

TRUST WIDE DOCUMENT

	Policy
DOCUMENT TITLE:	Policy for use of the Ambulatory Syringe Pump in adults (Palliative Care)
DOCUMENT NUMBER	ELHT/CP22 Version 6.4
DOCUMENT REPLACES	ELHT/CP22 Version 6.3
LEAD EXECUTIVE DIRECTOR DGM	Director of Nursing
AUTHORS	Syringe Pump Policy Task and Finish Group chaired by Palliative Medicine Consultant

TARGET AUDIENCE	Medical and Nursing Staff		
DOCUMENT PURPOSE	 To provide a clear governance framework to ensure a safe consistent approach to the use of the Ambulatory Syrin Pump. To provide easily accessible information about the commedicines used in a Syringe Pump. 		
To be read in conjunction with	 Palliative Care Clinical Practice Summary. Guidance on consensus approaches to managing palliative care symptoms. Lancashire and South Cumbria consensus guidance – 2nd Edition, November 2021 CO64 Medicines Management Policy IC24 Aseptic non touch technique (ANTT) policy SOP115 Procedure for setting up and using an Ambulatory Syringe Pump 2nd edition SOP115 Procedure for setting up and using an Ambulatory Syringe Pump 3rd edition SOP079 Procedure for the Administration of Subcutaneous PRN Medication Using a Prescribed Range of Doses for Symptoms in the Last Days of Life 		

SUPPORTING REFERENCES	 Royal Pharmaceutical Society – Professional guidance on the safe and secure handling of medicine, Dec 2018 Royal Pharmaceutical Society/Royal College of Nursing: Professional guidance on the administration of medicines in healthcare settings, Jan 2019 Advisory guidance – administration of medicines by Nursing Associates – Health Education England, Dec 2017 Dickman et al (2016) The Syringe Driver, 4th edition, Oxford Press T. Mitten (2000) Subcutaneous drug infusions, a review of problems and solutions. International Journal of Palliative Nursing Vol 7 No. 2 Palliative Care Formulary online, https://www.medicinescomplete.com/#/ accessed June 2021 Twycross R., Wilcock A., (2001) Symptom Management in Advanced Cancer 3rd edition Radcliffe Medical Press Oxon The Health and Safety (Sharp Instruments in Healthcare) Regulations 2013 British National Formulary online, accessed June 2021
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CONSULTATION				
	Committee/Group Date			
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Approval Committee	TWQG ELMMB Nursing & Midwifery Leaders Forum CIC CQSB January 2022 Sept 2023			
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AMENDMENTS	 SOP removed to be stand alone document – policy amended to reflect this Changes made regarding how to prescribe a syringe pump with transition to Electronic Patient Record (EPR) Sept 2023 to reflect the changes to the role of the 2nd checker within the District Nursing service (HCA/AP) Section 5 and Appendix 3. 			

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1. Introduction

- 1.1. A Syringe Pump is a portable battery-operated device that is used to deliver a continuous subcutaneous infusion of medicines.
- 1.2. The Syringe Pump is a minimally invasive route of medicine administration commonly used in Palliative Care.
- 1.3. The Ambulatory Syringe Pump is the mode recommended for use in Palliative Care.

 This policy applies only to the Ambulatory Syringe Pump in adults.
- 1.4. A Syringe Pump can pose serious risk to human life if used incorrectly.

2. Purpose

- 2.1. To provide a clear governance framework to ensure a safe and consistent approach to the use of the Ambulatory Syringe Pump in adults.
- 2.2. To provide easily accessible information about the common medicines used in a Syringe Pump.

3. Scope

- 3.1. All healthcare professionals in ELHT prescribing, setting up, administering or monitoring medicines being given by an Ambulatory Syringe Pump to adults.
- 3.2. This policy can be used by other healthcare providers across Pennine Lancashire but each organisation is responsible for ensuring they have appropriate training and governance processes in place.
- 3.3. ELHT will not accept any liability or responsibility for care delivered by non-ELHT staff.
- 3.4. Information provided about individual medicines is for guidance purposes. It is not fully comprehensive and is subject to change.
- 3.5. Individual prescribers are responsible for ensuring they prescribe in line with the most up to date guidance available and have appropriate knowledge and understanding of the medicines they prescribe.

4. Roles and responsibilities (ELHT specific)

4.1. The Trust recognises that the Ambulatory Syringe Pump is commonly used to deliver medicines to palliative care patients and that all staff using these Syringe Pumps need to have adequate knowledge and training to do so safely.

4.2. The Chief Executive and Trust Board

- 4.2.1. The Chief Executive is responsible for:
- Ensuring that this policy adheres to professional guidance.
- Ensuring this policy and supporting documentation is reviewed every three years.
- Ensuring appropriate leadership and governance arrangements are in place to enable staff to comply with this policy.
- Ensuring this policy is agreed and monitored by the organisation's governance process.
- 4.2.2. The Trust Board will receive a report at least once a year including audit results related to end of life care.

4.3. Divisional General Managers, Divisional Directors and Divisional Directors of Nursing and other Managers of Services

- 4.3.1. Directors and Managers are responsible for the care provided within their services.
- 4.3.2. They must ensure that:
- Staff are aware of the policy and how to access it.
- Staff are required and enabled to acquire and maintain necessary competencies to safely use an Ambulatory Syringe Pump.
- Appropriate leadership and governance arrangements are in place to implement the policy and to monitor the safe use of the Ambulatory Syringe Pump.
- Incidents that occur involving the Ambulatory Syringe Pump are reported using the Trust incident reporting system.

4.4. All Health Care staff setting up, administering or monitoring the delivery of medication using an Ambulatory Syringe Pump

- 4.4.1. The responsibility for the safe and effective use of an Ambulatory Syringe Pump lies with the ward staff for patients in hospital, the GP and community staff for patients in community settings.
- 4.5. All Health Care staff prescribing medicines to be given by a Ambulatory Syringe Pump

- 4.5.1. Must prescribe within their competence and experience.
- 4.5.2. Must seek advice from Pharmacy or the Specialist Palliative Care Team if required. A 24hour Specialist Palliative Care Advice Line is available for professionals; see 4.7.2.
- 4.5.3. All information provided around medication in this document is for guidance purposes only. The prescriber is responsible for ensuring prescribing is carried out in line with the most up to date guidance.
- 4.5.4. Whilst every effort has been made to ensure accuracy in this document responsibility remains with the prescriber.
- 4.5.5. Should understand that although the medicines in this policy have marketing authorisation (are licensed) their use within a Syringe Pump is often beyond the specifications of the marketing authorisation, also known as off-licence/off-label prescribing. See C064 Medicines Management Policy section on unlicensed and "off-label" medicines.
 - The marketing authorisation regulates the marketing activities of the Pharmaceutical Industry, not the activities of the prescriber. Clinical experience may reveal other indications (i.e., off-label use).
 - The use of medicines beyond and without marketing authorisation in palliative care is both necessary and common and should be seen as a legitimate aspect of clinical practice.
 - Health Care Professionals involved in prescribing medicines beyond or without marketing authorisation should select those medicines that offer the best balance of benefit against harm for any given patient. Prescribers should use the resources available to ensure prescribing is evidence based.
 - Choice of treatment requires partnership between patients and health professionals, and informed consent should be obtained, whenever possible.
 This should occur before prescribing any medicine but particularly when medicines are being prescribed which are beyond or without marketing authorisation.
 - Patients should be offered accurate, clear, specific and appropriate information that meets their needs about the use medicines beyond or without a marketing authorisation in accordance with professional regulatory body guidance.
 - The information needs of carers and other health professionals involved in the care of the patient should also be considered and met as appropriate.

The use of information cards or leaflets may help with this. It is often unnecessary to take additional steps when recommending medicines beyond or without marketing authorisation.

 For further information see Palliative Care Formulary online, https://www.medicinescomplete.com/#/.

4.6. Pharmacy Team

- 4.6.1. The core service provided by the pharmacy team includes clinical pharmacy, medicines procurement, medicines information and counselling.
- 4.6.2. Pharmacy will advise on the prescription writing and compatibility of medicines in the Syringe Pump.
- 4.6.3. Ward pharmacists will carry out clinical checks for stability and compatibility of Syringe Pumps. This includes drug, diluent and final dose concentration checks.
- 4.6.4. Pharmacy will advise on the ordering, storage, administration, disposal and record keeping of medicines.
- 4.6.5. Pharmacy will ensure timely provision of medicine and help in assessing appropriateness of medicine orders.
- 4.6.6. Pharmacy offer a limited, centrally delivered aseptic additive service and emergency on-call service via hospital switchboard.

4.6.7. Contact details for Pharmacy:

Medicines	Mon-Fri	Telephone:
Information	08.30am-5.00pm	01282 803004/Ext. 13004
Pharmacy Aseptic Unit	Mon-Fri	Telephone:
	08.00am-4.30pm	01254 734680/Ext. 84680
	Sat/Sun 09.00am-1.00pm	
Pharmacy Dispensary	Mon-Fri	Telephone:
RBH	08.30am-5.00pm	01254 733507/Ext. 83507
	Sat/Sun 09.00am-4.00pm	
Pharmacy Dispensary	Mon-Fri 09.00am–	Telephone:
BGH	5.00pm	01282 804338/Ext. 14338

4.7. Specialist Palliative Care Team

- 4.7.1. The Specialist Palliative Care Team supports hospital and community staff with the use of the Syringe Pumps in palliative care. Health Care Staff should refer to the Specialist Palliative Care Team if:
 - Advice is needed about symptom management in Palliative Care patients or when initial measures have failed to provide adequate relief within 24 hours.
 - Advice is needed regarding the prescription of medicines to be given in a Syringe Pump.
 - Advice is needed about the set up or monitoring of a Syringe Pump.
- 4.7.2. The Specialist Palliative Care Team work Mon-Fri excluding bank holidays and weekends. Specialist Palliative Care telephone advice is available 24 hours per day, 7 days per week.

Contact details for Specialist Palliative Care advice:

Hospital	Mon-Fri	08.30-16.30	Ext 82316/82652
Community	Mon-Fri	08.30-16.30	Ext 86326/86428
Out of Hours	07730 639399 (advice line based at East Lancashire Hospice)		

5. Training and Education

- 5.1. Staff using Syringe Pumps are responsible for ensuring that they develop and maintain the skills and knowledge required to fulfil their professional role.
- 5.2. All staff setting up and monitoring the administration of medicines by a Syringe Pump must have attended an initial training session.
- 5.3. All staff using Syringe Pumps must attend an annual update thereafter.
- 5.4. Training needs should be identified and met as part of personal development review/appraisal processes.
- 5.5. All Trust Divisions will enable and support staff to acquire and maintain necessary skills and competencies to ensure safe use of the Ambulatory Syringe Pump.
- 5.6. The Trust provides Syringe Pump training and education sessions for nursing staff.
- 5.7. Training can be requested and organised for other groups such as medical staff by contacting the Specialist Palliative Care Team.
- 5.8. Additional Training requirements for Assistant Practitioners and Health Care Assistants. This additional training is required for Assistant Practitioners and Health Care Assistants to be able to be the second checker alongside a Registered Nurse
 - 5.8.1. Attendance at a face-to-face training session delivered by the District Nurse Practice Educator.

- 5.8.2. Completion of ELHT Syringe Pump e-learning module
- 5.8.3. Completion of ELHT Medicines Management Human Factor training.
- 5.8.4. Undertaking and completion of a Competency Framework signed off by a Senior Nurse within the DN Service (Appendix 3)
- 5.8.5. Attendance at an annual update by a syringe pump trainer in the team.

6. Monitoring and Audit

6.1. Audit of the safe and effective use of the Ambulatory Syringe Pump will be undertaken at least biennially within the Trust. Results will be fed back to the ELHT End of Life Care Strategy Group and the nominated Trust Board members.

Monitoring Mechanism:

Measuring and monitoring compliance with the effective implementation of this procedural document is best practice and a key strand of its successful delivery. Hence, the author(s) of this procedural document has/have clearly set out how compliance with its appropriate implementation will be measured or monitored. This also includes the timescale, tool(s)/methodology and frequency as well as the responsible committee/group for monitoring its compliance and gaining assurance.

