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**Note:** This consensus algorithm excludes patients who are in the ICU, perioperative or pre-procedural settings, or are currently receiving epidural or intrathecal analgesia.

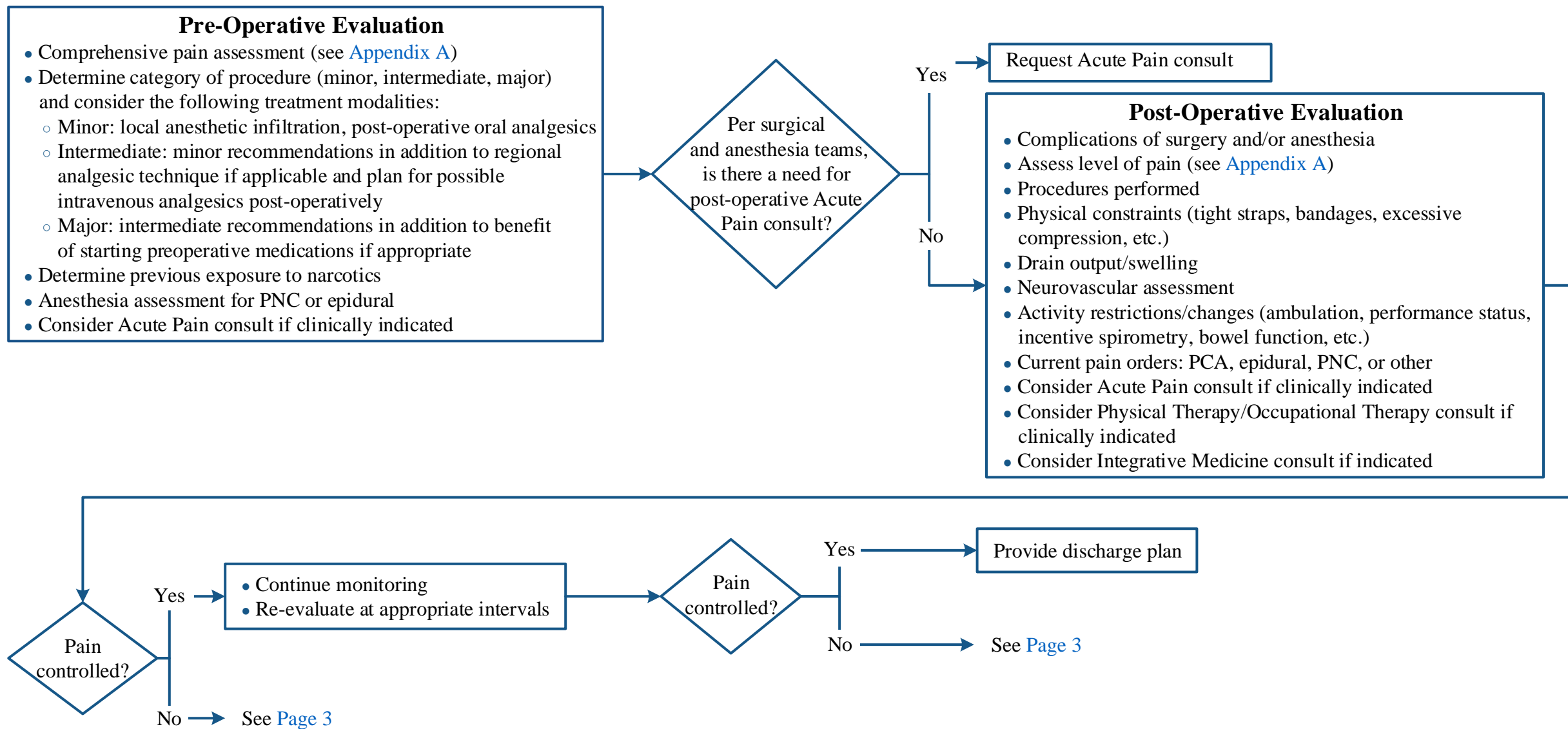
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## ASSESSMENT



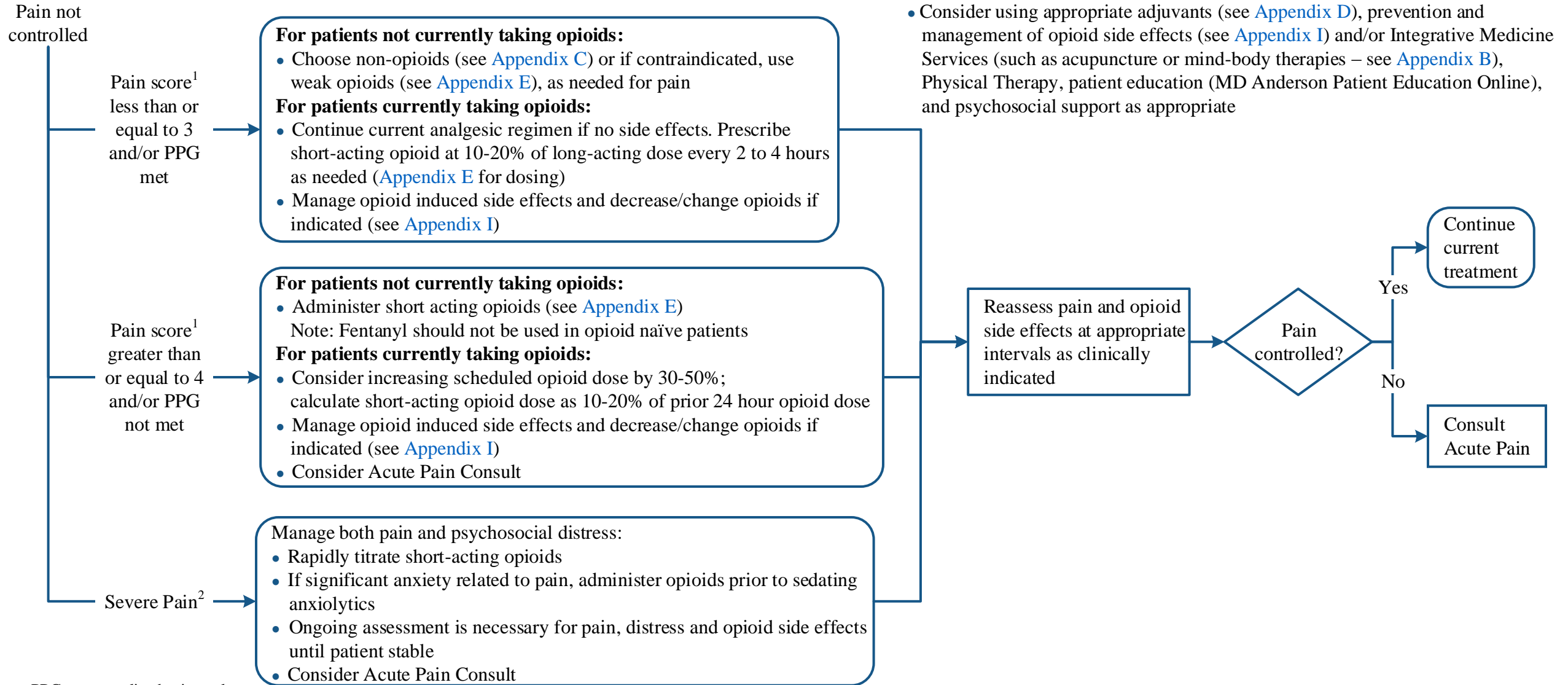
PNC = peripheral nerve catheter  
PCA = patient controlled analgesia

