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PHARMACIST'S LETTER / PRESCRIBER'S LETTER

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Equianalgesic Dosing of Opioids for Pain Management

Equianalgesic doses contained in this chart are approximate, and should be used only as a guideline. Dosing must be titrated to individual response. There is often incomplete cross-tolerance among these drugs. It is, therefore, recommended to begin with a 50% lower dose than the equianalgesic dose when changing drugs and then titrate to a safe/effective response. Dosing adjustments for renal or hepatic insufficiency, cytochrome P450 drug interactions, genetics, and other conditions or medications that affect drug metabolism, kinetics, or response may also be necessary. Also consider pain control at time of switch. In general, use cautious dosing for elderly or debilitated patients, and patients with renal or hepatic impairment. Some products have specific dosing recommendations for these populations (see footnotes). See our Opioid Conversion Algorithm for instructions on converting from one opioid to another.

A website with an equianalgesic dose calculator is available at http://www.hopweb.org

Drug ^{f,i,k,L}	Equianalgesic Doses (mg) ^{1,3,4}		Approximate Equianalgesic 24 hr Dose (Assumes Around-the- Clock Dosing) ^g		Usual Starting Dose (Opioid-Naïvo Adults) (Doses NOT Equianalgesic)	
	Parenteral	Oral	Parenteral	Oral/Other	Parenteral	Oral/Other
Morphine (immediate-release tablets, oral solution) ^{k,L}	10	30	3-4 mg q 4 h	10 mg q 4 h	2.5 mg q 4 h ⁵⁰	10 mg q 4 h (acute or chronic pain) ^{40,41,51, j} 2-10 mg q 4 h (hospice) ⁵
Controlled-release morphine (e.g., MS Contin, Kadian)	NA	30	NA	30 mg q 12 h (<i>Kadian</i> may be given as 60 mg q 24 h) ^{6,21}	NA	Not for initial dosing. 6,21,29,h, j
Extended-release morphine (Avinza [U.S.], Embeda [with naltrexone, U.S.])	NA	30	NA	60 mg q 24 h (Embeda may be given as 30 mg q 12 h) ¹²	NA	Avinza: 30 mg q 24 h ^{7, j} Embeda: 20 mg q 24 h ^{12, j}

Drug ^{f,i,k,L} Equianalgesic Doses (mg) ^{1,3,4}		Approximate Equianalgesic 24 hr Dose (Assumes Around-the- Clock Dosing) ^g		Usual Starting Dose (Opioid-Naïve Adults) (Doses NOT Equianalgesic)		
	Parenteral	Oral	Parenteral	Oral/Other	Parenteral	Oral/Other
Hydromorphone (Dilaudid)	1.5-2	7.5-8	0.5-0.8 mg q 4 h	2-4 mg q 4 h	See footnotes a,d.	See footnote a.
Controlled-release hydromorphone (<i>Hydromorph Contin</i> [Canada])	NA	7.5 mg	NA	6 mg q 12 h	NA	3 mg q 12 h ^{30, j}
Extended-release hydromorphone (<i>Exalgo</i> , <i>Jurnista</i> [Canada])	NA	See footnote b.	NA	See footnote b.	NA	See footnotes e, h, and j.
Oxycodone (e.g., Roxicodone [U.S.], Oxecta [U.S.], Oxy IR [Canada], also in Percocet, others)	NA	20-30	NA	5-10 mg q 4 h	NA	5-15 mg q 4-6 h (acute or chronic pain) ^{42,43} (Product labeling) 5-10 mg q 8-12 h ¹⁴ or 5 mg q 4-6 h ⁴¹ (chronic noncancer pain) (Guidelines)
Controlled-release oxycodone (<i>OxyContin</i> [U.S.], <i>OxyNeo</i> [Canada])	NA	20-30	NA	20-30 mg q 12 h	NA	10 mg q 12 h ^{9, j}
Oxymorphone (Opana [U.S.])	1	10	0.3-0.4 mg q 4 h	5 mg q 6 h	0.5 mg q 4-6 h ¹⁰	10-20 mg q 4-6 h (acute pain) ^{44,r} 5-10 mg q 4-12 h (chronic noncancer pain) ^{14,41}
Extended-release oxymorphone (<i>Opana ER</i> [U.S.]) ^{c,q}	NA	10	NA	10 mg q 12 h	NA	5 mg q 12 h ^{11, j}
Hydrocodone (in <i>Lortab</i> [U.S.], <i>Vicodin</i> [U.S.], others)	NA	30-45	NA	10-15 mg q 4 h	NA	5-10 mg q 4-6 h ⁴⁵ 5-10 mg q 4-12 h (chronic noncancer pain) ^{14,41}