Aspect of compliance being measured or monitored.	Individual responsible for the monitoring	Tool and method of monitoring	Frequency of monitoring	Responsible Group or Committee for monitoring
Use of syringe pumps in practice	Specialist Palliative care team audit lead	Audit tool	Every 2 years	Specialist Palliative Care Directorate

Indications for use of a Syringe Pump in Palliative Care

Palliative care patients often experience multiple symptoms that require the use of more than one medicine. If a patient's condition changes so that the oral route is no longer available, the Syringe Pump can be used to support continued symptom control.

A Syringe Pump is the chosen method for the administration of medicines when other routes are inappropriate due to:

- Nausea and vomiting
- Dysphagia
- Severe weakness/cachexia
- Unconsciousness
- Gastrointestinal problems e.g. diarrhoea, bowel obstruction
- Inability to administer medication via oral route i.e. Head/neck cancers
- Malabsorption
- Care in the last days and hours of life a Syringe Pump should only be started in the
 last hours or days of life if it is indicated for symptom management. Not all dying
 patients will require a syringe pump.

Many palliative care patients will require administration of 'as required' (prn) subcutaneous medication for symptom management.

If more than 2 or 3 doses of any 'as required' (prn) subcutaneous medication are required for symptom control over 24 hours, consider using a Syringe Pump.

In a patient with a Syringe Pump in place consider increasing the doses if more than 2 or 3 doses of any 'as required' (prn) medication is required for symptom control over 24 hours.

Advantages in the use of a subcutaneous Syringe Pump

- Increased patient comfort when oral route not available.
- Avoids repeated injections.
- Plasma concentration levels of medicines remain constant.
- Maintains patient's independence and mobility as pump is lightweight and portable.
- Ability to control multiple symptoms by infusing a combination of medicines.
- Accurate absorption.

Disadvantages in the use of a Syringe Pump

- Irritation, erythema or swelling can occur at the infusion site which may interfere with rate and absorption.
- Precipitation of medicines can occur. There is a lack of compatibility data for some mixtures.
- May be perceived as a 'terminal' event by patients and carers.

Guidance on prescribing opioid doses for a Syringe Pump and as required (prn)

Opioids given subcutaneously via a Syringe Pump are more potent than opioids administered orally. The dose of the opioid prescribed must therefore be adjusted when switching from oral to subcutaneous administration.

Different opioids also vary in their potency and therefore the dose prescribed must be adjusted when switching between different opioid medicines. For information on equivalent doses when changing the route of administration or the opioid given please refer to page 8 of the Lancashire and South Cumbria Consensus Guidance – Palliative Care Clinical Practice Summary November 2021. For doses or drugs out with this guidance please contact the Specialist Palliative Care Team – see page 67 for contact details.

Additional 'as required' (prn) medication

In addition to the medication prescribed in a Syringe Pump it is often necessary to prescribe other subcutaneous medicines for symptoms management that are available if required.

For breakthrough pain

It is best practice to prescribe additional subcutaneous doses of opioid analgesia equal
to 1/6th of the total daily dose of opioid for breakthrough pain. Refer to opioid
conversion charts on page 8 of the Lancashire and South Cumbria Consensus
Guidance – Palliative Care Clinical Practice Summary for further details of
recommended doses.

Other symptoms

- Please refer to the Lancashire and South Cumbria Consensus Guidance –
 Palliative Care Clinical Practice Summary for the management of symptoms in the last days of life for guidance on appropriate doses of other medicines
 - East Lancashire Health Economy Medicines Management Board Website
 Guidelines -> Palliative Care & Syringe Pump Guidelines
 (http://www.elmmb.nhs.uk/guidelines/palliative-care/)
 - ELHT Intranet —> Clinical Information —> Palliative Care/EoLC —>
 Symptom Control and Prescribing or Syringe Pump folder

Switching between other analgesic preparations and the Syringe Pump

1. Changing from twice daily modified release oral opioids to the Syringe Pump.

The Syringe Pump can be started when the next dose of oral modified release opioid is due. In some circumstances it may be appropriate to start the pump sooner. Seek specialist advice.

- 2. Concomitant use of Fentanyl or Buprenorphine patches with opioids in a Syringe Pump.
 - If a patient has a Fentanyl or Buprenorphine patch in situ and additional analgesia is required by a Syringe Pump the patch should be left in situ.
 - Maintain the current patch strength.
 - Continue to change the patch at the recommended interval.
 - When calculating the 'as required' (prn) dose for patients on a Syringe Pump and a Fentanyl or Buprenorphine patch take into account both methods of opioid delivery.
 - Calculate the breakthrough dose of 'as required' (prn) subcutaneous analgesia by ADDING the amount required for the Fentanyl or Buprenorphine patch to the amount required for the opioid dose in the Syringe Pump.

For example:

Patient on a 25microgram/hour Fentanyl patch and receiving 30mg Morphine by a Syringe Pump over 24 hours.

To calculate breakthrough dose of subcutaneous Morphine:

For 25microgram/hour Fentanyl patch:

- Breakthrough dose from conversion charts = 10mg oral Morphine.
- o Divide by 2 to calculate subcutaneous dose = 5mg subcutaneous Morphine

For 30mg Morphine in Syringe Pump:

- Breakthrough dose = 1/6 of total dose in Syringe Pump = Total dose = 30mg
- Divided by 6 = 5mg subcutaneous Morphine

Total dose for subcutaneous breakthrough Morphine

- = 5mg + 5mg
- = 10mg subcutaneous Morphine as required

Syringe Pump Prescribing and Medicines Information (Alphabetical)

All information provided on medicines in this document is for guidance purposes only. The prescriber is responsible for prescribing in line with the most up to date guidance. Please ensure that you have read sections 4.4 and 4.5 of the policy.

Prescribing a Syringe Pump

In community:

- An FP10 prescription needs to be issued for all medication prescribed.
- An Ambulatory Syringe Pump prescription form with all required medicines must be completed.
- Each medicine, dose, diluent and final volume must be clearly written on the prescription chart by the prescriber and signed.
- Doses of drugs MUST be written in words and figures.

In hospital:

- A Syringe Pump must be prescribed on the patient's electronic patient record. There
 is a quick reference guide that explains how to do this.
- When a patient is discharged from hospital with a syringe pump an Ambulatory Syringe prescription form with all required medicines must be completed.
 - Each medicine, dose, diluent and final volume must be clearly written on the prescription chart by the prescriber and signed.
 - o Doses of drugs **MUST** be written in words and figures.

In community and hospital:

- All medicines should be mixed with sterile water for injection unless known incompatibility or otherwise stated in drug monographs below. The final volume includes all prescribed medicines and diluent.
- The prescriber must complete the prescription in full and must indicate the time that the Syringe Pump needs to be commenced.
- If medicines are changed for any reason the previous prescription and authorisation must be discontinued by the prescriber and a new one written.
- When the patient's prescribed medicines are changed the changes should be commenced on the same day.
- The Saf-T giving set and the site should be changed at the same time.
- The Saf-T giving set and site should be changed after 7 days in one position.

- No more than 3 medicines are to be used in a single Syringe Pump.
- If more than 3 medicines are required consider the use of a second Syringe Pump.
- In exceptional circumstances, if more than 3 medicines are required, advice MUST be sought from the Specialist Palliative Care Team or Pharmacy.
- Medicine combinations should be reviewed on a regular basis to check efficacy and appropriateness of medicine and dose prescribed.
- The Syringe Pump giving set must not be used to give bolus doses of medication.
- If the total volume of medicines in a 20mL syringe exceeds 17mL it can be transferred into 30mL syringe and made up to 22mL. If the total volume exceeds 22mL in a 30mL syringe, discard medication and seek advice from Specialist Palliative Care or Pharmacy.
- Information on available medicine preparations is included on the reverse of the Ambulatory Syringe Pump Prescription for information to help with volume calculations.

Mixing of medicines in a Syringe Pump and compatibility

In palliative care the administration of medication by continuous subcutaneous infusion using a Syringe Pump is common. Situations routinely arise that require combinations of two or more medicines in the same syringe, however evidence for this practice is lacking. Most combinations used in palliative care are clear, colourless and free from precipitation. However, this does not confer stability because unrecognised chemical reactions may occur. For example, Dexamethasone and Glycopyrronium mix to form a clear, colourless solution that is free from precipitation. However, at the molecular level, the Dexamethasone is reacting with and therefore deactivating the Glycopyrronium.

Compatibility of medicines in a Syringe Pump

Prescribers must ensure that any drug combinations prescribed are recognised to be compatible. Information is included on 2 drug compatibility in the drug monographs that follow.

Information on 3 drug combinations for those drugs used commonly in the last days of life: Morphine, Oxycodone, Alfentanil, Glycopyrronium, Levomepromazine and Midazolam is included below.

Further information on drug combinations not included in the information below, including other 3 drug compatibilities is available from:

- The Specialist Palliative Care Team, hospital based Pharmacists or Aseptics Unit (see pages 67 for contact details).
- The Syringe Driver Dickman et al. 4th edition 2016, Oxford University Press medicinescomplete.comdrug compatibility checker only available to subscribers

Compatibilities must be checked for all drug combinations. If unfamiliar combinations or doses are involved advice MUST be sought from a hospital based pharmacist or the Specialist Palliative Care Team.

3 drug compatibility information for medicines commonly used in the last hours to days of life

Only the medicines recommended for first line use in the last hours to days of life in the Clinical Practice Summary Guidance are considered below.

These include: Morphine (or Oxycodone/Alfentanil if Morphine not suitable/tolerated), Midazolam, Levomepromazine, Glycopyrronium.

Morphine Sulfate combinations

Drug 1	Drug 2	Drug 3	Compatibility in water for injection
Morphine Sulfate	Midazolam	Levomepromazine	Compatible
Morphine Sulfate	Midazolam	Glycopyrronium	Compatible
Morphine Sulfate	Levomepromazine	Glycopyrronium	Limited data, watch for crystallisation*

Oxycodone hydrochloride (10mg/mL) combinations

Drug 1	Drug 2	Drug 3	Compatibility in water for injection
Oxycodone	Midazolam	Levomepromazine	Compatible
Oxycodone	Midazolam	Glycopyrronium	Compatible
Oxycodone	Levomepromazine	Glycopyrronium	Limited data, watch for crystallisation*

Alfentanil

Drug 1	Drug 2	Drug 3	Compatibility in water for injection
Alfentanil	Midazolam	Levomepromazine	Compatible
Alfentanil	Midazolam	Glycopyrronium	Compatible (extrapolated from 4 drug data)
Alfentanil	Levomepromazine	Glycopyrronium	Compatible if mixed with Sodium Chloride

Combination with no opioid

Drug 1	Drug 2	Drug 3	Compatibility in water for injection
Midazolam	Levomepromazine	Glycopyrronium	Compatible

*If crystallisation occurs:

- Stop infusion
- Seek immediate advice from Specialist Palliative Care Team, Pharmacy or 24 hour Advice Line (pages 67)
- Contact Specialist Palliative Care Team to alert them to allow reporting and recording of the incompatibility

Resources assessed to update the drug monographs:

- 1) PCF online, June 2021, https://www.medicinescomplete.com/#/.
- 2) The Syringe Driver, Andrew Dickman and Jennifer Schneider, 4th Edition.
- 3) BNF online, June 2021.

Alfentanil

Usual dose: Starting dose will depend on previous opioid requirements (see

table below) and patient factors, for example renal function -

seek specialist advice

There is no maximum dose of Alfentanil providing it is carefully

titrated

BE AWARE: It is 30 times more potent than oral Morphine

Alfentanil must only be prescribed after discussion with a Consultant in Palliative Medicine or a specialist palliative care clinical nurse specialist with appropriate experience or, if unavailable out of hours, then with a Senior Hospice Physician.

Special instructions

Dilute with water for injection. Sodium Chloride 0.9% may also be used. Use an alternative opioid for breakthrough analgesia.

Indications for Use

• Pain in patients with renal failure or intolerable side effects from other opioids. It is an alternative to other opioids such a Morphine, Oxycodone and Diamorphine, particularly in patients with renal failure. It should only be used as a continuous subcutaneous infusion.

Mechanism of Action

Alfentanil is a strong opioid analgesic. It is a synthetic opioid with strong activity at mu opioid receptors. It has a rapid onset of action and a short duration of action.

Side effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Side effects are as for other strong opioids; nausea and vomiting, constipation, drowsiness, cognitive impairment, myoclonus, pruritus and many others (see BNF).

