Pain Management and Opioids:

Balancing Risks and Benefits



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DISCLOSURE:

Dr. Carron declares that she has no relevant financial relationships with ineligible companies to disclose.



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https://www.opioidanalgesicrems.com/Resources/Docs/List_of_RPC_Companies.pdf for a listing of REMS Program Companies. This activity is intended to be fully compliant with the Opioid Analgesic REMS education requirements issued by the U.S. Food and Drug Administration. For more information about the Opioid Analgesics REMS visit https://opioidanalgesicrems.com/RpcUI/products.u.



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BY THE END OF THIS SESSION YOU WILL BE ABLE TO:

- Describe the pathophysiology of pain as it relates to the concepts of pain management.
- Accurately assess patients in pain.
- Develop a safe and effective pain treatment plan.
- Identify evidence-based non-opioid options for the treatment of pain.
- Identify the risks and benefits of opioid therapy.
- Manage ongoing opioid therapy.
- Recognize behaviors that may be associated with opioid use disorder.



WHY ARE WE HERE?



CO*RE STATEMENT

Historical over-prescribing, a massive and sustained exposure to opioids, and a gap in treatment availability have fueled the opioid overdose epidemic in the United States.

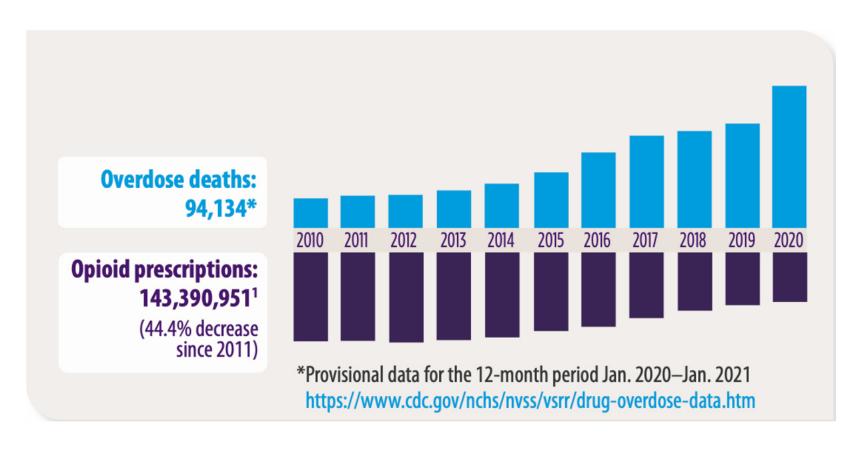
When prescribed well, and used as prescribed, opioids can be valuable tools for effective pain management.

Unintended consequences may occur from both under-prescribing (unmanaged pain) and over-prescribing (injudicious use of opioids).

This course does not advocate for or against the use of opioids. We intend to help health-care providers manage pain without putting vulnerable patients at risk for misuse or opioid use disorder. The goal is to keep our patients, our communities, and ourselves SAFE.



REDUCTIONS IN OPIOID PRESCRIBING HAVE NOT LED TO REDUCTIONS IN OVERDOSE DEATHS



SOURCE: https://www.ama-assn.org/delivering-care/overdose-epidemic/physicians-progress-toward-ending-nation-s-drug-overdose-epidemic



TYPES OF OPIOIDS



NATURALLY OCCURRING OPIATES	SEMI-SYNTHETIC OPIOIDS	SYNTHETIC OPIOIDS
Codeine	Buprenorphine	Alfentanil
Morphine	Hydrocodone	Fentanyl
	Hydromorphone	Methadone
	Oxycodone	Remifentanil
	Oxymorphone	Tapentadol
		Tramadol

AGONISTS	PARTIAL AGONISTS	ANTAGONISTS
Codeine	Buprenorphine	Naloxone
Methadone	Nalbuphine	Naltrexone
Morphine		
Oxycodone		



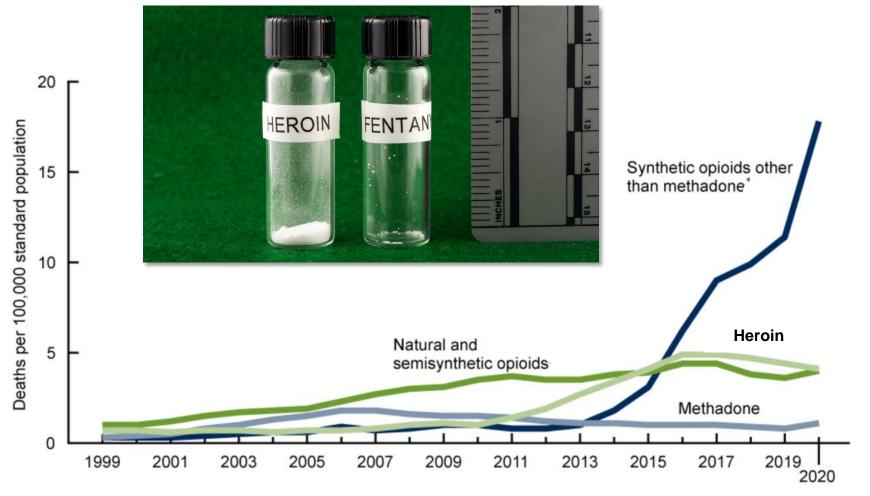
DEA SCHEDULED DRUGS

SCHEDULE	DESCRIPTION	EXAMPLES
I	High potential for abuse; no currently accepted medical use	Cannabis, ecstasy, heroin, LSD, peyote
II	High potential for abuse, which may lead to severe psychological or physical dependence	Codeine, fentanyl, hydrocodone combination products, hydromorphone, meperidine, methadone, morphine, opium, oxycodone,
III	Potential for abuse, which may lead to moderate or low physical dependence or high psychological dependence	Products containing ≤ 90 mg codeine per dose, buprenorphine, benzphetamine, phendimetrazine, ketamine, anabolic steroids
IV	"Low potential" for abuse	Alprazolam, carisoprodol, clonazepam, clorazepate, diazepam, lorazepam, midazolam, temazepam, tramadol
V	Low potential for abuse	Cough preparations containing ≤ 200 mg codeine/100 ml

Complete list of products covered under the Opioid Analgesic REMS available at: https://www.opioidanalgesicrems.com/products.html



OPIOID OVERDOSE DEATHS BY TYPE OF OPIOID



SOURCE: https://www.cdc.gov/nchs/images/databriefs/401-450/db428-fig4.png, https://www.cdc.gov/nchs/products/databriefs/db428.htm
PHOTO SOURCE: New Hampshire State Police Forensic Lab



FENTANYL AND FENTANYL ANALOGUES

ON THE PLUS SIDE:

- Short-acting and easily metabolized
- Useful for controlling acute pain



- Cheap and easy to manufacture
- Easy to store, easy to smoke, easy to conceal
- Unpredictable dosing with non-prescribed fentanyl
- Two causes of fentanyl OD death: opioid-induced respiratory depression and rigid chest wall syndrome; higher or repeated doses of naloxone are required to reverse a fentanyl overdose.





RISKS VERSUS BENEFITS OF PRESCRIBED OPIOIDS

POTENTIAL RISKS

- Life-threatening respiratory depression/overdose
- Development of SUD/OUD
- Diversion
- Inadvertent exposure to family and pets
- Interactions with other meds and substances
- Neonatal abstinence syndrome
- Physiologic dependence and withdrawal

POTENTIAL BENEFITS

- Analgesia
- Option for patients with contraindications for non-opioid analgesics
- Relieves suffering
- May improve function and quality of life



Opioid Prescribing Rates & Overdose Deaths



2018

2019

2020

WV

69.3

59.4

53.7

Office of National Drug Control Policy (ONDCP)
Non-Fatal Opioid Overdose Tracker

Opioid Overdose Death Rate Per 100,000 Population

WV 2021

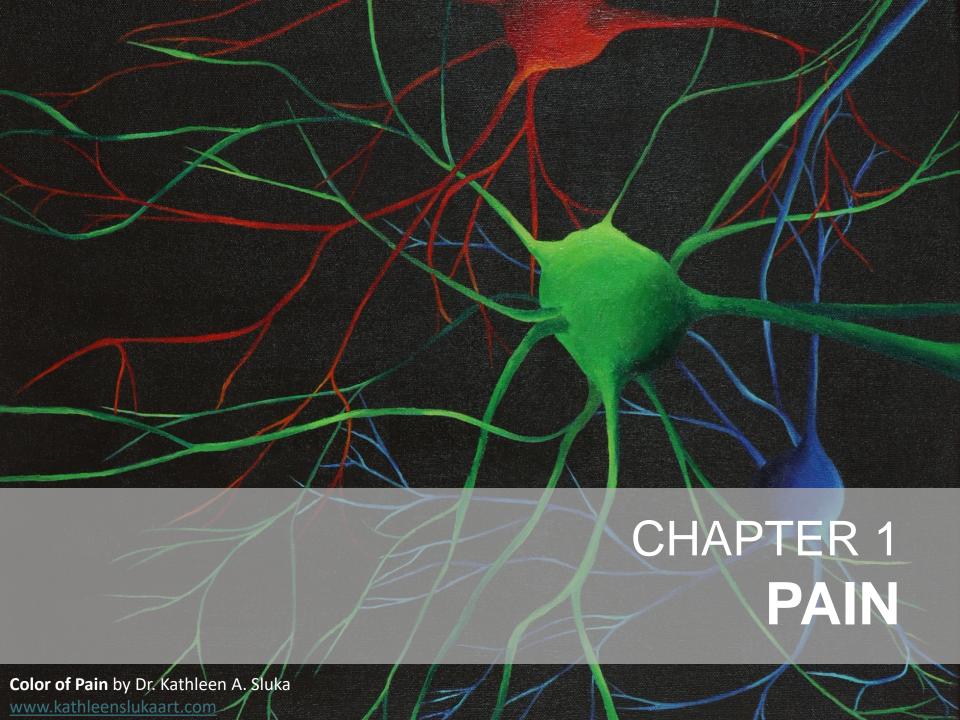
77.2

US 2021

24.7

https://www.cdc.gov/drugoverdose/rxrate-maps/ https://www.kff.org/state-category/health-status/opioids/





