haemorrhage, fits or tracheal obstruction are unusual and can often be anticipated and a management plan discussed with nursing staff in advance. Prescribe appropriate p.r.n. medication, e.g. midazolam, to relieve distress and to sedate if necessary. Seek advice from palliative care team if unsure. If possible, try and discuss the possibility of such events with family or carers if they are thought to be more likely, e.g. known head and neck tumours near major vessels. This may allow for better preparation if such an event were to happen.

Do Not Attempt Cardiopulmonary Resuscitation (DNACPR) Decisions

If a patient is in the last days of life, cardiopulmonary resuscitation (CPR) will not be of clinical benefit. The resuscitation status of the patient should be discussed within the clinical team and documented as per local policy. It is a legal obligation to offer a discussion re DNACPR and to explain to the patient and their carers: why CPR will not be attempted; that the focus of care is on palliation and comfort and in the home setting, to ensure that family members know what to do when the patient dies.

Advise them to keep the DNACPR/ ReSPECT form somewhere safe so that it can be shown to all healthcare providers.

Recommended Summary Plan for Emergency Care and Treatment (ReSPECT)

ReSPECT forms will eventually replace the regional DNACPR nationally but please check locally for current practice. ReSPECT forms document personalised recommendations for a person's clinical care in a future emergency in which they are unable to make or express choices. It provides health and care professionals responding to that emergency with a summary of recommendations to help them to make immediate decisions about that person's care and treatment. ReSPECT can be complementary to a wider process of advance/anticipatory care planning.

Implantable Cardiac Defibrillator (ICD) Deactivation

Implantable cardiac defibrillators (ICD) and cardioverter resynchronisation devices with a defibrillator (CRT-D) may need to be deactivated during the palliative phase to prevent painful shocks being delivered unnecessarily at the end of life. This may already have been discussed at the time of device insertion. The defibrillator function can be turned off separately to the pacemaker and resynchronisation functions and turning it off does not

cause death. Deactivation should be considered as part of a wider advance care planning conversation and is different from a DNACPR decision.

Devices can be turned off by the local hospital Cardiac Physiology team, and most teams will be able to arrange this urgently. In an emergency where a patient is receiving painful and inappropriate shocks, devices can be temporarily deactivated by taping a ring magnet securely over the device whilst arranging for permanent deactivation. Ring magnets should not be applied to pacemakers without an ICD function as this may cause the pacemaker to fire faster than needed. Ring magnets are stored in a variety of locations depending on your local area; if advice is needed you can contact your local cardiac physiology department or palliative care consultant.

Psychological and Spiritual Care of Patient and Family

The patient, relative and carer should be given the opportunity to discuss what is important to them at this time. Decisions about the plan of care should be communicated to the patient where appropriate and to the relative or carer. The patient may be anxious for themselves or others and addressing psychological and spiritual needs may alleviate symptoms of agitation.

Consider barriers to communication such as hearing, vision and speech difficulties, learning disabilities, dementia, neurological conditions, language barriers and confusion. The relative or carer may know how specific signs indicate distress if the patient is unable to articulate their own concerns.

Encourage open communication and explore fears and concerns:

- Facilitate expressions of emotions
- Involve children and those with learning disabilities
- Remember spiritual care and religious needs (offer to contact chaplain, priest, rabbi, imam etc. if appropriate)
- Consider music, art, poetry, reading, photographs or something else that has been important to the well-being of the patient

Care After Death

Practical and legal aspects to attend to after death.

Arrangements may vary depending on place of death and local bereavement service provision.

- Inform relatives when referral to the coroner might be necessary, e.g. mesothelioma. It is preferable to do this before the patient's death
- Ensure prompt verification of death, personal care after death and provision of death certificate
- Provide information about the role of the undertaker and how to register a death
- For deaths not occurring in the patient's own home, ensure patient's Primary Care Team is informed within 24 hours
- For deaths occurring at home, ensure planned visits are cancelled and arrangements made to return equipment

• Ensure hospital appointments (and transport) are cancelled and hospitals/consultants involved with the patient's care are informed

Bereavement

It is good practice when someone has died, to provide written information about the common feelings of grief and available support and to identify those at increased risk in bereavement. Risk factors include:

- previous multiple losses or recent losses
- ambivalent relationship
- dependent children involved
- bereaved parent
- previous psychological or psychiatric problems or substance abuse
- · people living alone or feeling unsupported

Seek advice from colleagues and the relative's Primary Care Team, with their permission, if one or more of these risk factors are present.

It is good practice to review how someone is coping 6-8 weeks after the death.