Opioid withdrawal symptoms can occur when switching from Morphine to Alfentanil, prn doses of the original opioid should help relieve symptoms.

Caution

- Renal impairment dose adjustments not usually required.
- Hepatic impairment a dose reduction may be needed seek further advice.
- Accumulation can occur in elderly or obese patients.
- Drug interactions: Alfentanil is metabolised by CYP3A4 AND CYP3A5.
- The effect of Alfentanil may be increased by drugs including aprepitant (short term effect) azoles (e.g., Fluconazole, Voricanozole), Bicalutamide, Cimetidine, Diltiazem, Haloperidol, Macrolide antibiotics (e.g., Clarithromycin, Erythromycin) protease inhibitors (e.g., Indinavir, Nelfinavir, Ritonavir).
- The effect of Alfentanil may be reduced by discontinuing aprepitant (short term effect) Carbamazepine, high dose Dexamethasone, Efavirenz, Phenobarbital, Phenytoin, Rifampicin.
- Do not administer concurrently with MAOI's or within 2 weeks of their use.

<u>Compatibilities:</u> There is 2-drug compatibility data for Clonazepam, Glycopyrronium, Haloperidol, Hyoscine Butylbromide, Levomepromazine, Metoclopramide, Midazolam, Octreotide and Ondansetron.

<u>Incompatibilities:</u> Possible concentration dependent incompatibility with Cyclizine, no data on Hyoscine Hydrobromide.

Preparations

Alfentanil 1mg/25mL, 5mg/10mL, 5mg/mL ampoules 5mg/ml strength may have restricted availability to reduce risk of errors being made.

Dose conversion between s/c Morphine, s/c Oxycodone and Alfentanil

Morphine syringe pump s/c in 24 hours	4 hourly x/c morphine*	Oxycodone syringe pump s/c in 24 hours	4 hourly s/c oxycodone*	Alfentanil syringe pump s/c in 24 hours
10mg	2.5mg	5mg	1.25mg	500micrograms
20mg	2.5mg - 5mg	10mg	2.5mg	1mg
30mg	5mg	15mg	2.5-5mg	2mg
60mg	10mg	30mg	5mg	3mg
90mg	15mg	45mg	7.5mg	4mg
120mg	20mg	60mg	10mg	6mg
150mg	25mg	75mg	15mg	8mg
200mg	30mg	100mg	15mg	10mg
240mg	40mg	120mg	20mg	12mg

*Please note: The most common reason for a patient requiring Alfentanil is severe renal impairment. In this instance the dose and frequency of administration of prn/breakthrough opioids may need to be adjusted to reduce the risk of accumulation and side effects. Seek further advice if needed.

Clonazepam

Usual dose: 500micrograms to 4mg over 24 hrs

Clonazepam subcutaneously is only available as an unlicensed preparation. Patients, or if relevant, a relative with lasting power of attorney, should be informed of this and given an information leaflet. In cases where the patient is not able to consent and there is no relative with power of attorney, then it is appropriate for the Clinician to make the decision and take responsibility to treat the patient without informing them of the unlicensed status of the medicine or providing the leaflet.

The relevant information leaflet can be found on the ELHT hospital intranet or can be accessed by contacting pharmacy.

Special Instructions

Dilute with Water for injections.

Sodium Chloride 0.9% may also be used (the 1mg/mL injection of Clonazepam must be diluted with the supplied WFI prior to parenteral administration: Sodium Chloride 0.9% can be used to further dilute a Syringe Pump).

Indications for Use

- Terminal restlessness
- Neuropathic pain

Clonazepam is an alternative to Midazolam, but it is recommended to be reserved for the treatment of terminal restlessness associated with a previous history of neuropathic pain. Although there are no randomised trials supporting the use of Clonazepam in neuropathic pain, it is an accepted treatment in several centres.

Mechanism of Action

Clonazepam is a long-acting Benzodiazepine.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

- Dose-dependent adverse effects commonly reported include dizziness, drowsiness, fatigue and muscle weakness.
- Increased risk of falls and fracture in the elderly with Benzodiazepines.

Caution

- Use with caution in patients with chronic respiratory disease, renal impairment and hepatic impairment if possible, avoid in severe hepatic impairment and myasthenia gravis.
- Clonazepam is metabolised by CYP3A4 and is susceptible to drug interactions:
 - Clonazepam effect may be reduced by co-administration of enzyme inducers, such as Carbamazepine, Phenobarbitone and high dose Dexamethasone.
 - Clonazepam effect may be enhanced by co-administration of enzyme inhibitors, such as Bicalutamide, Erythromycin, high-dose Fluconazole and Haloperidol.

- Effect on Clonazepam metabolism may persist for several days after cessation of these drugs.
- Use non-PVC giving sets as absorption into PVC infusion sets may occur.

Compatibilities

There are 2-drug compatibility data for Clonazepam in water for injection with Alfentanil, Diamorphine, Glycopyrronium, Haloperidol, Hyoscine Butylbromide, Hyoscine Hydrobromide, Morphine Sulphate and Oxycodone.

Preparations

Clonazepam 1mg/1mL in solvent ampoule, with 1mL Water for injection ampoule.

Cyclizine

Usual dose: 75 - 150mg over 24hrs

Special Instructions

Dilute with Water for injection.

Cyclizine is incompatible with 0.9% Sodium Chloride and will precipitate.

Indications for Use

- Nausea and vomiting due to:
 - o Raised intracranial pressure (in conjunction with dexamethasone).
 - Bowel obstruction.

Vertigo due to vestibular cause (should be avoided long-term as they may inhibit compensatory mechanisms).

Mechanism of Action

Cyclizine has antihistamine and antimuscarinic activity.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects

Anti-muscarinic effects:

- CNS drowsiness, cognitive impairment, delirium, restlessness, agitation.
- Visual mydriasis, loss of accommodation causing blurred vision.
- Cardiovascular tachycardia, palpitations, arrhythmias.
- Gastro-intestinal dry mouth, heartburn (due to relaxation of lower oesophageal sphincter), constipation.
- Urinary tract hesitancy of micturition, retention of urine.
- The elderly are more susceptible to sedative and antimuscarinic effects.
- Rarely movement disorders can occur e.g., tremor, dyskinesia and dystonia.

May occasionally cause irritation at the injection site.

Contra-indications

Narrow-angle glaucoma, acute porphyria.

Caution

Severe heart failure, acute myocardial infarction, glaucoma, epilepsy, myasthenia gravis, prostatic hypertrophy, urinary retention renal or hepatic impairment.

Anticholinergic medicines can directly interfere with the prokinetic action of Metoclopramide.

The combination of Cyclizine and Metoclopramide should be avoided.

Compatibilities

Cyclizine is implicated in many compatibility problems. To reduce the precipitation risk, dilute Cyclizine with Water for Injections before mixing.

There are 2-drug compatibility data for Cyclizine in water for injection with Haloperidol, Hyoscine Hydrobromide and Morphine Sulfate.

Incompatibilities

Concentration-dependent incompatibility occurs with Alfentanil, Dexamethasone, Diamorphine and Oxycodone.

Incompatibility has been reported with Clonazepam, Hyoscine Butylbromide, Ketorolac, Midazolam, and Octreotide.

Dexamethasone

Usual dose: 4mg to 16mg over 24hrs (3.3 to 13.2mg base over 24 hours)

Special Instructions

Dexamethasone should be diluted with Water for Injections. Also compatible with Sodium Chloride 0.9% as diluent if necessary.

If Dexamethasone is effective, consider reducing the dose after 5-7 days.

Indications for Use

- Nausea and vomiting (especially due to intestinal obstruction, raised intracranial pressure or associated with chemotherapy or radiotherapy). Dexamethasone is usually used in addition to other antiemetic for nausea and vomiting rather than used alone.
- For obstructive symptoms e.g. bowel obstruction, upper airway obstruction causing dyspnoea, superior vena cava obstruction.
- For symptoms relating to increased intracranial pressure (usually due to brain tumour/metastases).
- Cerebral oedema secondary to brain metastases.
- For patients established on long term Dexamethasone who need to continue this for symptomatic reasons.
- Pain (particularly if caused by nerve compression, liver capsule pain or bone pain).
- Metastatic spinal cord compression.

Consider prescribing

Consider prescribing gastro protective medicines if the patient is able to take oral medication (e.g.,, oral Omeprazole 20mg od) especially if high doses of steroids are being used, if an NSAID is co-prescribed or if there are other risk factors for gastric irritation (e.g. age, multiple recent courses of steroids, previous GI bleed).

Mechanism of Action

The main benefits of steroids in palliative care are due to their anti-inflammatory effects which are mediated by several different mechanisms. These anti-inflammatory effects can reduce peri-tumour oedema and so relieve compression and associated symptoms. There may also be other direct antiemetic effects. These various mechanisms make steroids very effective drugs, but also cause wideranging side effects, especially if used long-term.

Compared with other steroid drugs, Dexamethasone has high anti-inflammatory effects (due to being a highly potent glucocorticoid), with less fluid retention (due to having negligible mineralocorticoid effects). It is especially useful when high doses of steroids are needed and has a long duration of action, so once daily dosing is effective.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Steroids have wide-ranging side effects, particularly if used long-term.

Common short-term side effects include:

- Insomnia prescribe steroids to be given before 2pm to minimise this.
- Psychiatric effects including restlessness, depression, mania, psychosis and delirium.
- Peptic ulceration, especially if given with an NSAID.
- Hyperglycaemia (especially in the evening) and deterioration in diabetic control.
- Adrenal suppression with multiple or prolonged courses.
- Susceptibility to infection including candida infection.

- Cushingoid features skin changes, susceptibility to bruising, proximal myopathy, truncal obesity.
- Osteopenia/osteoporosis (long term use/multiple courses).

Contra-indications and cautions

In general, if the patient is in the last days or weeks of life, there are no absolute contra-indications. Potential benefits need to be weighed up against risks.

Caution in diabetes mellitus (risk of hyperglycaemia), psychotic illness (symptoms can emerge within a few days of starting steroids), other risk factors for gastric irritation (e.g. concurrent NSAID administration), hypokalaemia, can increase susceptibility to serious infections and mask their symptoms, prescribe cautiously if recent surgery.

Usually, contra-indicated if systemic infection is present (unless treatment not possible or appropriate and benefit of steroid outweighs risk).

Interactions

Refer to the BNF and the manufacturer's SPC for a detailed list of interactions.

Steroids reduce effect of insulin, oral hypoglycaemics, anti-hypertensives and diuretics.

Steroids can increase the INR of patients on Warfarin.

CYP3A4 is involved in metabolising Dexamethasone, so it is susceptible to drug interactions:

- Effect of Dexamethasone can be increased by enzyme inhibitors e.g. Itraconazole, Bicalutamide, Erythromycin
- Effect of Dexamethasone cab be reduced by enzyme inhibitors e.g. Carbamazepine, Phenobarbital, Phenytoin. Larger doses of Dexamethasone may be needed.
- Increases INR in patients taking Vitamin K antagonist (e.g., Warfarin) increase frequency of INR monitoring.

Compatibilities

Dexamethasone often causes compatibility problems therefore, if it is to be mixed with other medicines, as much diluent as possible should be added *before* the addition of Dexamethasone. Dexamethasone should be the last constituent added. If precipitate remains in the mixture, it is incompatible.

Some centres always use a separate Syringe Pump for Dexamethasone due to its liability to precipitate with other medicines.

Consider a once or twice daily dose, no later than 2pm, rather than syringe pump administration if possible.

There is 2-drug compatibility data for Dexamethasone in water for injection with Morphine, Oxycodone, Ketamine, Ranitidine, Hyoscine Butylbromide and Metoclopramide.

Incompatibilities

Glycopyrronium may be inactivated by Dexamethasone, but no precipitate forms. Therefore *avoid* combination.

Incompatibility has been reported with Midazolam.

There is concentration-dependent incompatibility with Ondansetron, Haloperidol, Levomepromazine and Cyclizine. Use these combinations with caution and seek advice from the Specialist Palliative Care Team or Pharmacy.

Preparations

Dexamethasone 3.3mg/mL vials (as 4.3mg Dexamethasone Sodium Phosphate). This is equivalent to 4mg of oral Dexamethasone.

Also available as 6.6mg/2mL vials and 3.8mg/mL vials.

Diamorphine

Usual dose:

Starting dose of Diamorphine dependent of the patient's present opioid requirements (see table below) and patient factors, for example renal function – seek specialist advice if needed.