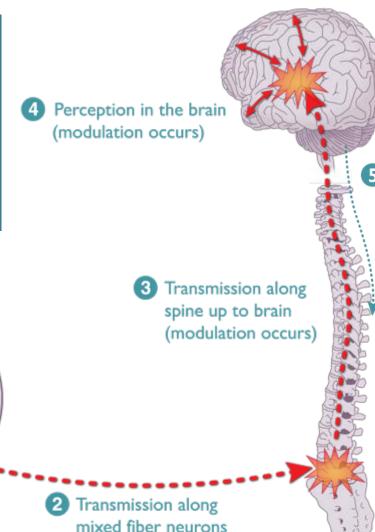
THE NEUROMECHANISMS OF PAIN

Peripheral Pain Modulators:

- Histamines
- Prostaglandins
- Cytokines
- Bradykinin
- Substance P

1 Injury

Others



(modulation occurs)

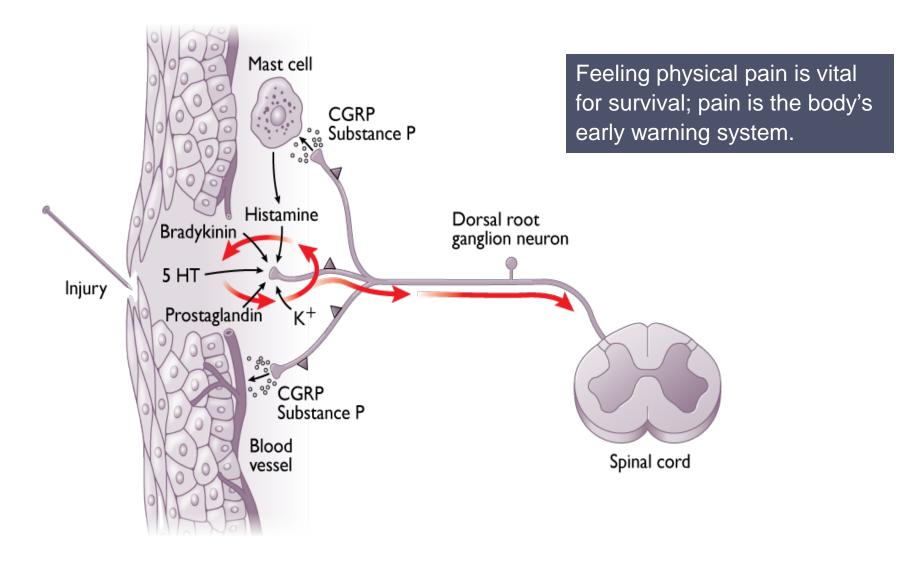
Descending pathway (down regulation)

Descending Neurotransmitters:

- Serotonin
- Norepinephrine
- Endogenous opiates
- Others

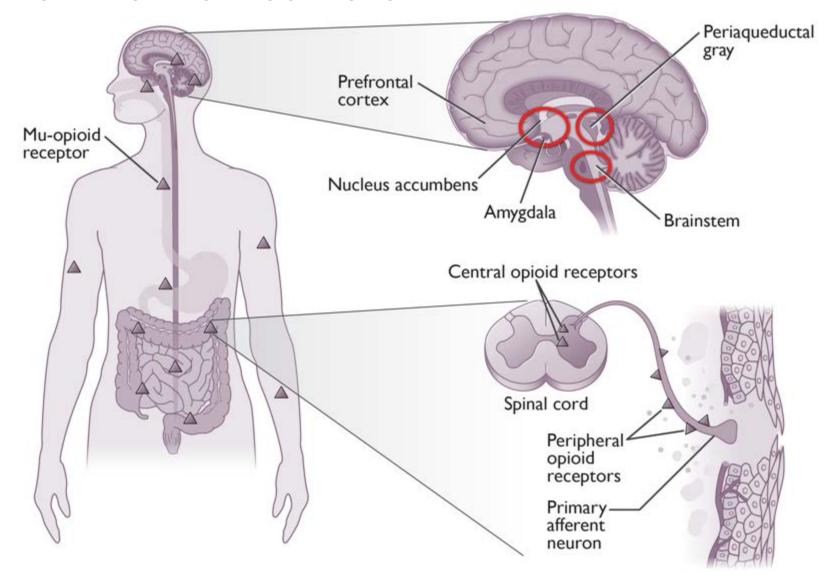


MEDIATORS OF PERIPHERAL NOCICEPTION





OPIOID RECEPTOR LOCATIONS



PAIN

"An unpleasant sensory and emotional experience associated with, or resembling that associated with, actual or potential tissue damage."

—IASP (July 2020)

ACUTE	CHRONIC
 Acute pain duration of < 1 month Sudden onset, self-limiting Ideally resolves with healing Triggered by tissue damage and inflammation Has protective value Inflammatory mediation Subacute, pain that continues for 1-3 months, can become chronic 	 Lasting 3 months or longer Generally steady-state or worsening Persists beyond normal healing period Serves no value Peripheral and central sensitization



TYPES OF PAIN

NOCICEPTIVE / INFLAMMATORY



Pain in response to an injury or stimuli; *typically* acute



Post-operative pain, sports injuries, arthritis, sickle cell disease, mechanical low back pain

NOCIPLASTIC



Pain that arises from altered nociceptive function; typically chronic



Fibromyalgia, irritable bowel syndrome, nonspecific low back pain

NEUROPATHIC



Pain that develops when the nervous system is damaged; chronic



Post-herpetic neuralgia, trigeminal neuralgia, distal polyneuropathy, CRPS, neuropathic low back pain

MIXED TYPES (NOCICEPTIVE / NEUROPATHIC)

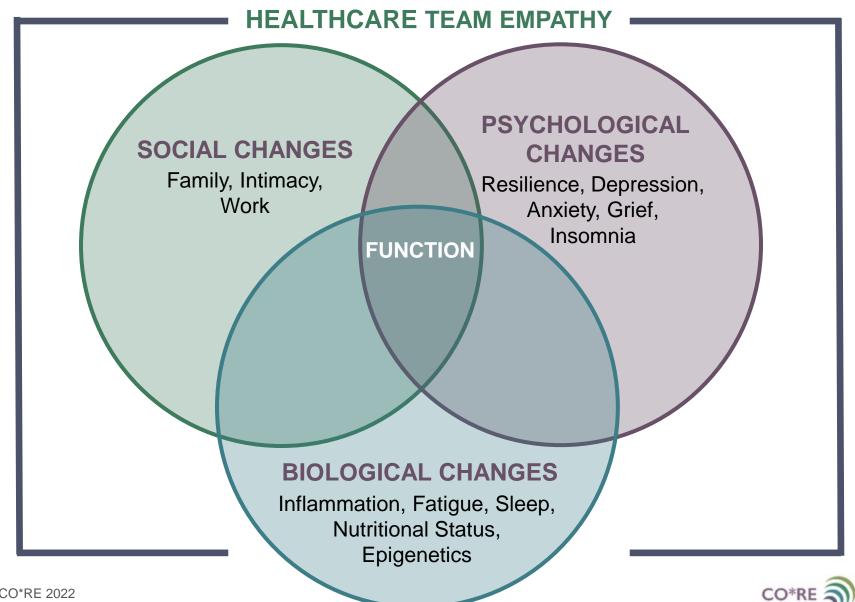


Primary injury and secondary effects

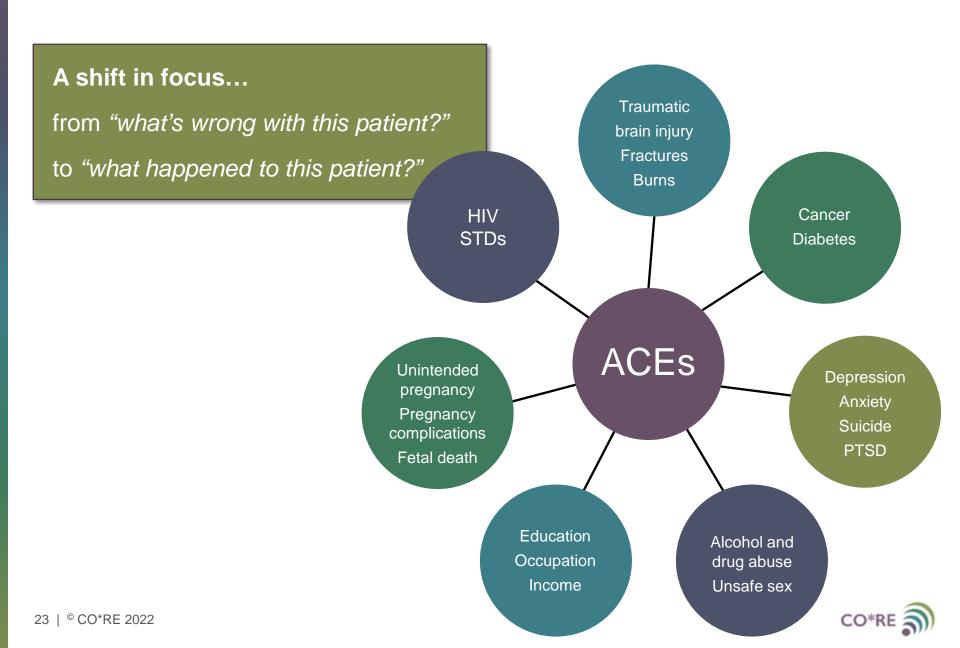




THE EXPERIENCE OF PAIN: A BIOPSYCHOSOCIAL MODEL



ADVERSE CHILDHOOD EXPERIENCES (ACEs)





CHAPTER 2 TERMINOLOGY

WORDS MATTER: LANGUAGE CHOICE CAN REDUCE STIGMA

"If you want to care for something, you call it a flower; if you want to kill something, you call it a weed." —DON COYHIS

COMMONLY USED TERM	PREFERRED TERM
Addiction	Substance use disorder (SUD) or opioid use disorder (OUD) [from the <i>DSM-5-TR</i> ®]
Drug-seeking, aberrant/problematic behavior	Using medication not as prescribed
Addict/user	Person with a substance use disorder (SUD) or an opioid use disorder (OUD)
Dirty urine/failing a drug test	Testing positive on a urine drug screen
Abuse or habit	Misuse or "use other than prescribed"

SOURCE: https://nida.nih.gov/research-topics/addiction-science/words-matter-preferred-language-talking-about-addiction