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## TREATMENT



### Note for all patients:

- If patient has a regional anesthetic block, notify appropriate managing service
- Consider using appropriate adjuvants (see [Appendix D](#)), prevention and management of opioid side effects (see [Appendix I](#)) and/or Integrative Medicine Services (such as acupuncture or mind-body therapies – see [Appendix B](#)), Physical Therapy, patient education (MD Anderson Patient Education Online), and psychosocial support as appropriate

PPG = personalized pain goal

<sup>1</sup> See [Appendix A](#)

<sup>2</sup> Severe pain, new onset, or exacerbation of previously stabilized pain, accompanied by significant distress or if present for more than 24 hours

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## Quick Reference Guide

- Opioid naïve: Includes patients who are not chronically receiving opioid analgesic on a daily basis and therefore have not developed significant tolerance.
- Opioid tolerant: Patients who are chronically receiving opioid analgesics on a daily basis. The FDA identifies this group as “receiving at least 60 mg of morphine daily, at least 30 mg of oral oxycodone daily, or at least 8 mg of oral hydromorphone daily or an equianalgesic dose of another opioid for a week or longer.”
- Incomplete cross-tolerance: Reduce dose of new opioid by 30 to 50% when switching from one opioid to another to account for tolerance to a currently administered opioid that does not extend completely to other opioids. Consequently, this phenomenon tends to lower the required dose of the new opioid.
- Dose titration: Adjusting the dose of an opioid should be individualized for each patient. Refer to [Page 3](#) of this algorithm for titration recommendations.
- Dosing frequency: For long-acting opioids, dosing frequency is typically every 12 hours to every 24 hours depending on the agent. Refer to [Appendix E](#) for Opioid Dose Considerations.
- Breakthrough pain: Doses of short-acting opioids for breakthrough pain should be 10 to 20% of the total daily dose given every 1 to 4 hours as needed. Breakthrough opioids can be given as frequently as every 1 hour for oral doses or every 15 minutes if IV (assuming normal renal/hepatic function).
- Elderly/organ dysfunction: Use additional caution when converting opioids in elderly patients (65 years and older), and/or patients with hepatic, renal, or pulmonary dysfunction. Codeine, morphine, hydromorphone, and oxycodone should be used with caution in patients with decreased renal function.
- Opioids **NOT** recommended for cancer pain: Meperidine and mixed agonist-antagonists (pentazocine, nalbuphine, butorphanol) should be avoided.
- Withdrawal symptoms: Nausea, vomiting, diarrhea, anxiety, and shivering are common symptoms of opioid withdrawal. A gradual taper is recommended when discontinuing opioids.
- Overdose: Symptoms may include respiratory depression, constricted pupils, and decreased responsiveness. Naloxone is used to reverse the effects of an opioid. To administer, dilute 0.4 mg/mL (1 mL) ampule into 9 mL of normal saline for total volume of 10 mL to achieve a 0.04 mg/mL concentration, and give 1 mL (0.04 mg) via slow IV push every 2 to 3 minutes until patient more awake and respiratory status improves. **DO NOT** administer undiluted due to risk of precipitating rapid withdrawal, which may cause severe pain or seizures.
- Constipation is a common side effect with opioid use. Consider starting a bowel regimen in all patients taking opioids. Refer to [Appendix I](#).
- Duration of drug effect: Any residual drug in the patient's system must be accounted for and an assessment of any residual effects from discontinued long-acting opioids must be made before any new opioid is started. Example: fentanyl will continue to be released from the skin 12 to 36 hours after transdermal patch removal.
- The Texas Prescription Monitoring Program is an electronic database that tracks controlled substance prescriptions. It can help identify patients who may be misusing prescription opioids or other prescription medications and who may be at risk for overdose. Clinicians are encouraged to check the Texas PMP prior to initial opioid prescribing and at regular intervals. It can be accessed at <https://texas.pmpaware.net/login>

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## APPENDIX A: Comprehensive Pain Assessment

### The comprehensive pain assessment should include the following:

#### 1. Pain:

- a. For each site of pain, determine intensity level: 0-10 numeric rating scale (NRS) (no pain = 0, mild = 1-3, moderate = 4-6, severe = 7-10).  
Assess at rest and with activity, location, onset (acute, chronic, acute exacerbation of chronic pain), pathophysiology (somatic, visceral, neuropathic), temporal factors (continuous, intermittent, breakthrough, incidental), etiology (*e.g.*, tumor, non-tumor related, fracture).
- b. Evaluation of medical history includes: oncologic or other significant medical illnesses, medication history, relevant imaging and laboratory studies.
- c. Physical examination.
- d. Assess for presence of sedation [inpatient setting, consider Richmond Agitation Sedation Scale (RASS) and common opioid side effects ([Appendix I](#))].

#### 2. Function:

- a. Evaluate patient's ability to ambulate, perform activities of daily living (ADL), range of motion (ROM), deep breathing, and coughing.
- b. Assess restrictions related to pain.
- c. Report patient's evaluation of functional ability.