There is no maximum dose of Diamorphine providing it is carefully titrated

Special instructions

Dilute with Water for Injections. Concentration dependent incompatibility can occur with 0.9% Sodium Chloride at higher doses or in combination with certain other drugs.

Usual dosage

A suitable starting dose for an opioid naïve patient would be 5mg to 10mg Diamorphine over 24 hours. For patients with uncontrolled opioid-responsive pain, who are tolerating Diamorphine, the Diamorphine should be increased by 30-50%. Rescue doses for breakthrough pain should be prescribed and are calculated to be one sixth of the total daily dose.

Although there is no maximum dose of Diamorphine, dosing should be titrated based upon a balance of analgesic effect versus undesirable effects.

Indications for Use

- Morphine is a similar drug and is usually used rather than Diamorphine. Diamorphine is useful
 if large doses of Morphine are needed, as it has better solubility and needs less volume for
 administration.
- Pain control
- Breathlessness

Mechanism of Action

Diamorphine is a derivative of Morphine. When given by subcutaneous injection it is rapidly absorbed and converted to the active metabolite, 6-monoacetylmorphine. This is then slowly converted to Morphine.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Opioids tend to cause similar side effects.

Common side effects include nausea and vomiting, constipation, drowsiness, cognitive impairment, myoclonus, pruritus.

Additional adverse effects that can develop with deteriorating renal function and accumulation of metabolites include delirium, hallucinations, myoclonic jerks and nightmares. Consider switching opiates, e.g. to Alfentanil or Oxycodone if occurs. Seek specialist advice.

For all patients prescribed regular Diamorphine, consider also prescribing a regular laxative (if able to take) and an antiemetic, regularly/prn.

Caution

The same considerations as for Morphine apply to the use of Diamorphine in patients with renal or hepatic impairment, i.e. metabolites are likely to accumulate and so alternatives may be needed. Seek specialist advice.

Compatibilities

There is 2-drug compatibility data for Diamorphine in water for injection with Clonazepam, Dexamethasone, Glycopyrronium, Hyoscine Butylbromide, Hyoscine Hydrobromide, Ketorolac, Levomepromazine, Metoclopramide, Midazolam, Octreotide and Ondansetron.

Incompatibilities

Concentration-dependent incompatibility occurs with Cyclizine and Haloperidol at higher concentrations.

Preparations

Diamorphine 5mg, 10mg, 30mg, 100mg and 500mg ampoules containing powder.

Dose conversion between s/c Morphine and Diamorphine

Morphine syringe pump s/c in 24 hours	PRN s/c Morphine	Diamorphine syringe pump s/c in 24 hours	PRN s/c diamorphine
7.5mg	1.25mg	5mg	1mg
15mg	2.5mg	10mg	2mg
30mg	5mg	20mg	3.5mg
45mg	7.5mg	30mg	5mg
60mg	10mg	40mg	7.5mg
90mg	15mg	60mg	10mg
120mg	20mg	80mg	12.5mg
135mg	20-25mg	90mg	15mg
150mg	25mg	100mg	17.5mg
180mg	30mg	120mg	20mg

Furosemide

Usual dose:	20-140mg via Syringe Pump over 24 hours
	If converting from oral to subcutaneous Furosemide because of fluid overload, consider using a 1:1 conversion ratio (this will be an increase in dose). If converting because a patient is not able to take oral Furosemide in the last days of life, consider a 2:1 conversion ratio (a dose reduction).

Furosemide via Syringe Pump must only be prescribed after discussion with a Consultant in Palliative Medicine or if a Consultant is unavailable out of hours, then with a Senior Hospice Physician.

Special Instructions

Dilute with 0.9% Sodium Chloride. Water for injection could also be used.

Furosemide is only available in 10mg/mL solution for injection. This volume means that SC doses are usually not practical, and a Syringe Pump needs to be used.

Indications for Use

If the patient is unable to swallow, consider a Syringe Pump containing Furosemide if needed for control of symptoms due to peripheral or pulmonary oedema or ascites. Titrate the dose until symptoms are managed.

Occasionally, Syringe Pump Furosemide may be used to manage decompensated congestive heart failure if intravenous therapy and/or hospital admission is not appropriate. Seek Specialist Palliative Care advice.

Mechanism of Action

Furosemide is a loop diuretic. It reduces the resorption of Sodium and therefore water within the kidney. It also increases urinary excretion of Potassium, Magnesium, Hydrogen and Chloride.

Side Effects

Refer to the manufacturer's SPC for a detailed list of adverse effects.

- Subcutaneous administration can cause pain and itching at the site of injection.
- Symptoms related to dehydration or electrolyte abnormalities (hypokalaemia, hypomagnesaemia) e.g. thirst, dizziness, weakness, muscle cramps.

Cautions

Refer to the manufacturer's SPC for detailed list of contraindications and precautions. Usually contra-indicated in patients with:

- Hepatic encephalopathy.
- Anuric renal failure.
- Renal failure due to nephrotoxic or hepatotoxic drugs.
- Dehydration/hypovolaemia.
- Hypersensitivity to sulphonamides.
- Severe hypokalaemia or hyponatraemia.

For patients in the last days and weeks of life, the prescriber must consider these conditions, but they may not necessarily be a deterrent to use, provided the dose is carefully titrated.

Use with caution in patients with:

- Prostatic hypertrophy.
- Diabetes mellitus (glucose levels may increase).
- Hepatic impairment monitor treatment closely and carefully titrate the dose to effect.
- Renal impairment monitor treatment closely and carefully titrate the dose to effect.
- Other medications affecting QT interval e.g. Citalopram, Methadone electrolyte disturbances caused by Furosemide increase the risk of cardiac effects from these medications.

If appropriate, ensure regular blood tests are performed to monitor electrolytes.

Compatibilities

Furosemide injection is alkaline and there is a high risk of incompatibility when mixed with acidic drugs. Because of this and the lack of compatibility date, Furosemide should not be mixed in the same syringe with any other drugs.

Preparations

10mg/mL solution for injection in 2mL, 4mL, 5mL, 25mL.

Glycopyrronium

Usual dose:	600 micrograms to 1200micrograms over 24 hours

Special Instructions

Glycopyrronium should be diluted with Water for Injections.

Can be diluted with Sodium Chloride 0.9%.

Indications for Use

- Excessive respiratory secretions. However, the development of terminal secretions must be anticipated because Glycopyrronium will not clear existing secretions.
- Bowel colic.
- May be of benefit in the treatment of large volume vomiting associated with bowel obstruction, possibly in combination with Octreotide.

Mechanism of Action

Glycopyrronium is a powerful anticholinergic causing inhibition of the parasympathetic autonomic system. It does not cross the blood brain barrier so is devoid of CNS effects such as paradoxical agitation and has less of an effect on the ocular and cardiovascular systems, at normal doses, than Hyoscine Hydrobromide.

Side Effects

Refer to the manufacturer's SPC for a detailed list of adverse effect.

The side effects of Glycopyrronium are dose related and are associated with its pharmacology. They include dry mouth, constipation and urinary retention.

Glycopyrronium may precipitate tachycardia.

Caution

The effect of Glycopyrronium accumulates in renal impairment and so dose adjustments may be necessary – check with medicines information for dosing.

Glycopyrronium should be avoided in patients with closed-angle glaucoma or paralytic ileus.

However, this is not a contraindication for patients with advanced disease.

Glycopyrronium may antagonise the prokinetic effects of Metoclopramide.

Compatibilities

There is 2-drug compatibility data for Glycopyrronium in water for injection with Alfentanil, Clonazepam, Diamorphine, Haloperidol, Levomepromazine, Metoclopramide, Midazolam, Morphine Sulfate, Oxycodone.

Limited 3 drug compatibility information can be found on pages 18-19.

Incompatibilities

Dexamethasone and Ketorolac. There may be a concentration-dependent incompatibility with Cyclizine.

Preparations

Glycopyrronium 200microgram/1mL ampoules Glycopyrronium 600microgram/3mL ampoules

Haloperidol

Usual dose: 500micrograms to 5mg over 24hrs (antiemetic)

500micrograms to 10mg over 24 hrs (agitation)

Special Instructions

Dilute with Water for Injections.

Can be diluted with Sodium Chloride 0.9%.

Indications for Use

- Nausea and vomiting due to chemical causes i.e. medicines, biochemical disturbance. Sedation is minimal at the low doses used for nausea and vomiting.
- Agitation and Confusion. Higher doses are sedating and it can be used for agitation and confusion. However higher doses may produce extrapyramidal side effects therefore Levomepromazine should be used if sedation is required.
- Hiccups.

Mechanism of Action

Haloperidol is a central dopamine D2 receptor antagonist with sedating properties.

When prescribing Haloperidol the subcutaneous dose should be lower than the corresponding oral dose (which undergoes significant first-pass metabolism).

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Extrapyramidal symptoms, especially if combined with other D2 antagonists e.g., Metoclopramide, Levomepromazine.

Anticholinergic effects such as drowsiness/apathy, dry mouth, constipation, difficulty with micturition.

Cautions

Refer to manufacturer's SPC for a detailed list of contraindications and precautions.

Use with caution with concurrent use of CYP2D6 and/or CYP3A4 inhibitors/inducers (e.g., Carbamazepine, Phenobarbital, Phenytoin, Rifampicin, Itraconazole, Fluoxetine).

Exacerbates Parkinson's disease so use alternatives where possible.

No specific guidance available in hepatic impairment, however, since Haloperidol undergoes extensive first pass metabolism, the lowest effective dose should be used in hepatic impairment.

The active metabolites of Haloperidol may accumulate in renal failure – check with medicines information for dosing.

Compatibilities

There is 2-drug compatibility data for Haloperidol in water for injection with Alfentanil, Clonazepam, Cyclizine, Glycopyrronium, Hyoscine Butylbromide, Hyoscine Hydrobromide, Metoclopramide, Midazolam and Oxycodone.

Incompatibilities

Incompatible with Dexamethasone and Ketorolac.

Concentration dependent incompatibility occurs with Morphine Sulfate and Diamorphine.

Preparations

Haloperidol 5mg/mL ampoules.

Hyoscine Butylbromide (Buscopan)

Usual dose: 60mg to 120mg over 24 hrs

Special Instructions

Dilute with Water for Injections. Sodium Chloride 0.9% may also be used.

Indications for Use

- Intestinal colic associated with bowel obstruction.
- Large volume vomiting associated with bowel obstruction (by reducing gastrointestinal secretions). Note: the maximum benefit may be seen only after three days.
- Spasm of the genito-urinary tract.
- Respiratory tract secretions.

Mechanism of Action

Hyoscine Butylbromide is an antimuscarinic.

Does not readily cross the blood brain barrier and so unlikely to cause sedation.

50% of the drug is excreted renally, unchanged.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Anticholinergic effects including dry mouth, constipation, urinary retention, tachycardia, palpitations, heartburn, mydriasis.

Cautions

Avoid in closed angle glaucoma and paralytic ileus unless patient has advanced disease.

Use with caution in patients with congestive cardiac failure or those with renal impairment. (Use the lowest effective dose).

The anticholinergic effects of Hyoscine Butylbromide can be additive with other drugs and may precipitate delirium or cognitive impairment in susceptible patients.

May antagonise the prokinetic effects of Metoclopramide.

Compatibilities

There is 2-drug compatibility data for Hyoscine Butylbromide in water for injection with Alfentanil, Haloperidol, Levomepromazine, Midazolam, Morphine Sulfate and Oxycodone.

Incompatibilities

Not compatible with Cyclizine.

Preparations

Hyoscine Butylbromide 20mg/1mL ampoules.

Hyoscine Hydrobromide

Usual dose: 400micrograms to 2400micrograms over 24 hrs

Special Instructions

To be diluted with Water for Injections. Sodium Chloride 0.9% may also be used.

Indications for Use

• Excessive respiratory tract secretions. However, the development of terminal secretions must be anticipated because Hyoscine Hydrobromide will not clear existing secretions.

Mechanism of Action

Hyoscine Hydrobromide is an antimuscarinic drug.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Dry mouth, constipation, urinary retention, blurred vision, drowsiness, paradoxical agitation. It crosses the blood brain barrier and as such can result in possible sedation and delirium.

Cautions

Bradycardia.

Tachycardia.

Congestive cardiac failure.