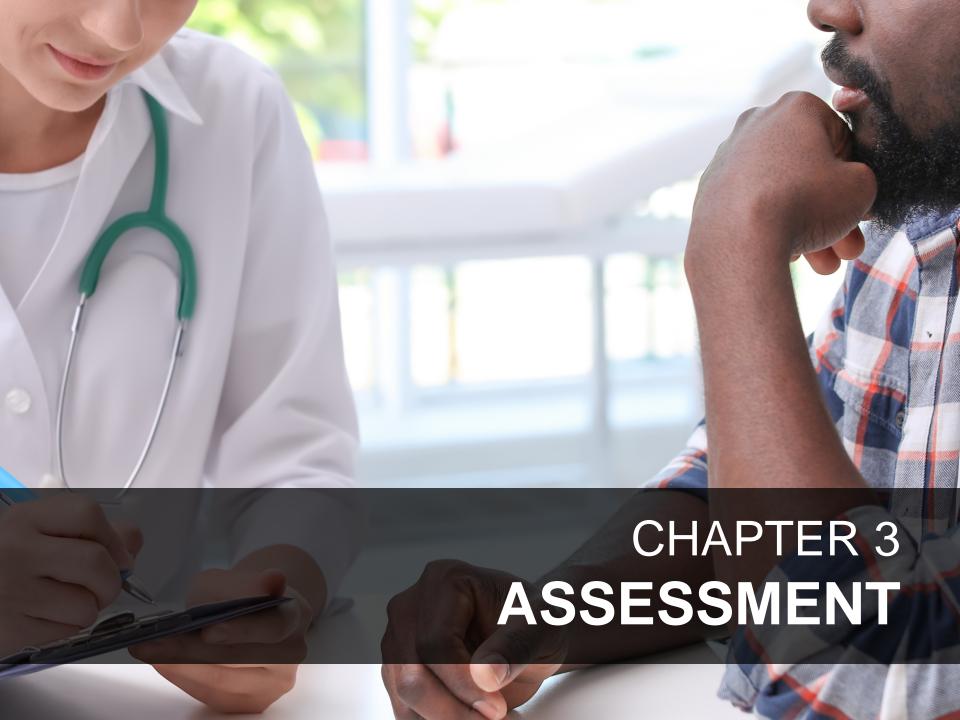


WORDS MATTER: DEFINITIONS

Misuse	Use of a medication in a way other than the way it is prescribed	
Tolerance	Increased dosage needed to produce a specific effect	
Dependence	State in which an organism only functions normally in the presence of a substance	
Diversion	Transfer of a legally controlled substance, prescribed to one person, to another person for illicit (forbidden by law) use	
Withdrawal	Occurrence of uncomfortable symptoms or physiological changes caused by an abrupt discontinuation or dosage decrease of a pharmacologic agent	
MOUD	Medication for Opioid Use Disorder, an approach to treating Opioid Use Disorder that combines FDA-approved medication with counseling and behavioral therapies	
мме	Morphine milligram equivalents; a standard opioid dose value based on morphine and its potency; allows for ease of comparison and risk evaluations	
Chronic non- cancer pain (CNCP)	Any painful condition that persists for ≥ 3 months, or past the time of normal tissue healing, that is not associated with a cancer diagnosis	

SOURCE: NIDA, https://nida.nih.gov/research-topics/addiction-science/words-matter-preferred-language-talking-about-addiction





PAIN ASSESSMENT

DESCRIPTION OF PAIN







Intensity



Quality



duration



Variations/ patterns/rhythms

WHAT RELIEVES THE PAIN?

WHAT CAUSES OR INCREASES THE PAIN?

EFFECTS OF PAIN ON PHYSICAL, EMOTIONAL AND PSYCHOSOCIAL FUNCTION

PATIENT'S CURRENT LEVEL OF PAIN AND FUNCTION

SOURCE: Hogans, B., Barreveld, A. (Eds.). Pain Care Essentials, New York, NY: Oxford University Press. 2020.



MEDICAL AND TREATMENT HISTORY

RELEVANT ILLNESSES



PAST AND CURRENT OPIOID USE

- Query your state's Prescription Drug Monitoring Program
 (PDMP) to confirm patient report
- Contact past providers and obtain prior medical records
- For opioids currently prescribed, note the opioid, dose, regimen, and duration
- Determine whether the patient is opioid-tolerant

NONPHARMACOLOGIC STRATEGIES AND EFFECTIVENESS

PHARMACOLOGIC STRATEGIES AND EFFECTIVENESS

BARRIERS TO PREVIOUS TREATMENT STRATEGIES



PRESCRIPTION DRUG MONITORING PROGRAMS (PDMPs)

PDMP DATABASES	BENEFITS
 Reports on opioid prescriptions filled by patient Nearly all are available online 24/7 54 operational PDMPs in the U.S. In some states, prescribers are required to access; know your state laws 	 Lower rates of prescription opioid-related hospitalization and ED visits Reduction in "doctor shopping" Reduction in prescribing high doses and over-prescribing Identify drugs that increase overdose risk when taken together (such as benzodiazepines, gabapentinoids, opioids, and other sedatives)

Limitations: Often under-used, can be time consuming, may not have access to bordering state data, lack of intuitive format, privacy issues, no national PDMP connection

Multiple prescriptions from different providers is most predictive of opioid misuse.



PDMP: Prescription Drug Monitoring Program

Ge	ne	ral

- CSMP (Controlled Substance Monitoring Program)
 https://www.csappwv.com/Account/Login.aspx?ReturnUrl=%2f
- Administered by the Board of Pharmacy
- Schedule II-V are monitored
- Dispensers and prescribers are required to register and input data
- Before prescribing, there is an obligation to review under certain circumstances
- Prescribers can authorize a registered delegate

Reporting

- Must be entered into PDMP within 24 hours after dispensing
- Unsolicited reports/alerts are sent to prescribers, dispensers, law enforcement and licensing boards
- West Virginia does share data with other states' PDMP
- Out-of-state pharmacies are required to report to the patient's home state
- Patient will be notified if their record has been accessed

https://namsdl.org/doc-library/?fwp_document_type=map January 2019 http://www.pdmpassist.org/content/pdmp-maps-and-tables January 2023



Prescribing Limits, Status & Education Requirements

Seven-day (short-term pain), four-day (emergency room prescriptions), three-day (prescribed by dentists or optometrists)

	Physician	PA	Advanced Practice Nurse
Prescriber Status	Licensed	Schedule II-V	Schedule II-V
Education Requirements	3 hrs./2 yrs.	3 hrs./2 yrs.	3 hr. x 1, and then 1 hour /1yr

^{*}Special rules apply for Schedule II

http://www.fsmb.org/siteassets/advocacy/key-issues/continuing-medical-education-by-state.pdf, January 2023 Opioid prescription limits and policies by state – Ballotpedia, April 4, 2022 www.netce.com/ce-requirements/

OBTAIN A COMPLETE SOCIAL AND PSYCHOLOGICAL HISTORY

SOCIAL HISTORY

Employment, cultural background, social network, relationship history, legal history, and other behavioral patterns

PSYCHOLOGICAL HISTORY

Screen for:

- Mental health diagnoses, depression, anxiety,
 PTSD, current treatments
- Alcohol, tobacco, and other drug use
- History of Adverse Childhood Experiences (ACES)
- Family history of substance use disorder and psychiatric disorders

Depression and anxiety can be predictors of chronic pain





PHYSICAL EXAM AND ASSESSMENT

Seek objective data

Conduct physical exam and evaluate for pain

Order diagnostic or confirmatory tests

General: vital signs, appearance, and pain behaviors

Neurologic exam

Musculoskeletal exam

- Inspection
- Gait and posture
- Range of motion
- Palpation
- Percussion
- Auscultation
- Provocative maneuvers

Cutaneous or trophic findings

SOURCE: Hogans, B., Barreveld, A. (Eds.). Pain Care Essentials, New York, NY: Oxford University Press. 2020.



PAIN ASSESSMENT TOOLBOX

http://core-rems.org/opioid-education/tools/

Pain Assessment Tools

BPI or 5 A's

Functional Assessment

SF-36, PPS, Geriatric Assessment

Pain intensity, Enjoyment of life, General activity

PEG

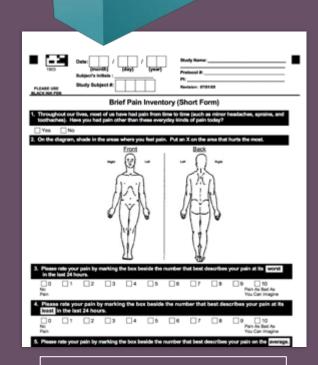
Adverse Childhood Experience Questionnaire

ACE

Assessment in Patients Unable to Self-Report

Hierarchy of Pain Assessment

PAINAD



Brief Pain Inventory (BPI)

Psychological Measurement Tools (PHQ-9, GAD-7, etc.)