#### 3. Psychosocial issues:

- a. Evaluate patient distress, family support, psychiatric history, patient/family knowledge and beliefs surrounding pain and its management, risk factors for under treatment of pain include: underreporting, prior treatment of pain and response to other pain medications, concerns about addiction to pain medications or side-effects, extremes of age, gender, cultural barriers, communication barriers, and prior history of drug abuse.
- b. Report patient's assessment of psychological distress.

#### 4. Personalized Pain Goal (PPG):

- a. Determine the verbal or written goal stated by the patient describing the desired level/intensity of pain that will allow the patient to achieve comfort in physical, functional, and psychosocial domains.

**In addition to Comprehensive Pain Assessment, rule out or treat pain related to oncologic emergencies.<sup>1</sup>**

<sup>1</sup> Pain related to an oncologic emergency requires assessment and treatment (*e.g.*, surgery, steroids, radiation therapy, antibiotics) along with an emergent consultation.

Oncologic emergencies include:

- Bowel obstruction/perforation
- Brain metastasis
- Leptomeningeal metastasis
- Fracture or impending fracture of weight-bearing bone
- Epidural metastasis/spinal cord compression
- Pain related to infection



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## APPENDIX B: Post-Op Specialty Services and Consultation Guidelines

**Integrative Medicine Clinic services** relative to pain management include:

- Acupuncture
- Meditation/mind-body techniques
- Massage
- Music therapy
- A variety of individual and group services are also available for long-term issues

Note: For Integrative Medicine Center services, submit a referral order for a specific service or an evaluation of the use of integrative therapies for symptom management.

### Rehabilitation Services

**Physical and Occupational therapy services and programs relative to pain management:**

- Education and activity modification
- DME and adaptive equipment recommendation and issue
- Modalities: thermal, ultrasound, electrical stimulation, IFC (interferential current), paraffin wax, TENS
- Positioning and joint protection
- Exercise and activity prescription

**Physical Therapy specialized inpatient and outpatient services:**

- Lymphedema management, assessment and treatment, compression garment assessment and recommendation
- TENS (IFC, biofeedback or equivalent) trial and issue
- Ergonomics assessment and training
- Low back pain programs
- Manual therapy, soft tissue mobilization and joint mobilization techniques
- Amputee programs: desensitization, mirror therapy
- Orthotics and prosthetics evaluation and recommendation
- Assistive device (walker, cane, crutches) evaluation and prescription
- Hydrotherapy and wound care

**Occupational Therapy specialized inpatient and outpatient services:**

- ADL, work and leisure activity modification
- Fatigue management/energy conservation
- Upper extremity amputee programs
- Custom splinting and bracing
- Psychosocial support
- Adaptive equipment assessment and prescription
- Wheelchair evaluation and recommendation (custom or rental)
- Seat and back cushion evaluation and recommendation

MD Anderson offers three coordinated pain specialty core services, consisting of Acute Pain Medicine, Chronic Pain Medicine, and Supportive Care. Guidelines for consultation to these services are indicated below.

### Specialty Service Consultation Guidelines:

- If upon pre-operative evaluation it is determined the patient will require either a peripheral nerve catheter (PNC) or epidural, consult Acute Pain Medicine
- If upon discontinuation of PNC or epidural it is determined the patient needs further specialized management by either Acute Pain Medicine or primary team, consider consulting Chronic Pain Management
- If at any time a post-operative patient has uncontrolled pain issues for more than 24 hours, consider consulting Chronic Pain Management
- If a post-operative patient has uncontrolled pain accompanied by multiple symptoms, consider consulting Supportive Care
- If a post-operative patient has uncontrolled pain in the context of cancer in the palliative stage or end of life, consider consulting Supportive Care
- For patients with suspected opioid addiction, request a consult to one of the specialty core services for a referral to a treatment program. See [Appendix J](#) for Treatment Services.

TENS = Transcutaneous electrical nerve stimulation      ADL = activities of daily living

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## APPENDIX C: Non-opioids<sup>1</sup>

**CAUTION:** All of these agents are antipyretic and may mask fever; use caution in patients on myelosuppressive chemotherapy. Non-steroidal anti-inflammatory drugs (NSAIDs) may have antiplatelet effects that can increase the risk of bleeding in patients who are thrombocytopenic or on myelosuppressive chemotherapy and likely to become thrombocytopenic. Non-acetylated salicylates (*e.g.*, salsalate, choline magnesium salicylate) and the COX-2 selected NSAID (celecoxib) may have less effects on platelets, but should still be used with caution in a patient on myelosuppressive chemotherapy.

**Non-opioids include** acetaminophen and nonsteroidal anti-inflammatory drugs (NSAIDs); may be used alone or in combination with opioids for pain management. NSAIDs are useful adjuvant analgesics for bone pain.

**Recommended Starting Doses:** The choice of non-opioid must depend on the individual risk/benefit balance for each patient. The mechanism of action and side effect profile of each option is different.

Drug	Recommended Starting Dose	Maximum Daily Dose	Comments
<b>Acetaminophen</b>	500-1,000 mg PO every 6 hours as needed	4,000* mg	Available PO and per rectum. At higher doses, can cause fatal hepatotoxicity and renal damage. Avoid use in hepatic dysfunction. Does not have anti-inflammatory effect.
	650 mg IV every 4 hours 1,000 mg IV every 6 hours	Single dose: 1,000 mg/dose; Daily dose: 4,000* mg daily	IV acetaminophen is formulary restricted.
<b>Aspirin</b>	500-1,000 mg PO every 4 hours as needed	4,000 mg	Available PO and per rectum. May be difficult to tolerate at analgesic doses due the wide range of side effects. Irreversibly inhibits platelet aggregation.
<b>Ibuprofen</b>	200-800 mg PO every 6 hours as needed	3,200 mg	Inhibits platelet aggregation, can cause gastrointestinal side effects or renal failure. Use with caution in patients at high risk <sup>1</sup>
<b>Naproxen</b>	500 mg PO initial, then 250 mg every 4 hours as needed	1,500 mg	Inhibits platelet aggregation, can cause gastrointestinal side effects or renal failure. Use with caution in patients at high risk <sup>1</sup>
<b>Celecoxib</b>	200-400 mg PO every 12 or 24 hours as needed	400 mg	Does not affect platelet aggregation; can cause renal insufficiency
<b>Ketorolac</b>	15-30 mg IV or PO every 6 hours as needed	120 mg	Limit treatment to 5 days. Reduce dose by 50% if over 65 years or weight less than 50 kg. Use is contraindicated in patients with advanced renal impairment or patients at risk for renal failure due to volume depletion. Inhibits platelet aggregation, can cause gastrointestinal side effects.