Hepatic impairment.

Myasthenia gravis.

Paralytic ileus.

Renal impairment.

Increased risk of seizures in epileptic patients.

Avoid in closed angle glaucoma and paralytic ileus. However this is not a contra-indication with advanced disease.

May block the prokinetic effects of Metoclopramide.

Compatibilities

There is 2-drug compatibility data for Hyoscine Hydrobromide in water for injection with Cyclizine, Diamorphine, Haloperidol, Levomepromazine, Midazolam, Morphine Sulfate and Oxycodone.

Preparations

Hyoscine Hydrobromide 400microgram/1mL ampoules.

Ketamine

Usual dose: Starting Dose: 50-100mg/over 24 hours

Increase by 50-100mg/24 hours until benefit achieved

Usual maximum 500mg/24 hours

To be used only on the recommendation of a Palliative Care Specialist for patients who have failed to obtain adequate relief from standard non-drug and drug treatments.

Special Instructions

As Ketamine is irritant it must be diluted with Sodium Chloride 0.9% w/v to the largest volume possible.

Ketamine can have an opioid sparing effect. If used concurrently with an opioid, consider a dose reduction of the opioid prior to initiating the Ketamine.

Indications for Use

• Pain unresponsive to standard analgesic treatments (including neuropathic, inflammatory, ischaemic limb and cancer related bone pain).

Concurrent Medicines

Consider the use of:

- Haloperidol (e.g. 2mg to 5mg/24hours) or Midazolam (e.g. 5-10mg/24hours) to treat dysphoria or hallucinations.
- Dexamethasone (500micrograms to 1mg/24hours) to reduce site toxicity.

Mechanism of Action

It is believed to produce an analgesic effect through antagonism of the N-methyl-D-aspartate (NMDA) receptor.

Side Effects

Refer to manufacturer's SPC for a detailed list.

The main side effects are:

- Psychotomimetic phenomena (including hallucinations, dysphoria and vivid dreams).
- Tachycardia, hypertension.
- Erythema and pain at the injection site.
- Urinary tract toxicity. Prescribers should consider discontinuing the Ketamine and seeking urology advice if patient develops urinary tract symptoms with no evidence of bacterial infection.

Cautions

Refer to manufacturer's SPC for a detailed list.

These include:

Cardiac failure, hypertension, cerebrovascular disease, ischaemic heart disease, raised intra-ocular pressure (glaucoma), epilepsy, current or past history of a psychiatric disorder. Concurrent opioid use my lead to opioid toxicity.

Contra-indications

Refer to manufacturer's SPC for a detailed list.

These include:

Raised intracranial pressure, severe cardiac disease, cerebrovascular accident or cerebral trauma or where a rise in blood pressure would pose a serious hazard.

Compatibilities

There is 2-drug compatibility data for: Haloperidol, Midazolam, low dose Dexamethasone, Diamorphine, Levomepromazine, Metoclopramide, Morphine Sulfate, Alfentanil, Clonazepam and Oxycodone.

Use a separate Syringe Pump from the opioids unless advised by a Palliative Medicine Consultant.

Preparations

Ketamine 10mg/mL – 20mL vial Ketamine 50mg/mL – 10mL vial

Ketorolac

Starting dose: Usually 60mg over 24hours

Increase by 15mg/24hours if necessary to

90mg/24 hours

(60mg/24hours is the recommended maximum dose in

those >65 years and/or <50kg)

To be used only on the recommendation of a Palliative Care Specialist

Special Instructions

Dilute maximally with Sodium Chloride 0.9% (preferred diluent) to avoid irritation at the infusion site. (Also compatible with water for injections).

Concurrent opioid dose reduction should be considered and other NSAID's (if any) must be discontinued.

Indications for Use

• Short term management of cancer pain when the use of other NSAID's has been exhausted or is impractical. Use the minimum effective dose for the shortest duration necessary in order to reduce risk of serious and undesirable side effects.

Concurrent medicines

High potential to cause upper gastrointestinal bleeds/perforation. The concurrent use of a gastro-protective drug e.g. a proton pump inhibitor must be considered to minimise this risk to the patient.

Mechanism of Action

NSAID with anti-inflammatory, analgesic and antipyretic activity.

Side Effects

Refer to manufacturer's SPC for a detailed list.

Gastro-intestinal tract (ulceration, haemorrhage, perforation) and renal function (hypercalcaemia, uraemia, acute renal failure). Anaphylaxis, drowsiness, dizziness, headache, thrombocytopenia, skin reactions.

Of all NSAID's Ketorolac has the highest risk for gastritis, duodenitis and upper gastrointestinal complications.

Cautions

Refer to manufacturer's SPC for a detailed list.

Hypovolaemia from any cause (including those taking diuretics or the elderly). History of cardiac failure left ventricular dysfunction or hypertension. Renal, cardiac or hepatic impairment. Risk of bleeding increased if co-prescribed with antiplatelet drugs, corticosteroids or SSRI's.

Contra-indications

Refer to manufacturer's SPC for a detailed list

History of hypersensitivity to Aspirin or NSAID, Asthma, active peptic ulceration (or history of gastro-intestinal bleeding, ulceration or perforation). Renal impairment, dehydration, pregnancy, severe heart failure, severe hepatic impairment, coagulation/bleeding disorders or cerebrovascular bleeds. Concurrent treatment with Aspirin, NSAIDS, probenecid, Lithium salts or anticoagulants.

Compatibilities

2-drug compatibility data is available in Sodium Chloride 0.9% w/v for Diamorphine (dependent upon concentration), Oxycodone and Ranitidine.

Incompatibilities

May precipitate in solutions with low pH e.g. Midazolam, Haloperidol and Cyclizine.

<u>Preparations</u> Ketorolac 30mg/mL 1mL ampoules.

Levetiracetam

As per oral dosing - will depend on previous oral

requirements

Usual dose: Use 1:1 oral subcutaneous conversion ratio

Max 2g/24hours in one syringe due to volume constraints

Levetiracetam must only be prescribed as a subcutaneous infusion after discussion with a Consultant in Palliative Medicine or, if a Consultant is unavailable out of hours, then with a Senior Hospice Physician.

Special Instructions

Dilute maximally with Water for Injection (i.e. to volume of 22mLs). Sodium Chloride 0.9% w/v may also be used.

Administer in a separate Syringe Pump.

Higher doses may require 2 Syringe Pumps.

Indications for Use

Levetiracetam is an antiepileptic which can be used to treat focal or generalised seizures. Its
use subcutaneously (unlicensed) offers the possibility of maintaining seizure control when oral
or intravenous routes of Levetiracetam administration are not possible and when increased
sedation (from alternatives such as Midazolam) is undesirable.

Mechanism of Action

Levetiracetam binds to a synaptic vesicle protein and is presumed to interfere with the release of the neurotransmitter stored within the vesicle. It readily crosses the blood-brain barrier. It is effective for a broad range of seizure types.

Side effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Common or very common: Abdominal pain; aggression; anorexia; anxiety; convulsion; cough; depression; diarrhoea; dizziness; drowsiness; dyspepsia; headache; insomnia; irritability; malaise; nasopharyngitis; nausea; rash; tremor; vertigo; vomiting.

Uncommon: Agitation; alopecia; amnesia; ataxia; blurred vision; confusion; diplopia; eczema; impaired attention; leucopenia; myalgia; paraesthesia; pruritus; psychosis; suicidal ideation; thrombocytopenia; weight changes.

Rare: Agranulocytosis; choreoathetosis; drug reaction with eosinophilia and systemic symptoms (DRESS); dyskinesia; erythema multiforme; hepatic failure; hyponatraemia; hypersensitivity; neutropenia; pancreatitis; pancytopenia; Stevens-Johnson syndrome; toxic epidermal necrolysis.

Caution

Renal impairment – dose needs to be reduced – seek further advice.

Hepatic impairment – dose may need to be reduced in severe hepatic impairment if associated renal impairment – seek further advice

Stopping Levetiracetam – do not stop abruptly; reduce by maximum of 500mg bd every 2-4 weeks to avoid rebound seizures.

Compatibilities

Data not available – use a separate Syringe Pump.

Preparations

Levetiracetam 100mg/mL 5mL ampoules.

Levomepromazine

Usual dose: 5mg to 25mg/24hours (antiemetic)

12.5mg to 200mg/24hours (agitation)

Special Instruction

To avoid infusion site reactions, dilute maximally with Sodium Chloride 0.9% where possible. Water for Injections can also be used.

Protect infusions from light.

Indications for Use

- Levomepromazine is used at low doses to treat intractable nausea and vomiting. It is a broad spectrum antiemetic usually used as a second or third line drug for patients who do not respond to more specific antiemetics.
- At higher doses it can be used to manage agitation/terminal restlessness due to its antipsychotic and potent sedative action.

Mechanism of Action

Levomepromazine is a phenothiazine antipsychotic drug with a half-life of 15-30 hours. It acts on the main receptor sites involved in the vomiting pathway.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Sedation, skin irritation at infusion site, dry mouth, postural hypotension, extra-pyramidal reactions. May also rarely cause prolongation of QT interval in cardiac disease and hypokalaemia.

Caution

Use with caution (i.e. low initial doses), especially in ambulatory patients with; concurrent antihypertensive medication, diabetes, epilepsy (lowers seizure threshold), liver dysfunction, Parkinson's disease or postural hypotension.

Avoid in patients with dementia unless patient at immediate risk of harm or severely distressed (increased mortality reported). Use lowest possible dose for shortest possible duration.

Irritation at infusion site may occur. For lower doses, a bolus subcutaneous injection can be given to overcome this problem (usually at night).

Compatibilities

There is 2-drug compatibility data for Morphine Sulfate, Oxycodone, Diamorphine, Glycopyrronium, Hyoscine Butylbromide, Hyoscine Hydrobromide, Ketamine, Metoclopramide, Midazolam, Octreotide and Ondansetron.

Limited 3 drug compatibility information can be found in the compatibility tables shown on pages 18-19.

Incompatibilities

Incompatibilities reported with Dexamethasone (concentration dependent), Ketorolac and Ranitidine.

Preparations

Levomepromazine 25mg/1mL ampoules.

Metoclopramide

Usual dose: 30mg to 60mg over 24 hours

There is an increased risk of neurological adverse effects at doses higher than 30mg/24hours and if used for longer than 5 days

Special Instructions

Dilute with Water for Injections or Sodium Chloride 0.9%.

Must not be used if complete intestinal obstruction is present or if there is a risk of gastrointestinal perforation or haemorrhage. Should not be used within 4 days of gastrointestinal surgery.

Indications for Use (Maximum of 5 days)

- Nausea and vomiting caused by medicines
- Gastric stasis
- Partial outflow obstruction
- Hiccups

Mechanism of Action

Metoclopramide is a central dopamine D2 receptor antagonist with non-sedating antiemetic and prokinetic properties.

Side Effects

Refer to manufacturer's SPC for detailed list of adverse effects.

Extrapyramidal reactions may occur especially if used concurrently with another D2 antagonist, particularly in children and young adults.

Drowsiness.

Diarrhoea.

Serotonin toxicity when combined with SSRI antidepressants.

Caution

Refer to manufacturer's SPC for a detailed list of contraindications and cautions.

Avoid concurrent IV Ondansetron used as this can cause cardiac arrhythmias.

Can cause irritation at the site of injection.

Anticholinergic medicines can directly interfere with the prokinetic action of Metoclopramide.

The combination of Cyclizine and Metoclopramide should be avoided.

Metoclopramide antagonises the treatment of Parkinson's disease.

Metoclopramide may increase the rate of absorption of Morphine via increased gastric emptying. Dose reduction of up to 75% may be necessary in moderate to severe renal impairment (refer to manufacturer's SPC for further details).

• Dose reduction of up to 50% may be necessary in patients with a significant degree of hepatic impairment.

Epilepsy – lowers seizure threshold.

Cardiac disease.

Compatibilities

There is 2-drug compatibility data for Metoclopramide in water for injection with Alfentanil, Diamorphine, Glycopyrronium, Haloperidol, Hydromorphone, Ketamine, Midazolam, Morphine Sulfate, Octreotide and Oxycodone. The injection should be discarded if it discolours. Sometimes compatible with Dexamethasone but may precipitate.

Incompatibilities

Metoclopramide may crystallise with Cyclizine. This combination should not be used routinely.