ASSESSMENT IS NOT A ONE-TIME OCCURRENCE

Assessment of a patient's response to pain treatment is a continual process:

- Routinely check the PDMP
- Check in with your patients
- Reassess to identify the underlying source of pain
- Investigate comorbid conditions that may arise
- Ask if patient is willing to engage with other modalities
- Modify plans as needed

















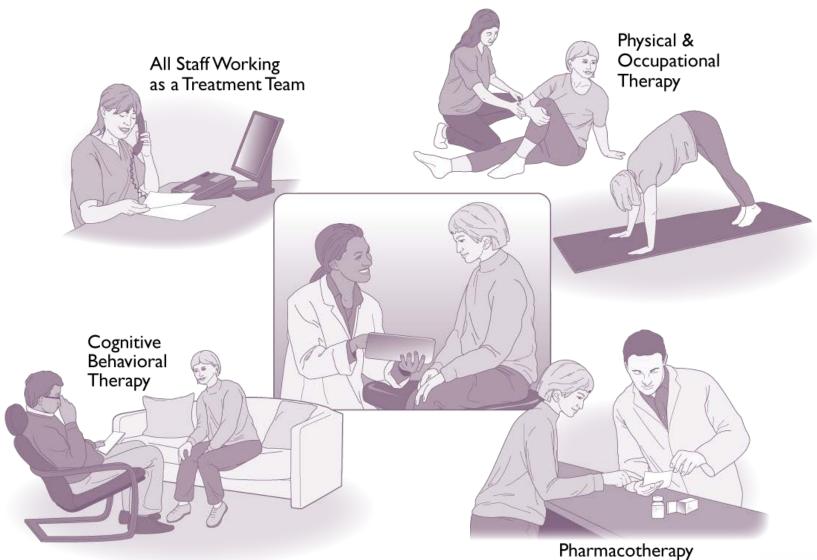




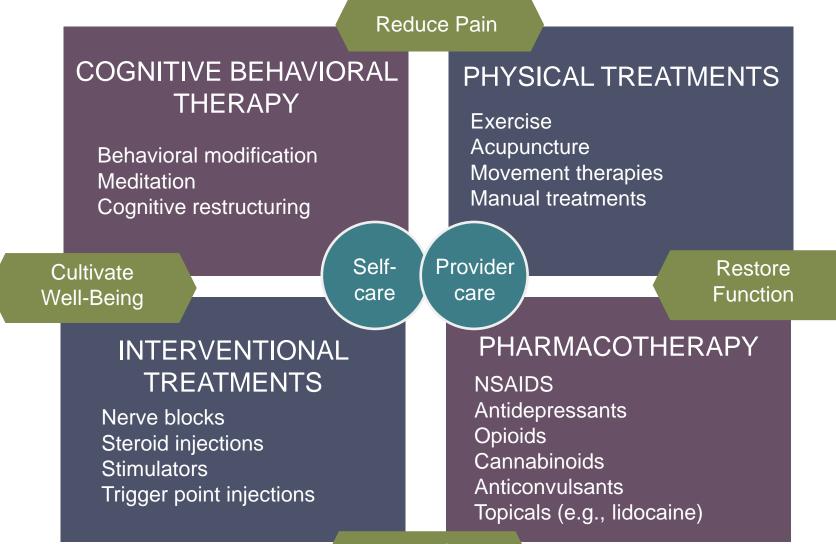
HOW IS PAIN MANAGED?



COMPONENTS OF A MULTIMODAL TREATMENT PLAN FOR PAIN



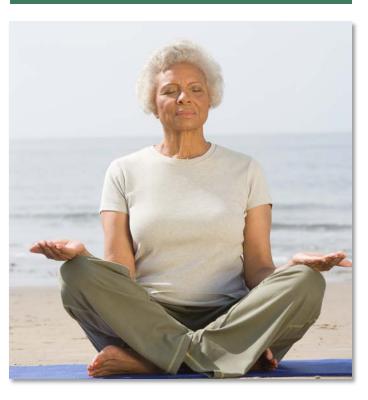
PAIN MANAGEMENT GOALS AND TREATMENT OPTIONS: A MULTIMODAL APPROACH





EVIDENCE-BASED NONPHARMACOLOGIC TREATMENTS

What is appropriate for your patient?



- Tai Chi
- Yoga
- CBT and ACT
- Acupuncture
- PT/OT/aquatic
- Mindfulness meditation
- OMT
- Massage therapy
- Chiropractic
- Neuromodulation or surgical approaches (in some situations)

CBT = cognitive behavioral therapy; ACT = acceptance commitment therapy; OMT = osteopathic manipulative therapy



PHARMACOLOGIC TREATMENTS BY TYPE OF PAIN

CONTINUE EFFECTIVE NONPHARMACOLOGIC OPTIONS

NOCICEPTIVE / INFLAMMATORY



IR opioids
Nerve blocks
NSAIDs
Topicals and patches

NOCIPLASTIC



Anticholinergic
Anticonvulsants
TCAs and SNRIs
Other serotonin agents



NEUROPATHIC



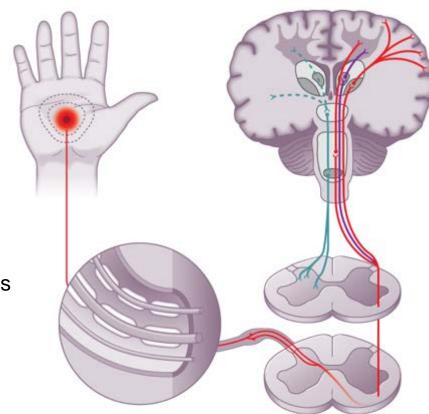
Anticonvulsants
IR and ER/LA opioids
Gabapentinoids
Nerve blocks
TCAs and SNRIs
Transdermal opioids



POTENTIAL SITES OF ACTION FOR ANALGESIC AGENTS

Peripherally Mediated Pain:

- Acetaminophen
- NSAIDs
- Opioids
- Topical anesthetics



Centrally Mediated Pain:

- Alpha-2 agonists
- Anticonvulsants
- Ca⁺ channel antagonists
- NMDA RAs
- Opioids
- TCA/SNRI antidepressants

Most commonly, pain conditions are a combination of peripherally and centrally mediated processes



DRUG CHARACTERISTICS TO CONSIDER BEFORE PRESCRIBING

Mechanism of Route of Strength Dosing interval administration action Key instructions Product-specific Specific drug **Formulation** (indications, uses, safety concerns interactions contraindications) Relative Specific information about product Use in opioid potency to conversions, if available tolerant patients morphine (MME)

Opioid product information available at https://opioidanalgesicrems.com/products.html.



2022 CDC GUIDELINE

- Clinician recommendations for patients aged ≥18 years
- Summary of current research
- Flexible; encourages patientcentered decision making
- Emphasizes the importance of the individual & clinical judgement
- This is a clinical tool, not a law, regulation or policy

CDC Clinical Practice Guideline for Prescribing Opioids for Pain —

*Dissison of Overnous Presention, National Center for Injury Presention and Control, CDG:

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**Office of the Director, National Center and Orogan Health & Science University Purland, Orogan

**Specific Northwest Evidence-based Practice Center and Orogan Health & Science University Purland, Orogan

This guideline provides recommendations for clinicians providing pain care, including those prescribing opioids, for outpatients Inst guideune provides recommendations for curricians providing pain care, including those prescribing opioids, for outpatients aged 218 years. It updates the CDC Guideline for Prescribing Opioids for Chronic Pain — United States, 2016 (MMWR). agea 2.10 years. It updates the CLO Guideline for Fresching Opioids for Caronic rain — Onited states, 2010 (Ann MR Recomm Rep 2016;65] No. RR-I]:1–49) and includes recommendations for managing acute (duration of c1 month), subacute Recomm Rep 2010;05][No. KK-1];1—92] and includes recommendations for managing active (auration of <1 months), students (duration of <3 months) pain. The recommendations do not apply to pain related to duration of 1–3 months), and chronic (duration of <3 months) pain. The recommendations do not apply to pain related to the recommendation of the recomm (duration of 1-5 months), and chronic (duration of >5 months) pain. The recommendations do not apply to pain retained to sickle cell disease or cancer or to patients receiving palliative or end-of-life care. The guideline addresses the following four areas: siekte cell ausease or cancer or to patients receiving paulative or ena-03-119 care. Loe guiaeune auareuso ire jouwwing jour areas.

1) determining whether or not to initiate opioids for pain. 2) selecting opioids and determining opioid dosages, 3) deciding 1) determining whether or not to initiate opioids for pain, 2) selecting opioids and determining opioid dosages, 3) deciding duration of initial opioid prescription and conducting follow-up, and 4) assessing risk and addressing potential harms of opioid the conduction of the conduc users that opious prescription and conducting jouone-up, and 1) assessing risk and adaressing potential narms of opioid issess. CDC developed the guideline using the Grading of Recommendations Assessment, Development, and Evaluation (GRADE) framework. Recommendations are based on systematic reviews of the scientific evidence and reflect considerations of benefits and framework. recommensations are oasea on systematic reviews of the scientific evidence and reflect considerations of oenefits and harms, patient and clinician values and preferences, and resource allocation. CDC obtained input from the Board of Scientific narms, passeus associational vasues and preperences, and resource association. CLD obtained input from the Board of occentific Counselors of the National Center for Injury Prevention and Control (a federally chartered advisory committee), the public, and Councesors of the vatuonal Center for injury revention and Comrol se feetauty character authorized monitoring the proper reviewers. CDC recommends that persons with pain receive appropriate pain treatment, with careful consideration of the peer reviewers. LLA recommenas inat persons with pain receive appropriate pain treatment, with careful consideration of the benefits and risks of all treatment options in the context of the patient's circumstances. Recommendations should not be applied to the patient's circumstances. venessis and rise, of an ireasment options in the context of the patients circumstances, recommendations should not be applied as inflexible numbered of care across patient populations. This clinical practice guideline is intended to improve communication anno of the across pattern populations. 1715 thrittal practice guintern is intended to improve communication

and patients about the benefits and risks of pain treatments, including opioid therapy; improve the effectiveness stituate pain; improve function and quality of life for patients with pain; and reduce risks associated id use disorder, overdose, and death.



Morbidity and Mortality Weekly Rep

November 4, 20

CDC Clinical Practice Guideline for Prescribing Opioids for Pain — United States, 2022

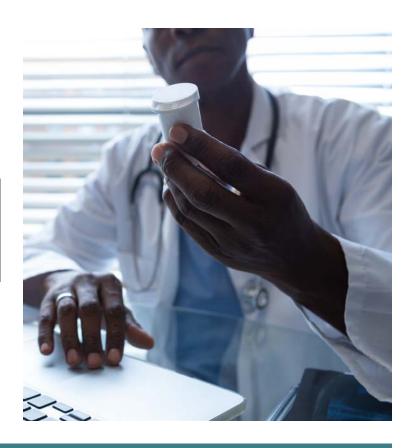


CONSIDER AN OPIOID ONLY WHEN:

Potential benefits are likely to outweigh risks

Patient has failed to adequately respond to non-opioid and nonpharmacological interventions

Patient has moderate to severe nociceptive or neuropathic pain



Begin as a therapeutic trial

SOURCES: Chou R, et al. J Pain. 2009;10:113-130. Department of Veterans Affairs, Department of Defense. VA/DoD Clinical Practice Guideline for Management of Opioid Therapy for Chronic Pain. 2017.