<sup>1</sup> Patients at high risk of serious gastrointestinal side effects or renal damage from NSAIDs include: elderly (greater than 60 years old), smokers, previous history of peptic ulcer, currently receiving corticosteroids, anticoagulants, or presence of existing renal disease, cardiac or liver impairment.

\*Manufacturers of over-the-counter acetaminophen recommend no more than 3,000 mg daily

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## APPENDIX D: Adjuvants “Co-analgesics” for Neuropathic Pain Syndromes and Chronic Pain

Drug Class and Uses)	Medication	Recommended Starting Dose	Maximum Daily Dose	Comments
<b>Anticonvulsants</b> (various NP types)	Gabapentin	100-300 mg PO daily	3,600 mg PO per day in 3 divided doses	Used in PHN and NP. May cause drowsiness, dizziness, peripheral edema. Dose adjust for renal impairment.
	Pregabalin	25-75 mg PO twice daily	600 mg PO per day in 3 divided doses	Used in DN, PHN, FM, and NP. May cause drowsiness, dizziness, peripheral edema. Dose adjust for renal impairment.
	Carbamazepine	100 mg PO twice daily	1,200 mg PO per day in 2 divided doses	Used in TGN and NP. Associated with aplastic anemia, agranulocytosis, bone marrow suppression, severe dermatologic reactions, hyponatremia. May cause drowsiness, dizziness, nausea. Significant drug interactions. Avoid in hepatic dysfunction.
	Oxcarbazepine	150-300 mg PO daily	2,400 mg PO per day in 2 divided doses	Used in TGN and NP. Associated with severe dermatologic reactions, hyponatremia. May cause drowsiness, dizziness. Dose adjust for renal impairment.
	Topiramate	25-50 mg PO twice daily	200 mg PO twice per day	Used in NP. May cause acidosis, drowsiness, dizziness, nausea. Dose adjust for renal impairment and hepatic dysfunction.
	Tiagabine	4 mg PO at bedtime	8 mg PO per day	Used in NP. May produce seizures in patients with prior seizure history. May cause drowsiness, dizziness, diarrhea. Use with caution if hepatic dysfunction. Higher doses resulted in increased side effects.
<b>Tricyclic Antidepressants</b> (TCA)	Amitriptyline	10-25 mg PO at bedtime	150 mg PO at bedtime	Consider for continuous and shooting neuropathic pain. Caution in elderly or frail, or patients with glaucoma or arrhythmias. May cause sedation, arrhythmias, dry mouth, orthostasis, urinary retention.  Consider duloxetine for NP, DN. Caution in patients with seizures; avoid MAOIs, other SSRIs or SNRIs due to potential for serotonin syndrome. Duloxetine may increase the risk of bleeding events. Concomitant use of aspirin, nonsteroidal anti-inflammatory drugs, warfarin, and other anti-coagulants may add to this risk. Taper slowly.
	Nortriptyline	10-25 mg PO at bedtime	75 mg PO at bedtime	
	Desipramine	10-25 mg PO at bedtime	150 mg PO at bedtime	
<b>Serotonin-Norepineprine Reuptake Inhibitors</b> (SNRI)	Duloxetine	20-30 mg PO daily	60 mg PO per day	
	Venlafaxine	37.5 mg PO daily	225 mg PO per day	

DN = diabetic neuropathy

FM = fibromyalgia

MAOI = monoamine oxidase inhibitors

NP = neuropathic pain

PHN = post herpetic neuralgia

SNRIs = serotonin-norepinephrine reuptake inhibitors

SSRIs = selective serotonin reuptake inhibitors

TCAs = tricyclic antidepressants

TGN = trigeminal neuralgia

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## APPENDIX D: Adjuvants “Co-analgesics” for Neuropathic Pain Syndromes and Chronic Pain - continued

Drug Class and Uses	Medication	Recommended Starting Dose	Maximum Daily Dose	Comments
<b>Muscle relaxants</b> (muscle pain, spasm)	Baclofen <sup>1</sup>	5 mg PO twice daily	80 mg PO per day in 3 to 4 divided doses	Caution in patients with seizures, cardiovascular disease, glaucoma, myasthenia gravis, renal or hepatic impairment, patients on tricyclic antidepressants or MAOIs, the elderly. May cause anticholinergic effects and significant drowsiness.  Methocarbamol: may repeat course after drug free interval of 48 hours.
	Cyclobenzaprine	5 mg PO three times daily	30 mg PO per day in 3 divided doses	
	Metaxalone	400 mg PO three times daily	3,200 mg PO per day in 3 to 4 divided doses	
	Methocarbamol	500 mg PO four times daily 1,000 mg IV every 8 hours	4,000 mg per day in 3 to 6 divided doses; IV for 3 days maximum if PO not possible	
	Tizanidine	2-4 mg PO at bedtime	36 mg per day in 2 to 3 divided doses	
<b>Corticosteroids</b> (inflammation, nerve compression)	Dexamethasone	Varies by clinical situation (IV or PO) Standard dose 4 -16 mg/day	Varies by clinical situation	May cause impaired healing, infection, thrush, hyperglycemia, weight gain, myopathy, stomach upset, psychosis, emotional instability.

MAOI = monoamine oxidase inhibitors

TCA = tricyclic antidepressants

<sup>1</sup>Intrathecal formulation not on MD Anderson Cancer Center Formulary

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## APPENDIX E: Opioid Dose Considerations (Weaker medications are listed at the beginning of Appendix E)