Preparations Metoclopramide 10mg/2mL ampoules.	

Midazolam

Usual dose: 10mg to 30mg over 24 hours

Higher doses may be used – seek Specialist Palliative

Care advice for doses above 30mg

Special Instructions

Dilute with Water for Injections or Sodium Chloride 0.9%.

Initial starting dose of Midazolam should be no more than 10mg per 24 hours. Higher doses may be used if required for seizure control. Specialist Palliative Care advice should be sought in this situation.

Indications for Use

- Terminal agitation
- Anticonvulsant
- Anxiety
- Breathlessness if patient already on regular opioids
- Myoclonus

Mechanism of Action

Midazolam is a Benzodiazepine.

Side effects

Refer to manufacturer's SPC for a detailed list of adverse effects. Drowsiness, confusion, ataxia, amnesia, cognitive impairment.

In <10% of people it can contribute to increased agitation.

Caution

Refer to manufacturer's SPC for a detailed list of contraindications and cautions.

Dose reduction may be necessary in liver disease (main site of metabolism) and renal disease (accumulation of metabolite).

Elderly.

Compatibilities

There is 2-drug compatibility data for Midazolam in water for injection with Alfentanil, Diamorphine, Glycopyrronium, Haloperidol, Hyoscine Butylbromide, Hyoscine Hydrobromide, Levomepromazine, Metoclopramide, Morphine Sulfate and Oxycodone.

Limited 3 drug compatibility information can be found on pages 18-19.

Incompatibilities

Likely to cause precipitation if mixed with Dexamethasone or Ketorolac.

Preparations

Midazolam 10mg/2mL ampoules.

Morphine

Usual dose: Starting dose of Morphine is dependent on the patient's

present opioid requirements and patient factors, for

example renal function - seek advice if needed.

There is no maximum dose of Morphine providing it is

carefully titrated

Special Instructions

Dilute with Water for Injections or Sodium Chloride 0.9%.

Usual Dosage

Morphine is the first line opioid of choice.

Although there is no maximum dose of Morphine, dosing should be titrated based upon a balance of analgesic effect versus undesirable effects.

The initial dose of Morphine is dependent on the patient's present opioid requirements. A suitable starting dose for an opioid naïve patient would be 10mg to 20mg Morphine over 24 hours. For patients with uncontrolled pain, the Morphine should be increased by 30-50%.

Additional 'as required' doses for breakthrough pain should be prescribed at a dose of one sixth of the total daily dose of the regular Morphine prescription.

Conversion ratio for oral to subcutaneous Morphine = 2:1
 i.e. 60mg oral Morphine daily = 30mg subcutaneous Morphine in 24 hours.

Indications for use

- Pain control.
- Breathlessness.
- Cough.

Mechanism of Action

Morphine is a strong opioid and acts primarily via µ-opioid receptor.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Adverse effects commonly include nausea and vomiting, constipation, drowsiness, cognitive impairment, myoclonus and pruritus.

<u>Caution</u>

Refer to manufacturer's SPC for a detailed list of contraindications and cautions.

The metabolites of Morphine are renally excreted, therefore patients with renal impairment are at risk of toxicity and may need a reduced dose of Morphine between 30-50% or an alternative drug e.g., Alfentanil. Seek specialist advice in these circumstances.

Compatibilities

There is 2-drug compatibility data for Morphine Sulfate in water for injection with Clonazepam, Cyclizine, Glycopyrronium, Hyoscine Butylbromide, Hyoscine Hydrobromide, Ketamine, Levomepromazine, Metoclopramide and Octreotide.

Incompatibilities

Morphine Sulfate is *incompatible* with Ketorolac and may be *incompatible* with higher concentrations of Haloperidol or Midazolam.

Limited 3 drug compatibility information can be found on pages 18-19.

<u>Additional medicines</u>
A regular laxative, or as required laxative and/or antiemetic may be necessary.

Preparations
Morphine 10mg/1mL ampoules.
Morphine 15mg/1mL ampoules.
Morphine 30mg/1mL ampoules.
Morphine 60mg/2mL ampoules.

Octreotide

Usual dose: 200micrograms to 600micrograms over 24 hours

Special Instructions

- Not to be used first line.
- To be used only on the recommendation of a Palliative Care Specialist.
- Dilute with Sodium Chloride 0.9% to the largest possible volume to reduce the likelihood of inflammatory reactions at the skin site.
- Avoid abrupt withdrawal after long-term treatment as this may precipitate biliary colic from gallstones or biliary sludge.

Indications for Use

- Anti-secretory effect.
- Ascites, bronchorrhoea, excessive diarrhoea, associated with for example, malignancy, chemotherapy or radiotherapy, malignant fistulae, large volume vomiting associated with inoperable bowel obstruction, malignancy related mucous secretion.
- Once control of symptoms is achieved it may be possible to reduce to a lower maintenance dose, maintaining control whilst minimising dose-dependent undesirable effects.

Mechanism of Action

Octreotide has various actions as a somatostatin analogue. For the anti-secretory effect, it acts by reducing intestinal secretions of water and Sodium, in addition to stimulating absorption of water and electrolytes from the gastrointestinal tract. It may also improve gastrointestinal motility.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Commonly observed effects include - constipation/diarrhoea at the beginning of therapy, dry mouth, flatulence (reduce dose and increase slowly), anorexia, abdominal pain, abdominal bloating, nausea, steatorrhoea (>500microgram daily), gallstones (10-20% of patients on long term treatment), pancreatitis associated with gallstones, hypoglycaemia shortly after starting treatment and hyperglycaemia (with chronic administration), hyperkalaemia, dizziness, hair loss.

Caution

- In type 1 diabetes insulin requirements may be reduced by up to 50%.
- For type 2 diabetes insulin and oral/parenteral hypoglycaemic agents may need adjusting. Close blood glucose monitoring should guide these adjustments.
- Hepatic impairment (dose reduction may be necessary).
- Monitor thyroid function (risk of hypothyroidism on long term treatment).

Compatibilities/Incompatibilities

Due to complexities around drug compatibilities, including Octreotide, please liaise with the Specialist Palliative Care Team, ELHT Pharmacy Team of out of hours from the 24 hour Palliative Care Advice Line before considering 2-drug combinations.

Presentations

Octreotide 50microgram/mL 1mL ampoule. Octreotide 100microgram/mL 1mL ampoule Octreotide 500micrograms/mL 1mL ampoule

Ondansetron

Usual dose: 8mg to 24mg over 24 hours

Special Instructions

Dilute with Sodium Chloride 0.9% or Water for Injection.

To be used only on the recommendation of a Palliative Care Specialist.

Indications for Use

Use of Ondansetron in Palliative Care remains limited to intractable or challenging cases of nausea and vomiting when the situation suggests that Serotonin release is the cause of the nausea and vomiting such as:

- · Chemotherapy.
- Radiation-induced damage of the GI mucosa.
- Intestinal distension.
- Leaky platelets in severe renal impairment.

Mechanism of Action

Ondansetron is a selective 5HT3 serotonin.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Common: Headache, constipation, flushing, injection site reactions.

Caution/Drug Interactions

Dose should be reduced in moderate or severe hepatic impairment (max, 8mg daily).

Additive effect with other drugs that cause QT interval prolongation, for example Citalopram, Erythromycin, Haloperidol, Domperidone, Levomepromazine, Methadone.

Additive effect with other drugs that cause Serotonin toxicity, for example SSRIs, Fentanyl, Tramadol, Metoclopramide, Tricyclic Antidepressants, Venlafaxine, Duloxetine.

Contraindicated with Apomorphine – risk of severe hypotension.

Compatibilities

Please liaise with the Specialist Palliative Care Team, ELHT Pharmacy Team or out of hours the 24 hour Palliative Care Advice Line before considering 2-drug combinations.

Preparations

Ondansetron 2mg/mL 2mL ampoule (4mg/2mL ampoule).

Ondansetron 2mg/mL 4mL ampoule (8mg/4mL ampoule).

Oxycodone

Usual dose: The initial dose of Oxycodone is dependent on the

patient's present opioid requirements and patient factors,

for example renal function - seek advice if needed.

There is no maximum dose of Oxycodone providing it is

carefully titrated

Special Instructions

Dilute with Water for Injection. Sodium Chloride 0.9% may also be used.

Indications for Use

Moderate to severe cancer and non-cancer pain. An alternative to other strong opioids in case
of intolerance.

Mechanism of Action

Oxycodone is a strong opioid and acts primarily via µ-opioid receptor.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects. Common: nausea and vomiting, constipation, drowsiness, dry mouth.

Less common: pruritus, sweating.

Neurotoxicity: hyperalgesia, allodynia, delirium, hallucinations, myoclonus - can develop with

deteriorating renal function.

Caution

Dose reduction may be needed in patients with hepatic and/or renal impairment

- Mild to moderate renal impairment start with lower dose and titrate cautiously/slowly.
- Severe renal impairment increased risk of toxic side effects due to drug and metabolite accumulation. Use with caution and monitor on a regular basis, especially for symptoms of neurotoxicity (see above). Alternative opioid, such as Alfentanil might be preferable depending on circumstances. Please discuss with the Specialist Palliative Care Team, or out of hours with the 24 hour Palliative Care Advice Line.
- Moderate to severe hepatic impairment avoid if possible. Please discuss with the Specialist Palliative Care Team or out of hours with the 24 hour Palliative Care Advice Line.

Preparations

Two strengths of injection are available, 10mg/mL and high strength 50mg/mL. The latter may be useful in situations where high doses cause volume difficulties in a Syringe Pump.

Syringe Pump compatibilities when using Oxycodone 10mg/mL

There is 2-drug compatibility data using Water for Injection as diluent with Glycopyrronium, Haloperidol, Hyoscine Butylbromide, Hyoscine Hydrobromide, Levomepromazine, Metoclopramide, and Midazolam.

Limited 3 drug compatibility information can be found on pages 18-19. For any other drug combinations please discuss with the Specialist Palliative Care Team, the ELHT Pharmacy Team or out of hours the 24 hour Palliative Care Advice Line.

Compatibilities when using Oxycodone 50mg/mL

Differences in compatibility with other drugs for the 10mg/mL and 50mg/mL formulations of Oxycodone have been reported. It is important not to extrapolate compatibility information from one formulation to the other. Please seek advice from Specialist Palliative Care or Pharmacy.

Incompatibilities

Concentration-dependent incompatibility may occur when mixed with Cyclizine.

Preparations

10mg/mL 1mL and 2mL ampoules. 50mg/1mL ampoule.

Parecoxib

Usual dose: 40mg subcutaneously once daily, can be increased to

twice daily

Can be given via syringe pump if preferred (40-80mg

over 24 hours)

Parecoxib must only be prescribed after discussion with a Consultant in Palliative Medicine or, if a Consultant is unavailable out of hours, then with a Senior Hospice Physician.

Ensure 0.9% Sodium Chloride solution for injection ampoules are prescribed with Parecoxib to enable administration

Special Instructions

Parecoxib is supplied in a vial containing 40mg of powder, this must be reconstituted with 2mL 0.9% Sodium Chloride. This can then be given by subcutaneous injection or diluted further with 0.9% Sodium Chloride to be given via syringe pump.

Consider gastroprotection if able to take. Stop other NSAIDs.

Consider if once or twice daily SC injection is preferable to syringe pump.

Indications for Use

Pain not responding to other measures, if an anti-inflammatory analgesic is indicated and the oral route is not available. Seek Specialist Palliative Care advice.

Mechanism of Action

Parecoxib is a selective inhibitor of cyclo-oxygenase-2 (COX2). By inhibiting COX2, parecoxib reduces the production of inflammatory prostaglandins, and so reduces inflammation and pain.

Parecoxib is a non-steroidal anti-inflammatory drug (NSAID), but whereas non-selective NSAIDs inhibit COX1 and COX2, Parecoxib mainly only inhibits COX2, and so there are some differences in side effects.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

Parecoxib can cause similar side effects to other NSAIDs:

- Parecoxib has a lower risk of causing gastro-intestinal side effects and has less effect on platelets and bleeding risk compared to non-selective NSAIDs, but a higher risk of cardiovascular events.
- Severe skin reactions can occur but are uncommon or rare. Parecoxib should be discontinued at the first appearance of a skin rash, mucosal lesion or sign of hypersensitivity.
- Renal function can deteriorate consider if monitoring is appropriate

Subcutaneous injection or syringe pump site reactions can occur.