OPIOID MISUSE RISK ASSESSMENT TOOLS

http://core-rems.org/opioid-education/tools/



ORT-OUD Opioid Risk Tool

SOAPP® Screener and Opioid Assessment for Patients with Pain

DIRE Diagnosis, Intractability, Risk, and Efficacy score

TOOLS FOR SUBSTANCE USE DISORDER

CAGE-AID Cut down, Annoyed, Guilty, Eye-Opener tool, Adapted to Include Drugs

TAPS Tobacco, Alcohol, Prescription Medication and Other Substances Tool

DAST Drug Abuse Screening Test

CTQ Childhood Trauma Questionnaire

ACEs Adverse Childhood Experiences



A CLOSER LOOK AT THE ORT-OUD

Opioid Risk Tool - OUD (ORT-OUD)

This tool should be administered to patients upon an initial visit prior to beginning or continuing opioid therapy for pain management. A score of 2 or lower indicates low risk for future opioid use disorder; a score of >/=3 indicates high risk for opioid use disorder.

Mark each box that applies	YES	NO
Family history of substance abuse		
Alcohol	1	0
Illegal drugs	1	0
Rx drugs	1	0
Personal history of substance abuse		
Alcohol	1	0
Illegal drugs	1	0
Rx drugs	1	0
Age between 16-45 years	1	0
Psychological disease		
ADD, OCD, bipolar, schizophrenia	1	0
Depression	1	0
Scoring totals		

Substance use disorder history does not prohibit treatment with opioids but may require additional monitoring and expert consultation or referral.

Scoring:

• ≤ 2: low risk

≥ 3: high risk

SOURCE: Cheatle, M., et al. J Pain 2019; Jan 26.



OPIOID SIDE EFFECTS AND ADVERSE EVENTS

SIDE EFFECTS	ADVERSE EVENTS
Respiratory depression	Death
Opioid-induced constipation (OIC) (most common)	Addiction
Myoclonus (twitching or jerking)	Overdose
Sedation, cognitive impairment	Hospitalization
Sweating, miosis, urinary retention	Disability or permanent damage
Allergic reactions	Falls or fractures
Hypogonadism	
Tolerance, physical dependence, hyperalgesia	

Prescribers should report serious AEs and medication errors to the FDA: https://www.fda.gov/media/76299/download or 1-800-FDA-1088



OPIOID-INDUCED RESPIRATORY DEPRESSION

MORE LIKELY TO OCCUR:

- In elderly, cachectic, or debilitated patients
- If given concomitantly with other drugs that depress respiration (such as benzodiazepines*)
- In patients who are opioid-naïve or have just had a dose increase
- In patients with conditions causing respiratory compromise

HOW TO REDUCE RISK:

- Ensure proper dosing and titration
- Do not overestimate dose when converting dosage from another opioid product
 - Can result in fatal overdose with first dose
- Avoid co-prescribing benzodiazepines*
- Instruct patients to swallow tablets/capsules whole
 - Dose from cut, crushed, dissolved, or chewed tablets/capsules may be fatal, particularly in opioid-naïve individuals

*Greatest risk of respiratory depression



DRUG INTERACTIONS COMMON TO OPIOIDS

Other CNS Depressants

- Increased risk of respiratory depression, hypotension, profound sedation, or coma
- Reduce initial dose

Partial Agonists* or Mixed Agonist/Antagonists[†]

- Use caution with full opioid agonist
- May reduce analgesic effect and/or precipitate withdrawal

Skeletal Muscle Relaxants

 Concurrent use may enhance neuromuscular blocking action and increase respiratory depression

Anticholinergic Medication

- Concurrent use increases risk of urinary retention and severe constipation
- May lead to paralytic ileus



^{*}Buprenorphine; †Pentazocine, nalbuphine, butorphanol

FOR SAFER USE: KNOW DRUG INTERACTIONS, PHARMACODYNAMICS, AND PHARMACOKINETICS

CNS depressants can potentiate sedation and respiratory depression (e.g., benzodiazepines, gabapentin)

Some ER/LA products rapidly release opioid (dose dump) when exposed to alcohol Some drug levels may increase without dose dumping

Opioid use with MAOIs may increase respiratory depression

Certain opioids with MAOIs can cause serotonin syndrome (e.g., tramadol)

Opioid use can reduce efficacy of diuretics

Inducing release of antidiuretic hormone

Many opioids can prolong QTc interval, check the PI; methadone requires extra caution

Drugs that inhibit or induce CYP enzymes can increase or lower blood levels of some opioids



OPIOIDS AND CYP450 ENZYME INTERACTIONS

Metabolism of several commonly used opioids occurs through the cytochrome P450 system

Be aware of potential inhibitors (e.g., macrolides, azole antifungals) and inducers (e.g., carbamazepine)

Genetic and phenotypic variations in patient response to certain opioids

Refer to product-specific information in the drug package insert before prescribing

SOURCE: https://dailymed.nlm.nih.gov/dailymed/index.cfm



TRANSDERMAL/TRANSMUCOSAL DOSAGE FORMS



Do not cut, damage, chew, or swallow

Prepare skin: clip (not shave) hair and wash area with water

Rotate location of application

Do not apply buccal film products if film is cut, damaged, or changed in any way—use the entire film

Note that metal foil backings are not safe for use in MRIs

Monitor patients with fever for signs or symptoms of increased opioid exposure

Note that exertion or exposure to external heat can lead to fatal overdose





OLDER ADULTS

RISK FOR RESPIRATORY DEPRESSION

 Age-related changes in distribution, metabolism, excretion; absorption less affected



ACTIONS

- Monitor
 - Initiation and titration
 - Concomitant medications (polypharmacy)
 - Falls risk, cognitive change, psychosocial status
- Reduce starting dose to 1/3 to 1/2 the usual dosage in debilitated, non-opioid-tolerant patients
- Start low, go slow, but GO
- Routinely initiate a bowel regimen
- Patient and caregiver reliability/risk of diversion

CO*RE

SOURCEs: American Geriatrics Society Panel on the Pharmacological Management of Persistent Pain in Older Persons. J Am Geriatr Soc. 2009;57:1331-46; Chou R, et al. J Pain. 2009;10:113-30.

WOMEN OF CHILDBEARING POTENTIAL

Neonatal opioid withdrawal syndrome is a potential risk of opioid therapy

GIVEN THIS POTENTIAL RISK, CLINICIANS SHOULD:

- Discuss family planning, contraceptives, breastfeeding plans with patients
- Counsel women of childbearing potential about risks and benefits of opioid therapy during pregnancy and after delivery
- Encourage minimal/no opioid use during pregnancy, unless potential benefits outweigh risks to fetus
- Refer to a qualified provider who will ensure appropriate treatment for the baby
- Perform universal screening to avoid neonatal opioid withdrawal syndrome (NOWS)
- For women using opioids on a daily basis,
 ACOG recommends buprenorphine or methadone



ACOG = American College of Obstetricians and Gynecologists SOURCES: Chou R, et al. J Pain. 2009;10:113-30; ACOG Committee on Obstetric Practice, August 2017



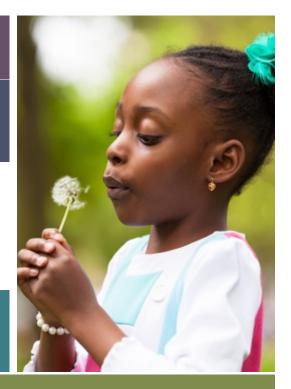
CHILDREN AND ADOLESCENTS

HANDLE WITH CARE: JUDICIOUS AND LOW-DOSE USE OF IR FOR BRIEF THERAPY

THE SAFETY AND EFFECTIVENESS OF MOST OPIOIDS ARE UNESTABLISHED

- Pediatric analgesic trials pose challenges
- Transdermal fentanyl approved in children ≥ 2 years
- Oxycodone ER dosing changes for children ≥ 11 years

ER/LA OPIOID INDICATIONS ARE PRIMARILY LIFE-LIMITING CONDITIONS



WHEN PRESCRIBING ER/LA OPIOIDS TO CHILDREN:

 Consult pediatric palliative care team or pediatric pain specialist or refer to a specialized multidisciplinary pain clinic

SOURCES: Berde CB, et al. *Pediatrics*. 2012;129:354-364; Gregoire MC, et al. Pain Res Manag 2013;18:47-50; Mc Donnell C. Pain Res Manag. 2011;16:93-98; Slater ME, et al. Pain Med. 2010;11:207-14.



OTHER POPULATIONS NEEDING SPECIAL TREATMENT CONSIDERATIONS

Persons with...

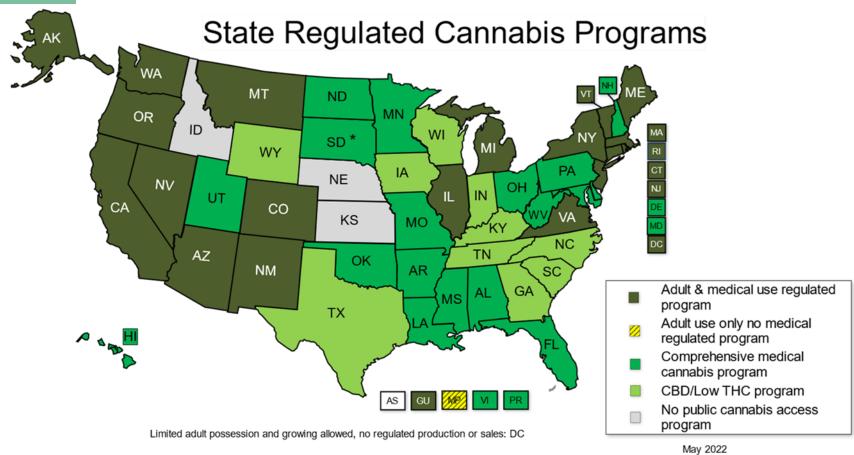
- Sleep disorders or sleep-disordered breathing (sleep apnea)
- Dementia/nonverbal patients
- Obesity
- Renal/hepatic impairment
- Psychiatric disorders
- Life-limiting illness
- Substance use disorder



Marijuana Status



Medical



Recreational

Not legal for recreational use in West Virginia

https://www.ncsl.org/research/health/state-medical-marijuana-laws.aspx, Sept. 2022



INFORMED CONSENT

When initiating a pain treatment plan, confirm patient understanding of informed consent to establish:

ANALGESIC AND FUNCTIONAL GOALS OF TREATMENT

EXPECTATIONS

POTENTIAL RISKS

ALTERNATIVES

PATIENT'S UNDERSTANDING

PATIENT'S DECISION





Telehealth technology allows new, effective, and efficient options for clinicians and patients to work in partnership to manage chronic medical issues