Drug ^{f,i,k,L}	Equianalge (mg)	esic Doses	Approximate Equianalgesic 24 hr Dose (Assumes Around-the- Clock Dosing) ^g		Usual Starting Dose (Opioid-Naïve Adults) (Doses NOT Equianalgesic)		
	Parenteral	Oral	Parenteral	Oral/Other	Parenteral	Oral/Other	
Codeine ⁿ	100-130	200	30-50 mg q 4 h	60 mg q 4 h	10 mg q 3-4 h ⁵²	15-60 mg q 4 h (mild to moderately severe pain) ⁴⁶ 15-30 mg q 4-12 h (chronic noncancer pain) ^{14,41}	
Controlled-release codeine (Codeine Contin [Canada]) ^{m,n}	NA	200	NA	180 mg q 12 h	NA	50 mg q 12 h ⁴⁹	
Methadone (<i>Dolophine</i> [U.S.], <i>Metadol</i> [Canada]) ^o	Variable	Vari- able	For opioid-tolerant patients only. The conversion ratio of methadone is highly variable depending on factors such as patient tolerance, morphine dose, and length of dosing (short-term versus chronic dosing). Because the analgesic duration of action is shorter than the half-life, toxicity due to drug accumulation can occur with just a few doses. For conversion methods, see http://www.cancer.gov/cancertopics/pdq/supportivecare/pain/HealthPr ofessional/page3. Some experts recommend that only those with substantial experience with its use should prescribe methadone. 39,55				
Meperidine (Demerol)	75	300	Should be used for acute dosing only (short duration of action [2.5 to 3.5 hours]) and neurotoxic metabolite, normeperidine. ¹ Avoid in renal insufficiency and use caution in hepatic impairment and in the elderly (potential for toxicity due to accumulation of normeperidine). ^{1,16-18} Seizures, myoclonus, tremor, confusion, and delirium may occur. ¹				





Drug ^{f,i,k,L}	Equianalgesic Doses (mg) ^{1,3,4}		24 hr Dose (As	Equianalgesic sumes Around- k Dosing) ^g	Usual Starting Dose (Opioid- Naïve Adults) (Doses NOT Equianalgesic)			
	Parenteral	Oral	Parenteral	Oral/Other	Parenteral	Oral/Other		
Fentanyl ^p	0.1	NA	only. Do not corpatch, transmuco (U.S.)], buccal fi sublingual tablet Drugs@FDA; Ca dosing. Or, for U. Products for Brecancer patients:	nvert mcg for mcg psal lozenge [Actiq lm [Onsolis], nasal [Abstral]). See sp anada: Health Can J.S. products only, akthrough Pain. So oral morphine 60 r Round up or down	among fentanyl pr (U.S.)], buccal table I spray [<i>Lazanda</i> (ecific product laberada Drug Product see our <i>PL Chart</i> some experts use the ing total daily dose	Parenteral Oral/Other are for opioid-tolerant patients among fentanyl products (i.e., U.S.)], buccal tablet [Fentora spray [Lazanda (U.S.)], ecific product labeling (U.S.: ada Drug Product Database) for see our PL Chart, Fentanyl ome experts use this conversion in ag total daily dose = 25 mcg/hr based on patient factors and		

NA = not available.

Most of the above oral opioids are available as generics. Exceptions (prices are AWP [U.S.]) include: *Kadian* (\$6.63/30 mg cap), *Avinza* (\$5.45/30 mg cap), *Opana* (\$6.53/10 mg tab), *Opana ER* (\$4.35/10 mg tab), *OxyContin* (\$2.43/10 mg tab), *Embeda* (\$4.98/20 mg cap), and *Exalgo* (\$8.99/8 mg). As a comparison, generic morphine controlled-release = \$1.69/30 mg tab.