Opioid	Initial short-acting dose in an opioid naïve patient		Onset (minutes)	Peak effect (hours)	Duration (hours)	Initial scheduled dosing in opioid naïve patients	Available oral dose formulations	Comments
	Route	Dose						
<b>Codeine</b>	PO IV/SC	30 - 60 mg N/A	30 - 60 -	1-1.5 -	4 - 8 -	Short-acting: 30-60 mg every 6 hours Long-acting: N/A	Short-acting: 15, 30, 60 mg tablets Long-acting: N/A	Available alone or in combination with 300 mg acetaminophen <sup>1</sup> . Avoid use in renal and/or hepatic dysfunction.
<b>Tramadol</b>	PO IV/SC	25-50 mg N/A	30 - 60 -	1.5 -	3-7 -	Short-acting: 25 mg PO every 6 hours Long-acting: 100 mg ER daily	Short-acting (IR): 50 mg tablets; Long-acting (ER): 100, 200, 300 mg tablets	Increased risk of serotonin syndrome. <sup>2</sup> May lower seizure threshold. Maximum daily dose 400 mg; consider lower doses if history or increased risk of seizures. Use with caution in renal dysfunction. <sup>3</sup>
<b>Tapentadol</b>	PO	50 -100 mg	less than 60	1.25-1.5	4 - 6	Short-acting: PO every 4-6 hours Long-acting: PO every 12 hours	Short-acting: 50, 75, 100 mg tablets Long-acting: 50, 100, 150, 200, 250 mg tablets	Avoid MAOIs, SSRIs, or SNRIs due to potential risk for serotonin syndrome. New medication orders are restricted to Anesthesiology and Perioperative Medicine, Pain Medicine, or Palliative/Supportive Care. Maximum daily doses: tapentadol IR 600 mg and tapentadol ER 500 mg. Avoid use if creatinine clearance is less than 30 mL/min.
<b>Hydrocodone</b>	PO IV/SC	5-10 mg N/A	10 - 20 -	1 - 3 -	4 - 8 -	Short-acting: 5-10 mg PO every 6 hours Long acting: hydrocodone ER (Hysingla® ER) 20 mg PO once daily hydrocodone ER (Zohydro® ER) (non-formulary) PO 10 mg every 12 hours	Short-acting: 5, 7.5, 10 mg tablets; 2.5 mg/5 mL liquid, in combination with acetaminophen Long-acting: hydrocodone ER (Hysingla® ER) 20, 30, 40, 60, 80, 100, 120 mg tablets hydrocodone ER (Zohydro® ER) (non-formulary) 10, 15, 20, 30, 40, 50 mg tablets	Doses greater than 160 mg/day of hydrocodone ER (Hysingla® or Zohydro® ER) have been associated with increased risk of QTc prolongation. Use with caution in renal dysfunction.

<sup>1</sup> Must consider all forms of acetaminophen (combination and individual) when determining total daily dosing. Manufacturers of over-the-counter acetaminophen recommend no more than 3,000 mg daily.

<sup>2</sup> When used with TCAs, MAOIs, SSRIs, SNRIs, or 2D6 or 3A4 inhibitors

<sup>3</sup> Avoid use of tramadol ER when creatinine clearance is less than 30 mL/min

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## APPENDIX E: Opioid Dose Considerations - continued

(Weaker medications are listed at the beginning of Appendix E)

Opioid	Initial short-acting dose in an opioid naïve patient		Onset (minutes)	Peak Effect (hours)	Duration (hours)	Initial scheduled dosing in opioid naïve patients	Available oral dose formulations	Comments
	Route	Dose						
<b>Morphine</b>	PO	5-15 mg	30	0.5-1	3-6	Short-acting: PO: 5-10 mg every 4 hours; IV: 2-4 mg every 4 hours Long-acting: 15 mg PO every 12 hours, or 20 or 30 mg PO once daily	Short-acting: 15, 30 mg tablets; 10 mg/5 mL, 20 mg/5 mL, 20 mg/mL liquid Long-acting: 15, 30, 60, 100 mg tablets	Available as tablet or liquid preparation. Short-acting preparations can be given via PEG tube. Avoid use in renal dysfunction.
	IV/SC	2-3 mg	5 -10	-	-			
<b>Oxycodone</b>	PO	5-10 mg	10-15	0.5-1	3-6	Short-acting: 5 mg PO every 4 hours. Long-acting: 10 mg PO every 12 hours	Short-acting: 5, 15, 30 mg tablets; 5 mg/5 mL, 20 mg/mL liquid Long-acting: 10, 20, 40, 80 mg tablets	Available alone or in combination with acetaminophen <sup>1</sup> (e.g., oxycodone 5 mg with acetaminophen 325 mg in Percocet®). Use with caution in renal dysfunction.
	IV/SC	N/A	N/A	N/A	N/A			
<b>Oxymorphone</b>	PO	5-10 mg	no data	0.5-1	3-6	Short-acting: 5 mg PO every 4 hours Long-acting: 5 mg PO every 12 hours	Short-acting: 5, 10 mg tablets Long-acting: 5, 10, 20, 40 mg tablets	Poor bioavailability - must be taken on empty stomach. Use with caution in renal dysfunction.
	IV/SC	0.5 mg	5-10	N/A				
<b>Hydromorphone</b>	PO	1-3 mg	15-30	0.5-1	3-5	Short-acting: 2 mg PO every 4 hours IV/SC: 0.5-1 mg every 4 hours Long-acting: 8 mg PO every 24 hours	Short-acting: 2, 4, 8 mg tablets; 1 mg/mL liquid Long-acting: 8, 12, 16, 32 mg tablets	Use with caution in renal dysfunction.
	IV/SC	0.5-1.5 mg	15-30	N/A	4-5			

<sup>1</sup> Must consider all forms of acetaminophen (combination and individual) when determining total daily dosing. Manufacturers of over-the-counter acetaminophen recommend no more than 3,000 mg daily

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## APPENDIX F: Equianalgesic Opioid Dose Conversion

**Note:** The equianalgesic doses (oral and parenteral) can be affected by interpatient variability, type of pain (for example, acute versus chronic), chronic administration, and tolerance. The following table should serve as a guide when switching from one opioid to another. It is recommended to reduce the dose of the new opioid by 30 to 50% to account for incomplete cross tolerance, and to periodically monitor for efficacy and adverse reactions and the dose adjusted accordingly.