OPTIMIZING PATIENT CARE THROUGH TELEHEATH

New CO*RE CE/CME Module

- Series of four short videos
- Help HCPs conduct successful telehealth patient visits
- Available online https://learningipma.org

PATIENT PROVIDER AGREEMENT (PPA)

Reinforce Expectations For Appropriate And Safe Opioid Use

- Clarify treatment plans and goals
- One prescriber
- Consider one pharmacy
- Safeguards
 - Do not store in medicine cabinet
 - Keep locked (medication safe)
 - Do not share or sell
- Instructions for disposal when no longer needed
- Prescriber notification for any event resulting in a pain medication prescription

- Follow-up plan
- Monitoring
 - Random urine drug test
 (UDT) and pill counts
- Refill procedure
- Identify behaviors indicating need for discontinuation
- Exit strategy
- Signed by both



PATIENT PROVIDER AGREEMENT NONADHERENCE

Behavior outside the boundaries of agreed-on treatment plan

Unsanctioned dose escalations or other noncompliance with therapy on 1 or 2 occasions

Unapproved use of the drug to treat another symptom

Openly acquiring similar drugs from other medical sources

Multiple dose escalations or other noncompliance with therapy despite warnings

Prescription forgery

Obtaining prescription drugs from nonmedical sources

Any of the above behaviors merits **investigation**: proceed with caution





MANAGING PATIENTS ON OPIOID ANALGESICS

INITIATING OPIOIDS

- Begin a therapeutic trial with an immediate release (IR) opioid
- Prescribe the lowest effective dosage
- Use caution at any dosage, but particularly when:
 - Increasing dosage to ≥ 50 morphine milligram equivalents (MME)/day
 - Carefully justify a decision to titrate dosage to ≥ 90 MME/day
- Always include dosing instructions, including daily maximum
- Be aware of interindividual variability of response
- Have PPA, baseline UDT, and informed consent in place
- Co-prescribe naloxone and bowel regimen
- Re-evaluate risks/benefits within 1–4 weeks (could be as soon as 3–5 days) of initiation or dose escalation
- Re-evaluate risks/benefits every 1–3 months; if benefits do not outweigh harms, optimize other therapies and work to taper and discontinue

There are differences in benefits, risks, and expected outcomes for patients with chronic pain and cancer pain, as well as for hospice and palliative care patients.



ONGOING AND LONG-TERM MANAGEMENT OF PATIENTS ON OPIOID ANALGESICS

PERIODIC REVIEW OF PAIN

- Is the patient making progress toward functional goals?
- Reset goals if required or indicated; develop reasonable expectations
- Monitor for breakthrough pain
- Review adverse events/side effects at each visit
 - Evaluate bowel function
 - Screen for endocrine function as needed
 - Report adverse events to the FDA website
 - Implement opioid rotation, as indicated

Prescribers should report serious AEs and medication errors to the FDA: https://www.fda.gov/media/76299/download or 1-800-FDA-1088



ONGOING AND LONG-TERM MANAGEMENT OF PATIENTS ON OPIOID ANALGESICS

MONITORING FOR SAFETY

- Check Prescription Drug Monitoring Program (PDMP)
- Use urine drug testing (UDT)
- Reassess risk of substance use disorder (SUD) and/or OUD
- Monitor adherence to the treatment plan
 - Medication reconciliation
 - Evaluate for nonadherence

DISCONTINUING AND TAPERING

When is opioid therapy no longer necessary?



MONITORING PAIN AND SUBSTANCE USE DISORDER

PAIN – **5** A's

- Analgesia
- Activity/Function
- Aberrant/Problematic behavior, not present
- Adverse events
- Affect

SUD – 5 C's

- Control, loss of
- Compulsive use
- Craving drug
- Continued use
- Chronic problem



WHEN TO MOVE FROM IR TO ER/LA OPIOIDS

PRIMARY REASONS

- Maintain stable blood levels (steady state plasma)
- Longer duration of action
- Multiple IR doses needed to achieve effective analgesia
- Poor analgesic efficacy despite dose titration
- Less sleep disruption

OTHER POTENTIAL REASONS

- Patient desire or need to try a new formulation
- Cost or insurance issues
- Adherence issues
- Change in clinical status requiring an opioid with different pharmacokinetics
- Problematic drug-drug interactions





CONSIDERATIONS FOR CHANGE FROM IR TO ER/LA OPIOIDS

DRUG AND DOSE SELECTION IS CRITICAL

Some ER/LA opioids or dosage forms are only recommended for opioid tolerant patients

- ANY strength of transdermal fentanyl or hydromorphone ER
- Certain strengths/ doses of other ER/LA products (check drug prescribing information)

MONITOR PATIENTS
CLOSELY FOR
RESPIRATORY
DEPRESSION

 Especially within 24–72 hours of initiating therapy and increasing dosage INDIVIDUALIZE
DOSAGE BY
TITRATION BASED ON
EFFICACY,
TOLERABILITY,
AND PRESENCE OF
ADVERSE EVENTS

- Check ER/LA opioid product PI for minimum titration intervals
- Supplement with IR analgesics (opioid and non-opioid) if pain is not controlled during titration

SOURCES: Chou R, et al. J Pain. 2009;10:113-130; FDA. Education Blueprint Healthcare Providers Involved in the Treatment and Monitoring of Patients with Pain 09/2018, https://www.accessdata.fda.gov/drugsatfda_docs/rems/Opioid_analgesic_2018_09_18_FDA_Blueprint.pdf



EMERGENCE OF OPIOID-INDUCED HYPERALGESIA

- An increased sensitivity to pain
- Usually occurs at high MME dosages and over long periods of time
- A physiological phenomenon that can happen to anyone
- Consider this explanation if:
 - Pain increases despite dose increases
 - Pain appears in new locations
 - Patient becomes more sensitive to painful stimuli
 - Patient is not improving in the absence of underlying cause or disease progression

SOURCE: Yi P, Pryzbylkowski P. Opioid induced hyperalgesia. Pain Medicine 2015; 16: S32-S36





OPIOID TOLERANCE

If opioid tolerant, still use caution at higher doses

Patients considered opioid tolerant are taking at least

- 60 mg oral morphine/day
- 25 mcg transdermal fentanyl/hour
- 30 mg oral oxycodone/day
- 8 mg oral hydromorphone/day
- 25 mg oral oxymorphone/day
- An equianalgesic dose of another opioid

Also use caution when rotating a patient on an IR opioid to a different ER/LA opioid



SOURCE: The Opioid Analgesics Risk Evaluation & Mitigation Strategy product search, https://opioidanalgesicrems.com/products.html



OPIOID TOLERANCE VERSUS PHYSICAL DEPENDENCE

TOLERANCE

- Occurs when increased dose is needed to maintain the functional status no longer achieved by current dose
- Remember CNS and respiratory depression can develop with dose increase



PHYSICAL DEPENDENCE

- Occurs when an individual only functions normally in the presence of the substance
- Abrupt discontinuation or dosage decrease causes uncomfortable symptoms of withdrawal

Both **tolerance** and **physical dependence** are physiological adaptations to chronic opioid exposure and **DO NOT** equal addiction or opioid use disorder



OPIOID ROTATION

DEFINITION

A change from an existing opioid regimen to another opioid with the goal of improving therapeutic outcomes or to avoid AEs attributed to the existing drug



RATIONALE

Used when differences in pharmacologic or other effects make it likely that a switch will improve outcomes

- Effectiveness and AEs of different mu-opioids vary among patients
- Patient tolerant to first opioid might have improved analgesia from second opioid at a dose lower than calculated from an equianalgesic dosing table (EDT)

SOURCES: Fine PG, et al. J Pain Symptom Manage. 2009;38:418-425; Knotkova H, et al. J Pain Symptom Manage. 2009;38:426-439; Pasternak GW. Neuropharmacol. 2004;47(suppl 1):312-323.



EQUIANALGESIC DOSING TABLES (EDTs)

Many different versions:

Published

Online calculators



Smartphone apps

Vary in terms of:



Equianalgesic values

Whether ranges are used

Which opioids are included: May or may not include transdermal opioids, rapid-onset fentanyl, ER/LA opioids, or opioid agonist-antagonists



START WITH AN EDT FOR ADULTS



	EQUIANALGESIC DOSE		USUAL STARTING DOSE	
DRUG	SC/IV	РО	PARENTERAL	PO
Morphine	10 mg	30 mg	2.5–5 mg SC/IV q3–4hr (1.25–2.5 mg)	5–15 mg q3–4hr (IR or oral solution) (2.5–7.5 mg)
Oxycodone	NA	20 mg	NA	5–10 mg q3–4hr (2.5 mg)
Hydrocodone	NA	30 mg	NA	5 mg q3–4hr (2.5 mg)
Hydromorphone	1.5 mg	7.5 mg	0.2-0.6 mg SC/IV q2-3hr (0.2 mg)	1–2 mg q3–4hr (0.5–1 mg)



MU-OPIOID RECEPTORS AND INCOMPLETE CROSS TOLERANCE

MU-OPIOIDS BIND TO MU RECEPTORS

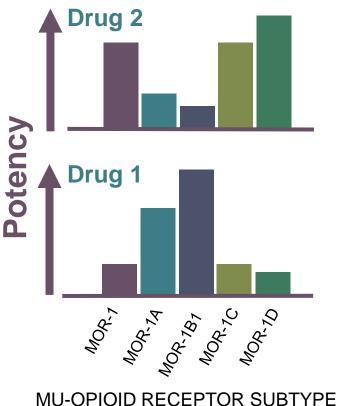
MANY MU RECEPTOR SUBTYPES

Mu-opioids produce subtly different pharmacologic responses based on distinct activation profiles of mu receptor subtypes

MAY HELP EXPLAIN:

Interpatient variability in response to mu-opioids

Incomplete cross tolerance among mu-opioids





GUIDELINES FOR OPIOID ROTATION

Calculate
equianalgesic
dose of new
opioid from
EDT

REDUCE CALCULATED EQUIANALGESIC DOSE BY 25%-50%*

SELECT % REDUCTION BASED ON CLINICAL JUDGMENT

CLOSER TO 50% REDUCTION

IF PATIENT...

- Is receiving a relatively high dose of current opioid regimen
- Is elderly or medically frail

CLOSER TO 25% REDUCTION

IF PATIENT...