Cautions

Refer to the manufacturer's SPC for a detailed list of contraindications and precautions. Contra-indicated if hypersensitivity to aspirin, other NSAIDs or sulfonamides.

Usually contra-indicated in patients with:

- Established heart failure
- Severe hepatic impairment (serum albumin <25g/L or Child-Pugh score ≥10)

- Active peptic ulceration or recent GI bleeding
- Inflammatory bowel disease
- Established heart disease, peripheral arterial disease and/or cerebrovascular disease

For patients in the last days and weeks of life, the prescriber must consider these conditions, but they may not necessarily be a deterrent to use, provided the dose is carefully titrated.

Use with caution in patients with:

- Significant risk factors for cardiovascular events
- High risk of developing GI toxicity, or prior history of peptic ulceration and GI bleeding
- Hepatic impairment if moderate impairment (Child-Pugh score 7-9) reduce initial dose to 20mg and maximum dose to 40mg
- Renal impairment if severe impairment (creatinine clearance <30mL/min), reduce initial dose to 20mg. No dose adjustment is needed in mild to moderate renal impairment
- Elderly patients weighing less than 50kg initial dose 20mg, maximum dose 40mg

Lower doses are recommended if the patient is also taking Fluconazole.

Compatibilities

Because the lack of compatibility data, parecoxib should not be mixed in the same syringe with any other drugs.

Consider if once or twice daily SC injection is preferable to syringe pump.

Preparations

40mg powder for solution for injection/ 40mg powder and solvent for injection (0.9% Sodium Chloride)

Phenobarbital

Usual dose: 200 to 1,600mg over 24 hours (doses up to 3,800mg/24h

have been used)

Phenobarbital must only be prescribed after discussion with a Consultant in Palliative Medicine, or if a Consultant is unavailable out of hours, with a Senior Hospice Physician.

Special Instructions

Dilute with water for injection. Sodium Chloride 0.9% may also be used.

Indications for use

- **Terminal agitation** third-line for patients who failed to respond to the combined use of Midazolam and an antipsychotic.
- **Status epilepticus** if other treatments have not been effective and admission to hospital or ICU is not appropriate given the patient's diagnosis and prognosis, i.e. in dying patients.
- Maintenance anti-epileptic in patients unable to swallow and where a Syringe Pump route is preferred over other routes Phenobarbital is a third-line alternative to Midazolam or Levetiracetam.

Mechanism of Action

Enhances post-synaptic inhibitory action of GABA by prolonging the opening of the chloride channel in the GABA-receptor-channel complex.

Side Effects

Refer to manufacturer's SPC for a detailed list of adverse effects.

- Paradoxical excitement.
- Local necrosis following s/c injections bolus/stat doses to be given by deep IM injection.

Caution

- Unless in the imminently dying, it is contraindicated in severe renal impairment or severe hepatic impairment.
- Use with caution in patients with severe respiratory depression, mild to moderate hepatic or renal impairment.
- Enzyme induction:
 - Phenobarbital is metabolised by CYP2C19 clinical significance of co-administration with CYP2C19 inhibitors is unknown. The effect of Phenobarbital might be enhanced by Fluconazole, Fluoxetine and Omeprazole.
 - Phenobarbital is a potent inducer of CYP3A4. Enzyme induction may take up to 2 weeks to develop, so clinical significance when used in the last days of life is unknown. It may reduce the effect of the following drugs: Alfentanil, Clonazepam, Dexamethasone, Fentanyl, Haloperidol, Methadone, Midazolam, Ondansetron, Oxycodone and Tramadol.

Compatibilities/Incompatibilities

Phenobarbital should be administered by a separate Syringe Pump. Due to its alkaline pH, it is likely to be incompatible with most palliative care drugs.

Preparations

Phenobarbital (Phenobarbital Sodium) 200mg/mL ampoules.

Preparations with lower concentrations are available, but due to volume constraints the 200mg/mL strength should be used.

Ranitidine

Usual dose: 100mg to 200mg over 24 hours

Special Instructions

Ranitidine can be diluted with Water for Injection or Sodium Chloride 0.9%.
Ranitidine should be the last drug added to an already diluted combination of drugs.

Indications for Use

• Inoperable bowel obstruction to reduce volume of gastric secretions.

Mechanism of Action

H2 – receptor antagonist inhibits gastric acid secretion and reduces volume of gastric secretions.

Side Effects

Refer to the manufacturer's SPC for a detailed list of adverse effects. Usually well tolerated. Constipation, nausea.

Caution

Dose reduction required in renal and hepatic impairment – seek advice. Dose reduction may be required in the elderly. Avoid in patients with acute intermittent porphyria.

Compatibilities

Seek specialist advice.

Incompatibilities

Seek specialist advice.

Preparations

Ranitidine 50mg/2mL ampoules.

AMBULATORY SYRINGE PUMP PRESCRIPTION (ETS443)

AMBULATORY SYRINGE PUMP PRESCRIPTION (ETS443)

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		NE	15	Trust
A University	Te	eachir	na	Trust

Patient Name:			Date of Birth	:	Instructions for use: Use one prescription chart for each
NHS/RXR No:			Patient Location:		Ambulatory syringe pump
Allergies:		(Consultant/G	P:	Commence a new chart where there are changes to the contents of the syringe pump
Approved name of modicine		Dose	Route/	Indication	It is the responsibility of the prescriber to ensure all prescribed drugs are compatible
Approved name of medicine (please delete unused lines)	(IN WORE	S AND FIGURES)	Rate	indication	If more than three medicines are required
	6	S	C/24hr		specialist advice MUST be sought
	ž	9	C/24hr		All medication should be mixed with water for injection unless known incompatibility
		S	C/24hr		Final volume includes all prescribed medication and diluent, if final volume exceeds 22mLs seek specialist advice
Specialist advice must be sought if 4 drugs to be used		S	C/24hr		On discharge: Keep original prescription. Write a new prescription for community.
Pharmacist clinical check (Hos	**************************************	ST BE COMPLETED			On admission: Send prescription details with patient
Pharmacist Name:	Sig	nature:	Date:	Time:	
Diluent required:	Final volume:	Buprenorphine/Fentanyl pa			Patient information leaflet given Y □ N □
Water for injection □	17mL □	Y □ N □ (If Y please contine Patch strengthm			For advice on syringe pumps please contact:
Sodium chloride 0.9%	22mL □	This is equivalent to	78 60		Specialist Palliative Care Team: Hospital: Mon-Fri 08:30-16:30 Tel: 01254 732316 Community: Mon-Fri 08:30-16:30 Tel: 01254 736326 Hospice 24/7 out of hours advice line: 07730 639399
Prescriber name (print):		scriber must ensure start e completed to authorise tion	Stop date: Reason for	Time: discontinuing:	Pharmacy: Medicines information Mon–Fri 08.30-17.00 Tel:
Prescriber signature: Start immediately □		nediately	Name (print	t):	01282 803004. At all other times contact the <u>on-call</u> pharmacist via hospital switchboard.
Date: Time:	33000.000.000	fy time	Signature:		Palliative care syringe pump compatibility ref. vww.pallcare.info/www.palliativedrugs.com

ETS443

AMBULATORY SYRINGE PUMP PRESCRIPTION (ETS443)

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Available
concentrations
1mg/2mL
5mg/1mL
50mg/ 1mL
200microgram/ 1mL
5mg/1mL
20mg/ 1mL
25mg/ 1mL
10mg/ 2mL
10mg/ 2mL
10mg/ 1mL
30mg/ 1mL
10mg/ 1mL
20mg/ 2mL
50mg/ 1mL

AMBULATORY SYRINGE PUMP ADMINISTRATION AND MONITORING RECORD (ETS442)

AMBULATORY SYRINGE PUMP ADMINISTRATION AND MONITORING RECORD (ETS442)

Hospital No: RXR		Dat	e Time	Patient location		ncashire Hospital NHS Tru A University Teaching Tru					
Checks to be recorded:	Syringe pump set u	up by:			Medicine	Dose					
Inpatient 4hourly	Name:										
Outpatient twice daily	Signature:										
Syringe pump prescription:	Checked by:										
 Must have all sections 	Name:										
 If any information is 	Signature:	Signature:									
	Diluent used: Water for injection Sodium Chloride 0.9%										
to medical staff before	Prescription fully o	Prescription fully completed? Y N N									
administration	0 Hr	+4 Hrs	+8 Hrs	+12Hrs	+16 Hrs	+20 Hrs					
Time	:	:	=	=	Ξ	-					
Volume in syringe (mL)											
Time remaining of infusion											
Record rate of infusion (mL/hr)											
Is the pump running to time? (Y/N)											
Remaining battery power (%)											
Green LED flashing (Y/N)											
Syringe secure in pump (Y/N)											
Is the fluid clear (Y/N)											
Is keypad lock on (Y/N)											
Site position											
Site inflamed/ red (Y/N)											
Checked by: Name											
Signature											

ETS442

PALLIATIVE CARE COMMUNITY ANTICIPATORY/SUPPLEMENTARY SUBCUTANEOUS MEDICINES AUTHORISATION SHEET

PALLIATIVE CARE COMMUNITY ANTICIPATORY/SUPPLEMENTARY SUBCUTANEOUS MEDICINES AUTHORISATION SHEET



Patient Name:	Patient location:	
NHS/RXR No:	Date of Birth:	
Consultant/GP:	Ward/Community Nursing Team:	
Known allergies/alerts:	Hospital/Community prescription (please circle)

indication	Medicine	Dose	Frequency	Max 24 hr dose to be given PRN	Route	Prescriber Signature		Date/Time discontinued (inc. signature)
						Prescriber's Signature		
						Print Name	Date	
						Prescriber's Signature		
						Print Name	Date	
						Prescriber's Signature		
						Print Name	Date	
						Prescriber's Signature		
						Print Name	Date	
						Prescriber's Signature		
						Print Name	Date	
						Prescriber's Signature		
						Print Name	Date	

PALLIATIVE CARE COMMUNITY RECORD OF ANTICIPATORY/SUBCUTANEOUS SUPPLEMENTARY MEDICINES AND INJECTABLE MEDICINES STOCK RECORD

PALLIATIVE CARE COMMUNITY RECORD OF ANTICIPATORY/ SUBCUTANEOUS SUPPLEMENTARY MEDICINES AND INJECTABLE MEDICINES STOCK RECORD (ETS446)



Patient Name:	Date of Birth:	NHS Number:1`	Medicine & Strength:

Date & Time Given	Batch No./Expiry Date	Balance	Medicine	Dose	No. of Ampoules Used	Site – Subcut	Site – Syringe Pump	New Stock	Stock Balance	Signature and Print Name
										HCP 1
										HCP 2
										HCP 1
										HCP 2
										HCP 1
										HCP 2
										HCP 1
										HCP 2
										HCP 1
										HCP 2
										HCP 1
										HCP 2
										HCP 1
										HCP 2

THE USE OF AN ADDITIONAL SUBCUTANEOUS CANNULAE TO ADMINISTER PRN ANTICIPATORY MEDICATION

ANTICIPATORY MEDICATIONS MUST NEVER BE GIVEN DOWN THE SYRINGE PUMP LINE

An additional cannulae can be inserted for administration of PRN anticipatory subcutaneous medication if necessary

Please refer to the Procedure for the Administration of Subcutaneous PRN Medication Using a Prescribed Range of Doses for Symptoms in the Last Days of Life. SOP0729Version 2

Appendix 2 gives details about the use of subcutaneous cannulae to administer PRN anticipatory medication

PATIENT AND CARER INFORMATION LEAFLET FOR YOUR SYRINGE PUMP

For further information please contact the following:

Hospital Specialist Palliative Care Team

Tel: 01254 732652

Community Specialist Palliative Care Team

Tel: 01254 736326



Safe Personal Effective

Community and Integrated Care Division East Lancashire Hospital Trust Royal Blackburn Hospital Haslingden Road Blackburn

BB2 3HH

INVESTORS IN PEOPLE







Patient and Carer Information Leaflet for your Syringe Pump



Produced with the public of East Lancashire

®NHS East Lancashire 2011 Author of leaflet: Specialist Palliative Care Team Version number: Version 5 Date of next review: June 2025 Doc ID – SPC-004-syringepump-2020



What is a Syringe Pump?