Opioid	Oral Dose (PO)	Parenteral Dose (IV/SC)	Conversion Factor: Parenteral to Oral Opioid	Conversion Factor: Oral Opioid to Oral Morphine
<b>Morphine</b>	30 mg	12 mg	2.5	1
<b>Oxycodone</b>	20 mg	N/A	N/A	1.5
<b>Hydrocodone</b>	30 mg	N/A	N/A	1
<b>Oxymorphone</b>	10 mg	1 mg	10	3
<b>Hydromorphone</b>	7.5 mg	3 mg	2.5	4
<b>Fentanyl<sup>2</sup></b>	N/A	120 mcg	N/A	Should be managed by clinicians experienced in pain management

**Methadone and buprenorphine should only be initiated and managed by clinicians experienced in pain management. Consider consult to pain specialists if needed.**

<sup>1</sup> This Equianalgesic Opioid Dose Conversion chart is based on the Centers for Disease Control and Prevention (CDC) recommendations (<https://www.cdc.gov/drugoverdose/resources/data.html>)

<sup>2</sup> See [Appendix G](#) for transdermal fentanyl conversion

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## APPENDIX F: Equianalgesic Opioid Dose Conversion - continued

### Steps for Opioid Rotation:

1. Stop current opioid regimen.
2. Calculate total dose of current opioid (scheduled and PRN doses) used in the previous 24 hour period.
3. Calculate the dose of the new opioid using the equianalgesic dose conversion table (from previous page) and conversion equation (below).

$$\frac{\text{Equianalgesic dose per route of CURRENT opioid}}{24 \text{ hour dose per route of CURRENT opioid}} = \frac{\text{Equianalgesic dose per route of NEW opioid}}{24 \text{ hour dose per route of NEW opioid}}$$

4. Calculate for incomplete cross-tolerance between opioids. Decrease the target dose from step 3 by 30-50% to obtain the new opioid dose.
5. Calculate scheduled pain dose. Divide the new opioid dose (from step 4) by number of doses to be given over 24 hours and administer as scheduled doses.
6. Calculate breakthrough pain dose as 10-20% of calculated new opioid dose to administer PRN every 1 hour.
7. Titrate new opioid regimen until adequate analgesia is achieved.

### Opioid Rotation Example: Rotation from morphine PCA (total daily dose of 120 mg IV) to oral oxycodone.

1. Stop current opioid regimen.
2. Calculate dose of current opioid (scheduled and PRN doses) used in the previous 24 hours which equals 120 mg IV morphine.
3. Calculate the dose of the new opioid using the equianalgesic dose conversion table and conversion equation (below).
  - a. Calculate IV morphine to PO morphine based on conversion table and conversion equation :
 
$$\frac{6 \text{ mg IV morphine}}{120 \text{ mg IV morphine over 24 hours}} = \frac{15 \text{ mg PO morphine}}{X \text{ mg PO morphine over 24 hours}} \quad X = 300 \text{ mg PO morphine}$$
  - b. Calculate PO morphine to PO oxycodone based on conversion table:
 
$$\frac{300 \text{ mg PO morphine}}{X \text{ mg PO oxycodone}} = \frac{15 \text{ mg PO morphine}}{10 \text{ mg PO oxycodone}} \quad X = 200 \text{ mg PO oxycodone}$$
4. Calculate for incomplete cross-tolerance. After a 30-50% dose reduction, the oxycodone dose calculated above should be between 100 and 140 mg per day.
5. Calculate scheduled pain dose. Extended release (ER) oxycodone is dosed every 12 hours; recommend ER oxycodone 60 mg every 12 hours (based on tablet availability).
6. Calculate breakthrough pain dose as 10-20% of 120 mg oxycodone dose and administer PRN every 1 hour.
  - Immediate release (IR) oxycodone is between 12 and 24 mg per dose and may be administered every 1 to 4 hours;
  - Based on tablet availability recommend IR oxycodone 10 to 20 mg every 1 to 4 hours as needed for breakthrough pain.
7. Titrate new opioid regimen until adequate analgesia is achieved.



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## APPENDIX G: Fentanyl

Dosage Forms	Onset	Peak	Duration	Doses Available per Formulary	Comments
Parenteral (IV/Subcutaneous)	Almost immediate	Several minutes	0.5-1 hour	0.05 mg/mL (5 mL vial for injection) PCA syringe supplied as 2,500 mcg/50 mL	
Transdermal patch <sup>1</sup>	12-24 hours	24-72 hours	48-72 hours	12 (delivers 12.5), 25, 50, 75, 100 mcg/hour	Bioavailability 90 %; Do <i>not</i> cut patch, apply heat, or use in patients who develop fever – results in faster onset, shorter duration, and possible overdose.
Transmucosal lozenge (Actiq®)	5-15 minutes	20-40 minutes	Related to blood level	200, 400, 600 mcg	Bioavailability: 50%
Sublingual Tablet (Abstral®)	5-15 minutes	30-60 minutes	2 hours	100, 200, 300, 400, 600, 800 mcg	Bioavailability: 54%

### **Drug specific characteristics:**

- Fentanyl is 80 to 100 times more potent than morphine. Fentanyl is not recommended for initial use in opioid naïve patients. Use in non-opioid tolerant patients may lead to fatal respiratory depression.
- Transdermal fentanyl should only be used in patients with stable opioid requirements. Due to its long half-life of 17 hours, the dose may be difficult to titrate if pain is not well-controlled.
- When initiating transdermal fentanyl, patients should use short-acting opioids as needed until efficacy is obtained (peak effect 24-72 hours).
- Titrate patients on transdermal fentanyl no more frequently than every 3 days after initial dose, and then every 6 days thereafter. Initial evaluation of maximum analgesic effect should not be made before 24 hours.
- Caution with CYP450 3A4 inhibitors, which can increase fentanyl plasma concentrations.
- May be used in patients with renal dysfunction.
- Prior to processing initial prescriptions for **rapid onset fentanyl**, the prescriber must register with the [TIRF REMS Access Program](#) and complete a Prescriber and Patient agreement.

### **Morphine to Fentanyl conversion: 1 mg of IV morphine or 2.5 mg of oral morphine = 10 micrograms of IV fentanyl**

Example: Conversion from oral morphine ER 90 mg every 12 hours to IV Fentanyl

1. 24 hour morphine dose is 90 + 90 = 180 mg
2. Decrease 180 mg by 30 % for incomplete tolerance = 126 mg
3. 1 mg IV morphine = 2.5 mg oral morphine = 10 micrograms IV fentanyl, then new 24 hour morphine dose of 126 mg = 24 hour IV fentanyl dose of 504 micrograms
4. Divide 24 hour fentanyl dose calculated by 24 hours = 21 micrograms/hour

Thus an appropriate starting dose for IV fentanyl/hour (as basal rate in PCA) is 20 micrograms/hour

<sup>1</sup> After Transdermal patch removal, continued absorption of fentanyl occurs from the skin. Delayed administration of another long-acting opioid should be considered due to persistent serum levels of fentanyl. Due to differences in bioavailability, fentanyl products are not interchangeable on a mcg to mcg basis.