- Does not have these characteristics
- Is changing route of administration



*75%–90% reduction for methadone



GUIDELINES FOR OPIOID ROTATION (continued)



IF SWITCHING TO METHADONE:

- Standard equianalgesic dosing tables are less helpful in opioid rotation to methadone
- For opioid tolerant patients, methadone doses should **not** exceed
 30–40 mg/day upon rotation
 - Consider inpatient monitoring, including serial EKG monitoring
- For opioid-naïve patients, do not give methadone as an initial drug

IF SWITCHING TO TRANSDERMAL:

• **Fentanyl:** calculate dose conversion based on equianalgesic dose ratios included in the drug package insert



GUIDELINES FOR OPIOID ROTATION: SUMMARY

VALUES FROM EDT* PATIENT OPIOID VALUES

SOLVE FOR X

AUTOMATICALLY REDUCE DOSE

Value of current opioid

Value of new opioid

24-hr dose of current opioid

X amount of new opioid

Equianalgesic 24-hr dose of new opioid

By 25%-50%[†]

Frequently assess initial response

Titrate dose of new opioid to optimize outcomes

Calculate supplemental rescue dose used for titration at 5%–15% of total daily dose[‡]



- * If switching to transdermal fentanyl, use equianalgesic dose ratios provided in Pl.
- [†] If switching to methadone, reduce dose by 75%–90%.
- ‡ If oral transmucosal fentanyl used as rescue, begin at lowest dose irrespective of baseline opioid.



BREAKTHROUGH PAIN (BTP)

PATIENTS ON STABLE ATC OPIOIDS MAY EXPERIENCE BTP

- Due to disease progression or a new or unrelated pain
 - Target cause or precipitating factors
- Dose for BTP: Using an IR, 5%–15% of total daily opioid dose, administered at an appropriate interval
- Never use ER/LA for BTP

CONSIDER ADDING

- PRN IR opioid trial based on analysis of benefit versus risk
 - There is a risk for problematic drug-related behaviors
 - High-risk: Add only in conjunction with frequent monitoring
 - and follow-up
 - Low-risk: Add with routine follow-up and monitoring
- Consider non-opioid drug therapies and nonpharmacologic treatments



ABUSE-DETERRENT FORMULATION (ADF) OPIOIDS

Drug formulations designed to discourage misuse

- An ER/LA opioid with properties to meaningfully deter misuse (less likely to be crushed, injected, or snorted)
- Consider as one part of an overall strategy
- Mixed evidence on the impact of ADF on misuse
- Overdose is still possible if taken orally in excessive amounts
- These products are expensive with no generic equivalents





URINE DRUG TESTING (UDT)



- Urine testing is done **FOR** the patient, not TO the patient
- Helps to identify drug misuse/addiction
- Assists in assessing and documenting adherence

CLINICAL CONSIDERATIONS

- Recommend UDT before first prescription (baseline), then intermittently, depending on clinical judgment and state regulations
- Document time and date of last dose taken
- Be aware of possible false positives or negatives
- Clarify unexpected results with the lab before confronting patient to rule out poor specimen or error



SCREENING VERSUS CONFIRMATORY UDTS





	SCREENING (Office-based)	CONFIRMATORY (Send to lab)
Analysis technique	Immunoassay	GC-MS or HPLC
Sensitivity (power to detect a class of drugs)	Low or none when testing for semi-synthetic or synthetic opioids	High
Specificity (power to detect an individual drug)	Varies (can result in false positives or false negatives)	High
Turnaround	Rapid	Slow
Cost/Other	Lower cost; intended for a drug- free population; may not be useful in pain medicine	Higher cost; legally defensible results



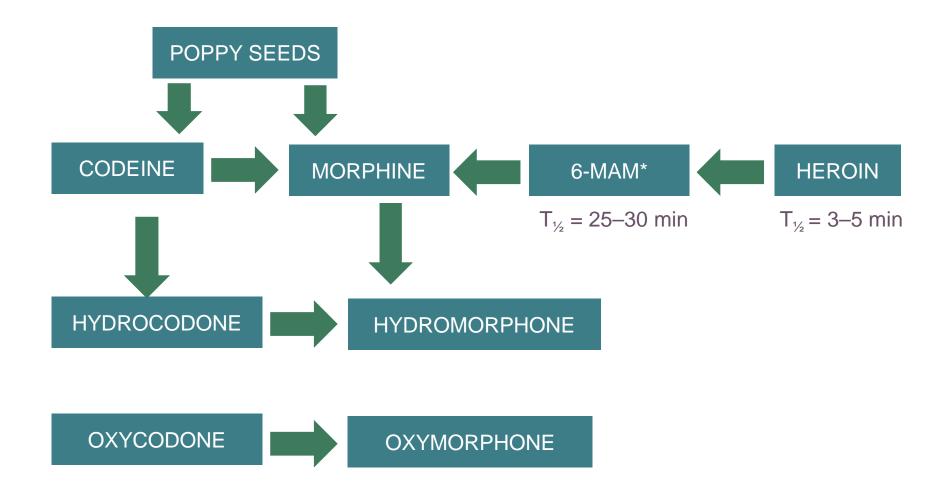
WINDOWS OF SPECIFIC DRUG DETECTION

Drug	How soon after taking drug will there be a positive drug test?	How long after taking drug will there continue to be a positive drug test?
Cannabis/ Tetrahydrocannabinol (THC)	1–3 hours	1–7 days (can be up to 1 month if long-term use)
Crack (cocaine)	2–6 hours	2–3 days
Heroin (opiates)	2–6 hours	1–3 days
Speed/uppers (amphetamine, methamphetamine)	4–6 hours	2-3 days
Angel dust/PCP	4–6 hours	7–14 days
Ecstasy	2–7 hours	2–4 days
Benzodiazepine	2–7 hours	1–4 days
Barbiturates	2-4 hours	1–3 weeks
Methadone	3–8 hours	1-3 days (up to 2 weeks)
Tricyclic antidepressants	8–12 hours	2-7 days
Oxycodone	1–3 hours	1–2 days

SOURCE: http://www.fda.gov/MedicalDevices/ProductsandMedicalProcedures/InVitroDiagnostics/DrugsofAbuseTests/ucm125722.htm



EXAMPLES OF OPIOID METABOLISM





CONSIDERATIONS FOR RE-EVALUATING OPIOID USE

PATIENT MOVES
PAST THE POINT
OF NEED

INTOLERABLE AND UNMANAGEABLE AEs

NO PROGRESS TOWARD THERAPEUTIC GOALS

RISKS OUTWEIGH BENEFITS

MISUSE BEHAVIORS

- One or two episodes of increasing dose without prescriber knowledge
- Sharing medications
- Unapproved opioid use to treat another symptom (e.g., insomnia)

- Use of illicit drugs or unprescribed opioids
- Repeatedly obtaining opioids from multiple outside sources
- Prescription forgery
- Multiple episodes of prescription loss
- Diversion

Even at prescribed doses, opioids carry the risk of misuse, abuse, opioid use disorder, overdose, and death



TOOLS TO REASSESS OUD/SUD RISK



SBIRT

Screening, Brief Intervention, and Referral to Treatment

TAPS

Tobacco, Alcohol, Rx, and Other Substances

PDUQ

Prescription Drug Use Questionnaire

PMQ

Pain Medication Questionnaire

COMM

Current Opioid Misuse Measure



APPROACHES TO SUPPORT THE DISCONTINUATION DECISION

- Discontinue through a taper schedule
- If OUD suspected:
 - Begin treatment: Medications for Opioid Use Disorder (MOUD)
 - Refer to an OUD specialist
- Consider rotation to partial agonist (e.g., buprenorphine)
- No single approach is appropriate for all patients
- May use a range of approaches, from a slow 10% dose reduction per week to a more rapid 25%–50% reduction every few days
- To minimize withdrawal symptoms in patients physically dependent on opioids, consider medications to assist with withdrawal (clonidine, NSAIDs, antiemetics, antidiarrheal agents)



CONSULTING A PAIN SPECIALIST

- Appropriate when you feel you cannot provide the level of care needed
- First ensure you have a reliable specialist to refer to
- To find a pain specialist in your area:
 - Consult with state boards
 - Consult with colleagues
 - Use online resources
 - Consult payment source
- Prior to referral, contact the specialist and ask what is needed for referral



Adequately **DOCUMENT**all patient interactions,
assessments, test results,
treatment plans,
and expectations.





EDUCATING YOUR PATIENTS AND THEIR CAREGIVERS

COUNSEL PATIENTS ABOUT PROPER USE

- Take opioid as prescribed
- Use least amount of medication necessary for shortest time
- Use caution with long-term opioid use patients; avoid abrupt discontinuation or dose reduction; taper safely to avoid withdrawal symptoms
- Notify HCP if pain is uncontrolled
- Report side effects to HCP
- Inform HCP of ALL meds and supplements being taken
- Never share or sell opioids: can lead to others' deaths, against the law
- Use caution when operating heavy machinery and driving





USE FDA PATIENT COUNSELING DOCUMENT

- What are opioids?
- What are the risks and benefits?
- How to take safely

What You Need to Know About Opioid Pain Medicines

This guide is for you! Keep this guide and the Medication Guide that comes with your medicine so you can better understand what you need to know about your opioid pain medicine. Go over this information with your healthcare provider. Then, ask your healthcare provider about anything that you do not understand.

What are opioids?

Opioids are strong prescription medicines that are used to What are the serious risks of using opioids? manage severe pain.