Syringe Pump Principles

In palliative care a syringe pump or driver is a way of administering medication continuously via the subcutaneous route when the patient is unable to swallow or absorb oral drugs due to:

• Persistent vomiting, intestinal obstruction, dysphagia, profound weakness, unconsciousness or mouth, throat and oesophageal lesions

Where indicated, syringe pumps can be used for a short period for symptom control, or for longer in the terminal phase. It is the responsibility of healthcare staff to discuss with the patient and those identified as important to them:

- The reason for the use of a syringe pump. Other than in exceptional circumstances, this should be done before it is used
- The likely side effects of specific interventions, especially those that may make the person sleepy, must be discussed with the dying person to enable them to make informed decisions, and explained to those important to the dying person if they so wish
- All medications, including anticipatory medicines, must be targeted at specific symptoms, have a clinical rationale for the starting dose, be regularly reviewed, and adjusted as needed for effect

Considerations

- Doses of medication are calculated on the basis of patients' previous requirements
- Following commencement of a syringe pump it will be many hours before therapeutic levels are achieved, so consider giving a stat dose of medication equivalent to the normal breakthrough/p.r.n. dose
- Syringe pumps require careful monitoring and should be prescribed on prescription/syringe pump charts as per local syringe pump policies
- Inadequate pain control is not an indication for syringe pump use unless there is reason to believe oral medications are not being absorbed or the patient has nausea or vomiting
- Recommended sites for insertion of the subcutaneous cannula are the anterior chest wall, upper arms, abdominal wall and thighs

NOTE:

A variety of models of ambulatory infusion devices (syringe pumps) are in use. T34 syringe pumps are widely used in the region. Alternative MHRA approved syringe pumps may also be available. Please follow your local syringe pump policy.

Subcutaneous Medications & Infusions

COMMON MEDICINE DOSAGES FOR SUBCUTANEOUS MEDICATIONS & INFUSIONS

All the medicines on the following two pages (52-53) can be given as subcutaneous infusions in a syringe pump.

Remember to prescribe subcutaneous p.r.n. medication. If using more than one drug in a syringe pump, check compatibilities with current PCF, pharmacy, or SPCT.

To convert from oral morphine see page 21. For other opioids, seek specialist advice. If symptoms are not controlled, other regimens may be needed. Seek specialist advice.

Table of Commonly Used Subcutaneous Medications

Commonly used subcutaneous medications	Usual starting p.r.n. dose	Usual 24-hour dose range	Usual ampoule size information	Comments			
PAIN AND BREATHLESSNESS							
Morphine	1mg to 2.5mg for opioid naive 1/6th of the 24-hour SC dose	Between 5mg and 10mg/24 hours if opioid naïve. 1/3 to 1/2 of previous 24-hour oral morphine dose/24 hours CSCI. See local guidelines. Seek advice.	10mg/1ml, 15mg/1ml, 20mg/1ml, 30mg/1ml, 20mg/2ml, 30mg/2ml, 40mg/2ml, 60mg/2ml	 Seek advice if: patient requiring rapidly escalating doses patient in renal failure 			
Diamorphine – Rarely used and generally reserved for high opioid doses	1mg to 2.5mg for opioid naive 1/6th of the 24-hour SC dose	1/3 of previous 24-hour oral morphine dose/24 hours CSCI. See local guidelines. Seek advice.	5mg, 10mg, 30mg, 100mg, 500mg	Seek advice if: • Thinking of using • rapidly escalating doses • renal failure			
NAUSEA AND VOMITING							
Haloperidol	500 micrograms	1mg to 5mg/24 hours CSCI	5mg/1ml	Haloperidol has anxiolytic and sedative properties			
Cyclizine	50mg (max t.d.s.)	100mg-150mg CSCI (max dose) Mix with water for injections	50mg/1ml	May be incompatible with hyoscine butylbromide			
Metoclopramide	10mg (max t.d.s.)	30mg to 60mg/24 hours CSCI	10mg/2ml	Prokinetic effect antagonised by cyclizine			
Levomepromazine	2.5-6.25mg	6.25mg to 25mg/24 hours CSCI	25mg/1ml	Can be sedating			

Commonly used subcutaneous medications	Usual starting p.r.n. dose	Usual 24-hour dose range	Usual ampoule size information	Comments			
RESPIRATORY SECRETIONS – Early intervention for "death rattle" is required							
Hyoscine butylbromide (Buscopan®)	20mg	Between 60mg to 120mg/24 hours CSCI	20mg/1ml	Not sedating may be incompatible with cyclizine			
Glycopyrronium	400 micrograms	600 micrograms to 1.2mg/24 hours CSCI	200 micrograms/1ml, 600 microgram/3ml	Not sedating			
Hyoscine hydrobromide	400 micrograms	1.2mg to 2.0mg (max dose 2.4mg/24 hours) CSCI	400 micrograms/1ml, 600 micrograms/1ml	Can be sedating (use other anti- secretory drugs 1st line)			
AGITATION – Consider haloperidol if patient is suffering from agitation and delirium							
Midazolam	2mg to 5mg	10mg to 20mg/24 hours CSCI (starting dose). May be increased up to 80mg/24 hours according to response. Seek SPCT advice.	10mg/2ml. Other preparations are available, store separately	Muscle relaxant, anxiolytic and anticonvulsant (see below). If ineffective seek specialist advice.			
Haloperidol	500 micrograms	1mg to 5mg/24hours CSCI	5mg/1ml	Haloperidol has anxiolytic and sedative properties			
ANTICONVULSANT							
Midazolam	10mg	20mg to 80mg/24 hours CSCI to replace oral anti-convulsants may be required.	10mg/2ml. Store high and low strength midazolam separately.	Seek SPCT advice regarding management of prolonged fits			