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## APPENDIX G: Fentanyl - continued

### Transdermal Fentanyl (TDF) Dosing:

**Option 1:** 2 mg oral morphine approximately 1 mcg *per hour* transdermal fentanyl

Example: Total daily dose of morphine 100 mg translates to approximately 50 mcg transdermal patch, to be applied every 72 hours

**Option 2:** Calculate the total daily dose of morphine and then use the following table to select the appropriate patch strength

Oral Morphine (mg/day)	Transdermal Fentanyl (mcg/hour)
30 to 90	25
91 to 150	50
151 to 210	75
211 to 270	100
Each additional 60 mg/day	An additional 25 mcg/hour

- **Note:** This table should **NOT** be used to convert from TDF to other therapies because this conversion to **TDF** is conservative. Use of this table for conversion to other analgesic therapies can overestimate the dose of the new agent.
- To convert patients to another opioid, remove the transdermal fentanyl patch and titrate the dose of the new analgesic based upon the patient's report of pain until adequate analgesia has been attained. Upon system removal, 17 hours or more are required for a 50% decrease in serum fentanyl concentrations.
- Must prescribe short-acting opioid for breakthrough pain.

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## APPENDIX H: Patient Controlled Analgesia (PCA)

**Suggested initial PCA settings:** All opioid doses must be individualized. (Use the institutional order set for all new PCA orders and dose changes.)

### 1. Opioid naïve patients

Opioid	Demand (PCA) dose (dose range)	Lock out interval (minutes)	1-hour dose limit (optional)	Continuous dose (Basal)	Nurse bolus prn pain	Nurse bolus interval (hours)
<b>Morphine</b> (milligrams)	1 mg (0.5-2.5)	10-30 minutes	4 mg	See below	2-4 mg	2
<b>Hydromorphone</b> (milligrams)	0.2 mg (0.1-0.5)	10-30 minutes	0.8 mg	See below	0.5-1 mg	2
<b>Fentanyl</b> (micrograms)	10 mcg (5-25)	10-30 minutes	40 mcg	See below	25 mcg	2

- a. Patient should be alert and demonstrate ability to administer demand dose for pain. If concerns about cognitive failure or significant anxiety, consider Specialty Consultation: Acute Pain, Chronic Pain, Supportive Care (see [Appendix B](#) for description of services).
- b. Carefully consider adding continuous (basal) dose after 12-24 hours if using frequent demand doses or if pain not controlled. Suggested basal dose is 30-50% of average hourly dose.  
 Example: The 12 hour total morphine demand dose is 20 mg, calculate continuous dose as  $20/12 = 1.7$  mg/hour then  $1.7 \times 0.3$  (30%) = 0.5 mg/hour basal rate.

### 2. Opioid tolerant patients (currently receiving opioid therapy).

PCA orders should take into account the patient's current opioid regimen, clinical situation (severity and etiology of the pain, side-effects from opioids, baseline drowsiness, need for opioid rotation). If there are significant side effects, drowsiness, confusion, respiratory or central nervous system concerns, it is recommended to call for Specialty Consultation: Acute Pain, Chronic Pain, Supportive Care (see [Appendix B](#) for description of services) for PCA ordering.

- a. Calculate total dose of opioid (scheduled and PRN doses) used in the previous 24 hour period.
- b. Use equianalgesic opioid dose conversion table ([Appendix F](#)) to calculate dose of IV opioid being considered for PCA. Decrease dose by 30-50% to adjust for lack of complete cross tolerance to obtain new IV dose.
- c. Divide new IV dose (from above step) by 24 hours, to obtain hourly (basal) dose.
- d. Calculate demand (PCA) dose as 10-20% of new IV opioid dose to use PRN every hour.

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## APPENDIX I: Opioid Side Effects – Prevention and Management

Side Effect	Prevention	Management
Sedation	<ul style="list-style-type: none"> <li>Discontinue other sedating medications if appropriate</li> <li>Educate all patients receiving opioids drowsiness may result for a few days following initiation or increase in opioid regimen.</li> </ul>	<ul style="list-style-type: none"> <li>Consider rotation or dose reduction of opioid if sedation persists</li> <li>Consider psychostimulant:                             <ol style="list-style-type: none"> <li>Methylphenidate (Ritalin®) 2.5-5 mg PO once or twice daily (last dose no later than 4 pm to avoid insomnia). Suggested time 8 am and 12 noon daily. Needs controlled substance (CII) prescription <b>or</b></li> <li>Consider modafinil 100 mg once or twice daily.</li> </ol> </li> </ul>
Opioid Induced Neurotoxicity Risk factors: <ul style="list-style-type: none"> <li>High opioid dose</li> <li>Dehydration</li> <li>Renal failure</li> <li>Pre-existing borderline cognition and/or delirium</li> <li>Use of other psychoactive drugs</li> </ul>	Eliminate nonessential CNS activating or depressing drugs (e.g., benzodiazepines)	<ul style="list-style-type: none"> <li>Consider reversible causes such as metabolic disorders, liver or renal dysfunction, dehydration, hypercalcemia, organic brain disease; treat as appropriate.</li> <li>Consider one or more of the following:                             <ol style="list-style-type: none"> <li>Opioid Rotation (see <a href="#">Appendix E</a>)</li> <li>Opioid dose reduction or discontinuation</li> <li>Discontinue other offending drugs (benzodiazepines)</li> <li>Hydration</li> <li>Symptomatic treatment with haloperidol 1-5 mg PO, IV, or SC every 4 hours PRN</li> </ol> </li> <li>Avoid using naloxone even if delirium is thought to be due to opioid use</li> </ul>
Respiratory depression	<ul style="list-style-type: none"> <li>Monitor sedation and respiratory status (respiratory rate and oxygen saturation) during the first 24 hours in opioid naïve patients</li> <li>Titrate opioids cautiously</li> <li>Consider dose reduction or opioid rotation if patient has excessive sedation.</li> </ul>	<ul style="list-style-type: none"> <li>Call MD, HOLD opioids, provide supplemental oxygen</li> <li>If patient minimally responsive or unresponsive and respiratory rate less than or equal to 6 breaths per minute, administer naloxone. Recommended dose: naloxone 0.4 mg diluted in 9 mL saline for total volume of 10 mL, give 1 mL (0.04 mg) via slow IV push every 2-3 minutes until patient more awake and respiratory status improves. (<i>Half life of naloxone is short and patient may need naloxone infusion for long acting opioids. If no change with naloxone, rule out other causes for the respiratory depression.</i>)</li> <li>If patient is actively dying, DNR (do not resuscitate) and receiving comfort care, naloxone administration may not be appropriate</li> </ul>

