PRESCRIBING A SAFER DOSAGE

HIGHER DOSAGE, HIGHER RISK.

Higher dosages of opioids are associated with higher risk of overdose and death—even relatively low dosages (20-50 morphine milligram equivalents (MME) per day) increase risk.

Higher dosages haven't been shown to reduce pain over the long term. One randomized trial found no difference in pain or function between a more liberal opioid dose escalation strategy (with average final dosage 52 MME) and maintenance of current dosage (average final dosage 40 MME).



THE IMPORTANCE OF CALCULATING DAILY DOSAGE

Patients prescribed higher opioid dosages are at higher risk of overdose death. In a national sample of Veterans Health Administration (VHA) patients with chronic pain receiving opioids from 2004–2009, **patients who died** of opioid

overdose were prescribed an average of 98 MME/day, while other patients were prescribed an average of 48 MME/day.

Dosages at or above 50 MME/day increase risks for overdose by at least 2x the risk at <20 MME/day. Calculating the total daily dose of opioids helps identify patients who may benefit from closer monitoring, reduction or tapering of opioids, prescribing of naloxone, or other measures to reduce risk of overdose.

HOW MUCH IS 50 OR 90 MME/DAY FOR COMMONLY PRESCRIBED OPIOIDS?

50 MME/day:

- 50 mg of hydrocodone (10 tablets of hydrocodone/acetaminophen 5/300)
- 33 mg of oxycodone (~2 tablets of oxycodone sustained-release 15 mg)
- 12 mg of methadone (<3 tablets of methadone 5 mg)

90 MME/day:

- 90 mg of hydrocodone (9 tablets of hydrocodone/acetaminophen 10/325)
- 60 mg of oxycodone (~2 tablets of oxycodone sustained-release 30 mg)
- ~20 mg of methadone (4 tablets of methadone 5 mg)

HOW SHOULD THE TOTAL DAILY DOSE OF OPIOIDS BE CALCULATED?

- DETERMINE the total daily amount of each opioid the patient takes.
- CONVERT each to MMEs—multiply the dose for each opioid by the conversion factor. (see table)
- 3. ADD them together.

CAUTION:

 These dose conversions are estimated and cannot account for all individual differences in genetics and pharmacokinetics

USE EXTRA CAUTION:

- Methadone: the conversion factor increases at higher doses
- Fentanyl: dosed in mcg/ hr instead of mg/day, and absorption is affected by heat and other factors

OPIOID (doses in mg/day except where noted)	CONVERSION FACTOR
Codeine	0.15
Fentanyl transder- mal (in mcg/hr)	2.4
Hydrocodone	1
Hydromorphone	4
Methadone	
1-20 mg/day	4
21-40 mg/day	8
41-60 mg/day	10
≥ 61-80 mg/day	12
Morphine	1
Oxycodone	1.5
Oxymorphone	3

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HOW SHOULD PROVIDERS USE THE TOTAL DAILY OPIOID DOSE IN CLINICAL PRACTICE?

- Use caution when prescribing opioids at any dosage and prescribe the lowest effective dose.
- Use extra precautions when increasing to ≥50 MME per day* such as:
 - Monitor and assess pain and function more frequently.
 - Discuss reducing dose or tapering and discontinuing opioids if benefits do not outweigh harms.
 - Consider offering naloxone.
- Avoid or carefully justify increasing dosage to ≥90 MME/ day.*
- Specify parameters for PRN. Ex. Must last 30 days.
- * These dosage thresholds are based on overdose risk when opioids are prescribed for pain and should not guide dosing of medication-assisted treatment for opioid use disorder.

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