

## **Co-prescribing Naloxone with Opioids**

Drug overdose is one of the leading causes of accidental death in the United States. Preventable drug overdose deaths increased from 98,268 in 2021 to 99,592 in 2022, a rise of 1.3%, with 77,603 (78%) of the 2022 deaths involving opioids.<sup>1</sup> Given rising opioid overdose death rates, the U.S. Surgeon General issued a national advisory in 2018 urging more Americans to carry naloxone.<sup>2</sup> As of January 2019, Assembly Bill 2760 requires California prescribers to provide education and offer a naloxone prescription to high-risk patients.<sup>3</sup>

## Why should naloxone be offered to patients?

Naloxone is a highly competitive mu-opioid receptor antagonist that displaces the opioid and temporarily reverses its fatal effects, such as respiratory depression.<sup>2,4</sup> Overdose education and naloxone distribution has been shown to reduce opioid overdose death rates by up to 46%.<sup>5</sup> A 2019 study showed an association between pharmacists' authority to furnish naloxone and significant reductions in fatal overdoses among Medicaid beneficiaries.<sup>6</sup> Despite a substantial increase in naloxone dispensing from 2019 to 2023, due to state-level standing orders and guideline updates, nationwide dispensing rates remain low.<sup>7</sup>

## Who should receive naloxone?

The 2022 Centers for Disease Control and Prevention Guideline for Prescribing Opioids for Chronic Pain recommends co-prescribing naloxone when one or more of the following risk factors for overdose are present:<sup>8</sup>

- Higher opioid dosages (greater than or equal to 50 MME per day)
- Concurrent benzodiazepine use (regardless of opioid dose)
- Respiratory conditions such as chronic obstructive pulmonary disease (COPD) or obstructive sleep apnea (regardless of opioid dose)
- History of overdose or substance use disorder
- Patients at risk for returning to a high dose for which opioid tolerance is lost (e.g., patients undergoing tapering)

Additionally, California Civil Code §1714.22 allows for licensed health care providers to prescribe naloxone to a family member, friend or another person in a position to assist a person at risk of an overdose.<sup>9</sup>

Consider the following opioid reversal agents for your patients at high risk for opioid overdose:

Formulations	Recommended Dosing <sup>4</sup>
naloxone 0.4 mg/mL carpuject	Inject 0.4 to 2 mg intramuscularly or subcutaneously.
naloxone 0.4 mg/mL, 4 mg/10 mL vial	May repeat every two to three minutes until medical assistance becomes
naloxone 0.4 mg/mL, 2 mg/2 mL syringe	available.
naloxone (Narcan) 4 mg nasal spray	Place one spray as a single dose in one nostril. May repeat every two to three minutes in alternating nostrils until medical
naloxone (Kloxxado) 8 mg nasal spray	assistance becomes available.

## **References**

- 1. National Safety Council Injury Facts. Drug Overdoses. https://injuryfacts.nsc.org/home-and-community/safety-topics/drugoverdoses/data-details. Accessed August 21, 2024.
- Office of the Surgeon General. U.S. Surgeon General's Advisory on Naloxone and Opioid Overdose. US Department of Health & Human Services. https://www.hhs.gov/surgeongeneral/reports-and-publications/addiction-and-substance-misuse/advisory-on-naloxone/index.html. Accessed August 21, 2024.
- 3. California Legislative Information. AB-2760 Prescription drugs: prescribers: naloxone hydrochloride and other FDA-approved drugs.
- https://leginfo.legislature.ca.gov/faces/billTextClient.xhtml?bill\_id=201720180AB2760. Published September 10, 2018. Accessed August 21, 2024.
  Naloxone hydrochloride. Lexi-Drugs. UpToDate Lexidrug. UpToDate Inc. http://online.lexi.com. Accessed August 21, 2024.
- Walley AY, Xuan Z, Hackman HH, et al. Opioid overdose rates and implementation of overdose education and nasal naloxone distribution in Massachusetts: interrupted time series analysis. BMJ. 2013;346: f174.
- 6. Abouk R, Pacula RL, Powell D. Association between State laws facilitating pharmacy distribution of naloxone and risk of fatal overdose. JAMA Intern Med. Published May 6, 2019. doi:10.1001/jamainternmed.2019.0272.
- 7. Rikard SM, Strahan AE, Schleber LZ, et al. Trends in naloxone dispensing from retail pharmacies in the US. JAMA. Published July 9, 2024. doi: 10.1001/jama.2024.8323.
- Dowell D, Ragan KR, Jones CM, et al CDC Clinical Practice Guideline for Prescribing Opioids for Pain United States, 2022. MMWR Recomm Rep 2022;71(No. RR-3):1–95. DOI: http://dx.doi.org/10.15585/mmwr.rr7103a1.
- 9. An act to amend Section 1714.22 of the Civil Code, relating to drug overdose treatment. Cal. Civil Code §1714.22 (2014).