Continued on next page

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## APPENDIX I: Opioid Side Effects – Prevention and Management - continued

Side Effect	Prevention	Management
Nausea, Vomiting	<ul style="list-style-type: none"> <li>• Titrate opioid dose slowly and steadily.</li> <li>• Provide antiemetic available with opioid prescription.</li> <li>• Metoclopramide 10 mg PO</li> <li>• Patients at high risk of nausea consider scheduled antiemetic for 5 days and then change to PRN.</li> </ul>	<ol style="list-style-type: none"> <li>1. Investigate for other causes of nausea (for example, constipation, bowel obstruction, chemotherapy or other medications) and treat per guidelines. Initiate scheduled antiemetics. Example: Metoclopramide 5 to 10 mg PO, IV, or SC every 6 hours.</li> <li>2. Add or increase non-opioid or adjuvant medications for additional pain relief so opioid dose can be reduced.</li> <li>3. If analgesia is satisfactory, reduce opioid dose by 25%.</li> <li>4. Consider opioid rotation if nausea remains refractory.</li> <li>5. If nausea remains refractory, consider opioid rotation. (See also Post-Op Nausea and Vomiting Management Algorithm).</li> </ol>
Constipation	<p>Unless alterations in bowel patterns such as bowel obstruction or diarrhea exist, all patients receiving opioids should be started on laxative bowel regimen and receive education for bowel management.</p> <ol style="list-style-type: none"> <li>1. Stimulant laxative plus stool softener: For example: Senokot-S (Senna 8.6 plus Docusate 50 mg), 2 tablets/day and titrate up maximum 9 tablets/day.</li> <li>2. Ensure adequate fluids, dietary fiber and exercise if feasible.</li> <li>3. Prune juice followed by warm beverage may be considered.</li> </ol>	<ol style="list-style-type: none"> <li>1. Assess potential causes of constipation (such as recent opioid dose increase, use of other constipating medications, new bowel obstruction)</li> <li>2. Increase Senokot-S® (or senna and docusate tablets if using separate) and add 1 or both of the following:             <ol style="list-style-type: none"> <li>a. Milk of magnesia oral concentrate (1,200/5 mL) 10 mL PO every 2-4 times daily.</li> <li>b. Polyethylene glycol (Miralax™) 17 grams in 8 ounce beverage daily.</li> </ol> </li> <li>3. If no response to above, perform digital rectal exam (DRE) to rule out low impaction (do not perform if neutropenic, thrombocytopenic, or post-operative bowel surgery). Continue above steps <b>and</b> <ul style="list-style-type: none"> <li>• If impacted: disimpact manually if stool is soft. If not, soften with mineral oil fleet enema before disimpaction. Follow up with milk of molasses enemas until clear with no formed stools.</li> <li>• Consider use of short-acting analgesics before disimpaction.</li> <li>• If not impacted on rectal examination, patient may still have higher level impaction. Consider abdominal imaging and/or administer milk of molasses enema along with 8 ounces of PO magnesium citrate.</li> <li>• See Bowel Management SmartSet</li> </ul> </li> <li>4. Methylnaltrexone (Relistor®) may be given to patients who meet the following criteria:             <ul style="list-style-type: none"> <li>• Patient experiencing opioid-induced constipation</li> <li>• Patient has not demonstrated an adequate response to other laxative therapy</li> <li>• Patient does not have a known or suspected mechanical gastrointestinal obstruction</li> </ul> </li> </ol>



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## APPENDIX J: Treatment Services

Note: Most treatment facilities require insurance coverage or sufficient money to cover treatment. If patient has insurance, call the customer service number to find a facility in-network to avoid a large out-of-pocket debt.

- Treatment Facilities for Alcohol and Drug Abuse  
Houston, Texas  
(1-800-304-2219)
- Bay Area Recovery Center  
1807 FM 517  
East Dickinson, Texas 77539  
(713) 705-3457
- The Council on Alcohol and Drugs  
Houston, Texas  
[www.councilonrecovery.org](http://www.councilonrecovery.org)
- Clearinghouse for treatment, education, and recovery groups, *etc.*  
303 Jackson Hill St.  
Houston, Texas 77007  
(713) 914-0556, (281) 866-7557
- UT Health Houston Behavioral and Biomedical Science Building  
941 East Rd. First floor  
Houston, Texas 77054  
(713) 500-3784
- Hazelden Betty Ford  
Multiple locations around the country  
1-866-831-5700
- The Treehouse  
Scurry, Texas (South of Dallas)  
(888) 683-1406
- St. Joseph Hospital  
1401 St. Joseph Parkway  
Houston, Texas 77002  
(713) 575-1000 (800) 466-0792
- West Oaks Hospital (Dr. George Santos)  
[www.westoaks.org](http://www.westoaks.org)  
6500 Hornwood  
Houston, Texas 77074
- UT Health Harris County Psychiatric Center (HCPC)  
2800 South MacGregor Way  
Houston, TX 77021  
713-741-5000
- SAMHSA, Substance Abuse and Mental Health Services Administration  
Behavioral Health Treatment Services Locator: [www.findtreatment.samhsa.gov](http://www.findtreatment.samhsa.gov)  
Enter patient's address and zip code on website  
(800) 622 4357
- The Menninger Clinic  
12301 S. Main St.  
Houston, Texas 77035-6207  
(713) 275-5000
- Narcotics Anonymous  
[www.na.org](http://www.na.org)  
Houston area Narcotics Anonymous  
[www.hascona.org](http://www.hascona.org)  
(713) 661-4200

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## SUGGESTED READINGS

American Society of Anesthesiologists Task Force on Acute Pain Management. (2004). Practice guidelines for acute pain management in the perioperative setting: an updated report by the American Society of Anesthesiologists Task Force on Acute Pain Management. *Anesthesiology*, 100(6), 1573-1581.

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## DEVELOPMENT CREDITS

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