- a. Product labeling for **hydromorphone** recommends a starting dose of 0.2 mg to 1 mg IV every two to three hours (Canadian labeling: 2 mg IV every four to six hours) as needed, or 2 mg to 4 mg orally every four to six hours as needed. An even lower oral starting dose (2 mg two or three times daily) has been recommended for chronic pain in opioid-naïve patients. Some institutions use even lower doses of parenteral hydromorphone (e.g., 0.2 mg to 0.5 mg every two hours as needed). One regimen starts opioid-naïve patients at 0.2 mg IV every two hours as needed for mild or moderate pain, with the option in moderate pain to give an extra 0.2 mg after 15 minutes if relief is inadequate after the first 0.2 mg dose. For severe pain, 0.5 mg IV every two hours as needed is used initially. In adults <65 years of age, the 0.5 mg dose can be repeated in 15 minutes if relief is inadequate, for a maximum of 1 mg in two hours.
- b. Per the product labeling, convert to *Exalgo* 12 mg *from* oral codeine 200 mg, hydrocodone 30 mg, morphine 60 mg, oxycodone 30 mg, oxymorphone 20 mg, or transdermal fentanyl 25 mcg/hr. After 50% dose reduction for incomplete cross-tolerance, reduce dose again by 50% for moderate renal impairment, and by 75% for severe renal or moderate hepatic impairment. Not for use in severe hepatic impairment. The *Jurnista* product monograph recommends a 5:1 oral morphine; oral hydromorphone conversion ratio.¹⁹
- c. Per the product labeling, oral **oxymorphone** 10 mg ER is approximately equivalent to hydrocodone 20 mg or oxycodone 20 mg. 11
- d. *Dilaudid* Canadian monograph recommends parenteral starting dose of 2 mg.²⁰ See footnote "a" for additional information and precautions.





- e. No initial dose for *Exalgo*. For opioid-tolerant patients only. Initial *Jurnista* dose (opioid-naïve or <40 mg daily oral morphine equivalents) is 4 to 8 mg q 24 h. In the second of the second
- f. **Tramadol** (e.g., *Ultram*, *Ralivia* [Canada]), potency is about one-tenth that of morphine, similar to codeine. The maximum daily dose of tramadol is 300 mg to 400 mg, depending on the product. 22-28,36,37 See product labeling for dosing in elderly, or in renal or hepatic dysfunction.
- g. Examples of doses seen in clinical practice, taking into account available dosage strengths.
- h. Labeling for some products (MS Contin [U.S.], Kadian, Jurnista [Canada]) recommends beginning treatment with an immediate-release formulation. 6,19,21,29
- i. **Tapentadol** controlled-release (*Nucynta CR*, Canada) and oxycodone controlled-release exhibit comparable pain relief in a dose ratio of 5:1 (tapentadol:oxycodone).³¹ The maximum dose of tapentadol CR is 250 mg twice daily.³¹ No specific dose conversion is given for *Nucynta* (U.S.), *Nucynta IR* (Canada) and *Nucynta ER* (U.S.).^{32,33} Not for use in severe in renal or hepatic dysfunction.^{31-33,38}
- j. Some experts do not recommend for chronic pain in opioid-naïve patients. 14
- k. The initial dose of transdermal **buprenorphine** (*Butrans*) for patients taking less than 30 mg of oral morphine or equivalent per day is a 5 mcg/hr patch applied once weekly (Canada: start with 5 mcg/hr patch in opioid-naïve patients, and 5-10 mcg/hr patch in patients taking up to 80 mg oral morphine equivalents per day). U.S.: When converting from 30 to 80 mg of morphine equivalents daily dose, first taper to 30 mg oral morphine equivalents, then start with the 10 mcg/hr patch.³⁴ The maximum dose is 20 mcg/hr patch once weekly.^{34,47}
- L. Parenteral morphine 10 mg is approximately equal to parenteral **pentazocine** 60 mg, oral pentazocine 180 mg, parenteral **butorphanol** 2 mg, and parenteral **nalbuphine** 10 mg. ⁴⁹ For buprenorphine transdermal patch (*Butrans*), see footnote "k." The analgesic efficacy of these drugs is limited by a dose ceiling. Furthermore, the mixed agonists-antagonists (i.e., pentazocine, butorphanol, nalbuphine) are contraindicated for use in patients receiving an opioid agonist because they can precipitate withdrawal and increase pain. They also pose a risk of psychotomimetic effects. ¹
- m. Reduce dose by 25% when switching from oral codeine phosphate to account for phosphate content of tablet.⁴⁹
- n. Analgesic efficacy limited by a dose ceiling. 46,49
- o. Relatively safe choice in renal or liver insufficiency. 54,55
- p. Relatively safe choice in renal or liver insufficiency.⁵⁵ Clearance reduced by uremia.⁵⁴ Do not start with patch in renal or liver failure.⁵⁴ Watch for delayed toxicity.^{54,55}
- q. Opana ER has received a notice of compliance (June 2012) by Health Canada. At time of publication, it is not yet available on the Canadian market.
- r. Start with an oral dose of 5 mg q 4-6 h for opioid-naïve elderly or opioid-naïve patients with renal or liver impairment.⁴⁴

Users of this PL Detail-Document are cautioned to use their own professional judgment and consult any other necessary or appropriate sources prior to making clinical judgments based on the content of this document. Our editors have researched the information with input from experts, government agencies, and national organizations. Information and Internet links in this article were current as of the date of publication.





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