A Syringe Pump is a small portable battery operated pump, holding a syringe. It allows medicines to be given steadily under the skin via a small needle over a 24 hour period. The pump will hopefully reduce the need for repeated injections.

Why do I need one?

A Syringe Pump is used to give you your medicines in an alternative way for various reasons – for example:

- · You may be struggling to swallow medicines
- You may have nausea/vomiting which can affect the way medicines are absorbed
- · To control your symptoms more effectively

Your Nurse or Doctor will discuss the reasons for starting a Syringe Pump with you and your family/carer. A Syringe Pump can be used at any stage of your illness to control your symptoms and if you become able to take oral medicines it may be possible to discontinue the Syringe Pump.

Who looks after the Syringe Pump?

Your nursing team will reload the syringe with your medicines every 24 hours. The device will be checked every time you are seen by a nurse to ensure the pump is operating correctly.

How do I know it is working?

- While the pump is running the indicator light will flash approximately every
 - 30 seconds.
- If the alarm sounds contact the nurses involved in your care immediately to check the device.

Taking care of yourself with a Syringe Pump

- Tell your Nurse if you have any redness or soreness where the needle is placed.
- · Contact your Nurse if the needle comes out or dislodges.

Side Effects:

Your Nurse or Doctor will explain possible side effects from the medicines in the syringe pump. All medicines have the potential to cause side effects. Please speak to your Nurse or Doctor if you have any concerns.

IN THE COMMUNITY



- Check your medicines are stored safely away from children/pets.
- Discuss bathing/showering needs with your Nurse.
- Contact your Nurse if the medicines change colour or become cloudy.



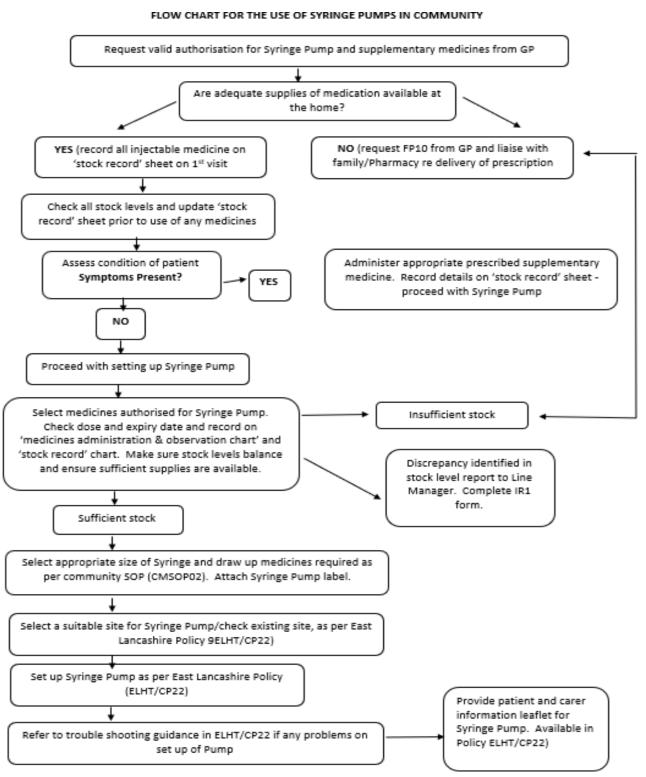
Some Don't

- · Do not interfere with the device.
- Do not place the device near extremes of heat, e.g. hot water bottles.
- Don't get the pump or needle site wet.

Please ensure the Syringe Pump is given to the District Nursing Team when no longer required

Contact Telephone Numbers:
Ward/Community Team:
GP/Doctor:

FLOW CHART FOR THE USE OF SYRINGE PUMPS IN COMMUNITY



Appendix 1 Equality Impact Assessment Screening Form

Department/Function	Specialist Palliative Care/Syringe Pump Policy					
Lead Assessor	Syringe pump task and finish group chair					
What is being assessed?	Syringe Pump Policy					
Date of assessment	24/1/22					
What groups have	Equality of Access to Health Group		Staff Side Colleagues			
you consulted with? Include details of	Service Users		Staff Inclusion Network/s			
involvement in the Equality Impact	Personal Fair Diverse Champions		Other (Inc. external orgs)			
Assessment Please give details: Syringe pump task and finis group						

1) What is the impact on the following equality groups?						
Positive: > Advance Equality of opportunity > Foster good relations between different groups > Address explicit needs of Equality target groups	haras victim > Failur explic	Negative: vful mination, esment and hisation re to address cit needs of lity target groups	Neutral: It is quite acceptable for the assessment to come out as Neutral Impact. Be sure you can justify this decision with clear reasons and evidence if you are challenged			
Equality Groups	Impact (Positive / Negative / Neutral)	Comments Provide brief description of the positive / negative impact identified benefits to the equality group. Is any impact identified intended or legal?				
Race (All ethnic groups)		Neutral				
Disability (Including physical and mental impairments)	Select	Neutral				
Sex	Select	Neutral				
Gender reassignment	Select	Neutral				
Religion or Belief	Select	Neutral				
Sexual orientation	Select	Neutral				
Age	Select	Neutral				
Marriage and Civil Partnership	Select	Neutral				
Pregnancy and maternity	Select	Neutral				

mamam riginio)	
2) In what ways does any impact identified contribute to or hinder promoting equality and diversity across the organisation?	Impact neutral

Select Neutral

- 3) If your assessment identifies a negative impact on Equality Groups you must develop an action plan to avoid discrimination and ensure opportunities for promoting equality diversity and inclusion are maximised.
- > This should include where it has been identified that further work will be undertaken to further explore
- > the impact on equality groups
- > This should be reviewed annually.

Action Plan Summary

Other (e.g. caring,

human rights)

Action	Lead	Timescale

This form will be automatically be inserted as an appendix in all Policies and Procedures which are presented for ratification at the Policy Council. Please do not hesitate to contact the qualityandsafetyunit@elht.nhs.uk if you have any queries.

Appendix 2 Useful Telephone Numbers

Hospital Specialist Palliative Care Team (Monday-Friday 08.30-16.30)	01254 732652/ 01254 732316	Ext 82652 Ext 82316
Community Specialist Palliative Care Team (Monday – Friday 08.30-16.30)	01254 736326	Ext 86326
Pendleside Hospice	01282 440100	
East Lancashire Hospice	01254 965830	
Specialist Palliative Care Out of Hours 24 Hour Advice Line (based at East Lancashire Hospice)	07730 639399	
Medical Equipment Library	01254 733660	Ext 83660
Medicines Information (Monday – Friday 08.30 17.00)	01282 803004	Ext 13004
Pharmacy Aseptic Unit (Monday – Friday 08.00 16.30 Saturday-Sunday 09.00-13.00))	01254 734680	Ext 84680
Pharmacy Dispensary RBH (Monday – Friday 08.30 17.00 Saturday-Sunday 09.00-16.00)	01254 732252	Ext 82252
Pharmacy Dispensary BGH (Monday – Friday 09.00 17.00)	01282 804338	Ext 14338
North West Medicines Information Centre (wmedinfo@nhs.net)	0151 794 8113	

APPENDIX 3 Community Nursing Competency Tool

Community Nursing Competency Tool

Competency Assessment for Health Care Assistants/ Assistant Practitioners to fulfil the role of the Second Checker of Controlled Drugs Prepared & Administered Via A Syringe Driver.

Name	Team	Competency Assessment
		, ,
DateCommenced		

Competency Number	KNOWLEDGE & ABILITY: The HCA / Assistant Practitioners can:	Self Assessment: HCA/AP Comments	Formal Assessment by Assessor (Achieved Yes/No)	Reviewer's signature	Reviewee's signature
1.	Identify relevant policies, procedures and SOPs relating to controlled drugs, syringe drivers and incident reporting?				
2.	What training have you undertaken to support you in this role?				
3.	Give the correct definition of a Controlled Drug				
4.	Articulate the rationale for the role of witness for preparing (controlled) drugs for administration via a syringe driver.				
5.	Demonstrate a clear understanding of the purpose and function of drug stock sheets				
6.	Demonstrate a clear understanding of the storage of controlled drugs in a person's own home or residential home				

Competency Number	Knowledge & Ability: The HCA/AP can:	Self Assessment: HCA/AP	Formal Assessment By Assessor (Achieved Yes/No)	Reviewer's Signature	Reviewee's Signature
7.	Describe what you would do if you identified any discrepancies with controlled drug stocks.				
8.	Describe the correct process for the disposal of controlled drugs in the community setting				
9.	Demonstrates correct hand hygiene before participating in the process				
10.	Remains with the administering nurse from start to finish of the administration & can articulate the rationale for this.				
11.	Makes an independent check that the identity of the patient is correct and corresponds with the prescription chart				
12.	Makes an independent check of the stock balance				
13.	Is able to complete drug stock sheets accurately-				
14.	Checks the prescription is legible				
15.	Checks the medication is due or can be given at this time if PRN				
16.	Confirms the medication has not already been administered				

Competency Number	Knowledge & Ability: The HCA/AP can:	Self-Assessment: HCA/AP	Formal Assessment By Assessor (Achieved Yes/No)	Reviewer's Signature	Reviewee's Signature
17.	Confirms the name on the medication box /box contents/ vial/ bottle corresponds to the medication name written on the prescription chart				
18.	Checks that the medication has not passed its expiry date				
	tional areas of competency assessed	below			
19.					
20.					
OUTCOME:					

The Assistant practitioner /health care assistant* is able to demonstrate safe practice to the assessing nurse to check controlled drugs administered via a syringe driver. Sign off when all areas competently achieved.

• Delete as appropriate

Final sign off achieved	
Sign/Print	Assessor, Designation of Assessor
Date	
Sign/Print	Nurse Assessed.
Date	

Notes on completing competency assessments:

Staff are required to attend for update medication management training as defined within the training matrix (see intranet for latest version) every three years. Staff may also attend the training session more frequently if they wish to do so or if a training need is identified as part of their PDP.

Performance Criteria	The AP/ Health care assistant will :
Locate the relevant trust protocols	Knows where and to find the relevant trust protocols
Give the correct definition of a Controlled Drug	Can explain the legal difference between controlled medication and general
_	medications
Articulate the rational for the role of witness	Can explain the role and responsibilities of the witness and why a witness is required
Demonstrate a clear understanding of the need for the	Can describe the storage and security requirements for CDs
storage and security measures taken for CDs	
Understands the need to inform the Clinical Team Leader	Can explain why discrepancies involving CD recording and stock balances must be
immediately of any discrepancies with CDs	reported immediately
Describe the correct process for the wastage of CDs	Knows how CDs should be disposed of and is able to point out when this is being
	done incorrectly
Ensures correct hand hygiene before participating in the	Follows ELHT hand hygiene policy and completed self- declaration on learning hub.
process	
Remains with the administering nurse from start to finish of	Knows how CD administration should be carried out, can explain the reasons for
the administration and can articulate the rational for this	observing the whole process and can point out when this is being done incorrectly
Makes an independent check that the identity of the	Is able to check independently the identity of the patient, understands the importance
patient is correct and corresponds with the prescription	of this and can explain why s/he would not simply accept the word of the qualified
chart	nurse
Makes an independent check of the stock balance	Knows the correct procedure for confirming stock balance for CDs
(counting remaining ampoules, making a visual estimate of	makes an independent count of CD's and can explain the importance of this as
liquids)	opposed to confirming what the qualified nurse states
Checks the prescription is legible	Ensures s/he can read the prescription clearly
Checks the medication is due.	Knows how to check that the drug is being given at the correct time if regular
	medication.
Confirms the medication has not already been	Knows how to check the prescription chart to confirm the medication has not already
administered	been given

Confirms the name on the medication box /box contents/ vial/ bottle corresponds to the medication name written on the prescription chart	Makes an independent check that the medication being prepared for administration corresponds to the medication named on the prescription chart
Checks that the medication has not passed its expiry date	Knows why and how to ensure the mediation being prepared for administration has not expired
Explain the need for the correct storage and security of medicines	Can explain where and how medicines should be stored in accordance with ELHT and UK guidelines and to meet any specific requirements e.g. away from sunlight etc.

Should a member of staff fail to successfully demonstrate an acceptable level of competency after supervised assessments, this may addressed via the following responses:

- address any additional training requirements
- shadowing other members of the team to observe how they achieve their role requirements
- Performance Improvement Policy and Procedure