- Opioids have serious risks of addiction and overdose.
- Too much opioid medicine in your body can cause your breathing to stop - which could lead to death. This risk is greater for people taking other medicines that make you feel sleepy or people with sleep apnea.
- Addiction is when you crave drugs (like opioid pain medicines) because they make you feel good in some way. You keep taking the drug even though you know it is not a good idea and bad things are happening to you. Addiction is a brain disease that may require ongoing

Risk Factors for Opioid Abuse:

- » a history of addiction
- » a family history of addiction

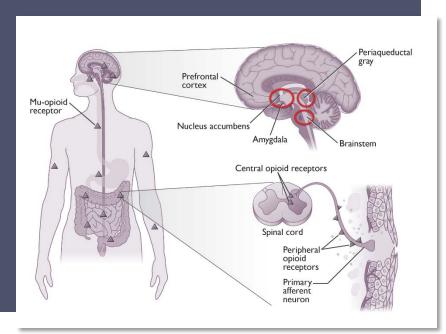
- Do not cut, break, chew, crush, or dissolve your medicine. If you cannot swallow your medicine whole, talk to your healthcare provider.
- When your healthcare provider gives you the prescription, ask:
- » What should I do if I need to taper off the opioid medicine » How long should I take it?
- Call your healthcare provider if the opioid medicine is not controlling your pain. Do not increase the dose on your own.
- Do not share or give your opioid medicine to anyone else. Your healthcare provider selected this opioid and the dose just for you. A dose that is okay for you could cause an overdose and death for someone else. Also, it is against the law.
 - Store your opioid medicine in a safe place where it cannot be reached by children or stolen by family or visitors to your home. Many teenagers like to experiment with pain medicines. Use a lock- box to keep your opioid





PROVIDE ANTICIPATORY GUIDANCE ON OPIOID SIDE EFFECTS AND ADVERSE EVENTS

- Overdose and death: respiratory depression
- Opioid-induced constipation (OIC): most common
- Nausea, vomiting, GERD
- Sexual dysfunction and other endocrine abnormalities (hypogonadism)
- Tolerance, physical dependence
- Hyperalgesia
- Allergic reactions
- Sedation, cognitive impairment
- Falls and fractures
- Sweating, miosis, urinary retention
- Myoclonus (twitching or jerking)
- Opioid use disorder (OUD)





COUNSEL PATIENTS AND CAREGIVERS

WARNINGS (Safe Administration)

- Never break, chew, crush, or snort an opioid tablet/capsule
- Never cut or tear patches or buccal films
- If patient cannot swallow, determine if appropriate to sprinkle contents on applesauce or administer via feeding tube
- Use of CNS depressants or alcohol with opioids can cause overdose

WHAT TO LOOK FOR (Safety Concerns)

- Cravings
- Being unable to fulfill work/family obligations
- Nodding off
- Taking more than prescribed



OPIOID-INDUCED RESPIRATORY DEPRESSION

If not immediately recognized and treated, may lead to respiratory arrest and death

More likely to occur in opioid-naïve patients during initiation or after dose increase

Instruct patients/family members to:

- Screen for shallow or slowed breathing
- Deliver NALOXONE
- CALL 911

Instructions may differ if patient is on hospice or near end of life

Greatest risk: when co-prescribed with a benzodiazepine



SIGNS OF ACCIDENTAL OPIOID POISONING: CALL 911

- Person cannot be aroused or is unable to talk
- Any trouble with breathing, heavy snoring is warning sign
- Gurgling noises coming from mouth or throat
- Body is limp, seems lifeless; face is pale, clammy
- Fingernails or lips turn blue/purple
- Slow, unusual heartbeat or stopped heartbeat

Administer Naloxone







NALOXONE

WHAT IT IS:

- An opioid antagonist administered intranasally (most common) or parenterally
- Reverses acute opioid-induced respiratory depression but will also reverse analgesia; may precipitate acute opioid withdrawal
- No misuse potential

WHAT TO DO:

- Discuss an overdose plan with patients; involve family/caregivers
- Ensure family/caregivers have access to naloxone; some states require coprescribing
- Involve and train family, friends, partners, and/or caregivers in the proper administration of naloxone
- Know your local naloxone resources (e.g., the library, community centers)
- Check expiration dates and replace expired naloxone
- In the event of known or suspected overdose, call 911 and administer naloxone



NALOXONE OPTIONS

- Available as auto-injector, intramuscular injection, or nasal spray
- Cost and insurance coverage vary
- Make use of tutorial videos or live demonstration to educate patient/family/caregiver on proper administration
- Store at room temperature







Naloxone vials

Narcan nasal spray

Evzio (auto-injector)

Trade names are used for identification purposes only and do not imply endorsement.

SOURCE: FDA Information About Naloxone, https://www.fda.gov/DrugSafety/PostmarketDrugSafety/InformationforPatientsandProviders/ucm472923.htm



Naloxone Regulation



Effective date	• June 2018	
Criminal Immunity	 Prescribers: Yes Dispensers: Yes Lay People: Yes 	
Also Available	 Without Prescription: Yes To 3rd Party: Yes By Standing Order: Yes 	
Carried by First Responders	• Yes	

https://www.networkforphl.org/wp-content/uploads/2021/05/NAL-Final-4-29.pdf, July 1, 2020 State Naloxone Access Rules and Resources - SAFE Project, January 2023 https://legislativeanalysis.org/wp-content/uploads/2021/05/Naloxone-Access-Summary-of-State-Laws-Final.pdf Oct 2020

SAFE OPIOID STORAGE AND DISPOSAL



STEP 1: MONITOR

- Note how many pills are in each prescription
- Keep track of dosage and refills
- Make sure everyone in the home knows meds are tracked (if appropriate)

STEP 2: SECURE

- Keep meds in a safe place (locked cabinet or box)
- Store away from children, family, visitors, and pets
- Extra precautions needed with adolescents in the home

STEP 3: DISPOSE

- Discard expired or unused meds
- Check your local disposal options (e.g., pharmacy, police)

SOURCE: McDonald E, Kennedy-Hendrick A, McGinty E, Shields W, Barry C, Gielen A. Pediatrics. 2017;139(3):e20162161



WHERE AND HOW TO DISPOSE OF UNUSED OPIOIDS





Authorized Collection Sites

Use the DEA disposal locator website to find sites near you:

Search Google Maps for "drug disposal nearby"

Options

- Check with local pharmacy for disposal options
- Flush
 - Fold patch in half so sticky sides meet, then flush
- Trash (mix with noxious element like kitty litter or compost)



Mail-Back Packages

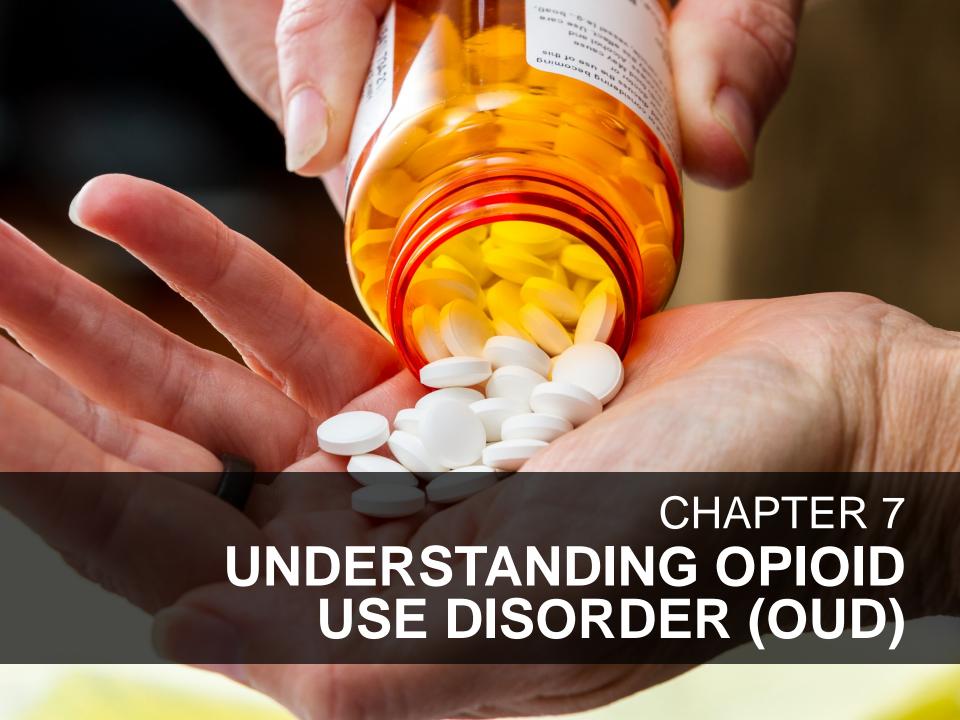
Obtain from authorized collectors

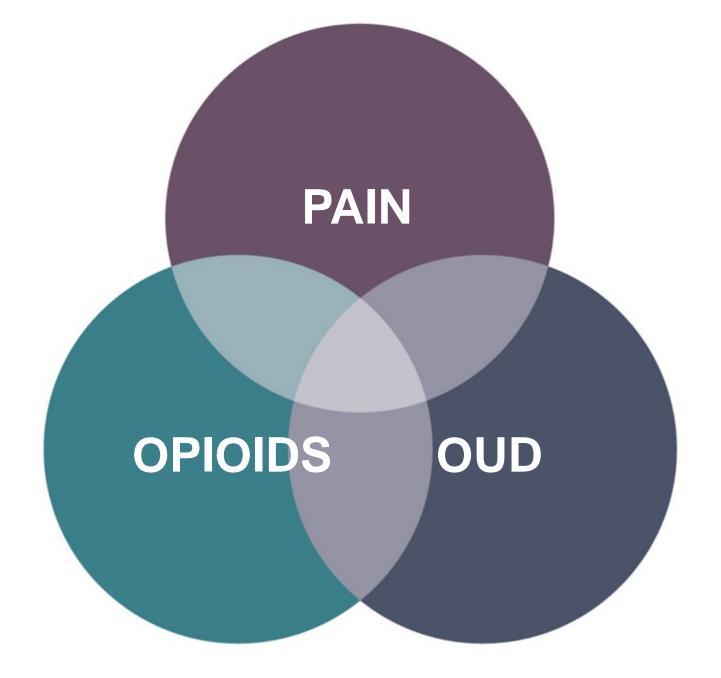


EPA. How to Dispose of Medicines Properly. https://archive.epa.gov/region02/capp/web/pdf/ppcpflyer.pdf









WHAT IS ADDICTION?



PRACTICAL DEFINITION:

Addiction is the continued use of drugs or activities, despite knowledge of continued **harm** to one's self or others.

OFFICIAL ASAM DEFINITION:

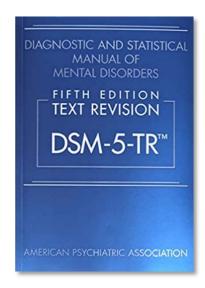
Addiction is a treatable, chronic medical disease involving complex interactions among brain circuits, genetics, the environment, and an individual's life experiences. People with addiction use substances or engage in behaviors that become compulsive and often continue despite harmful consequences. Prevention efforts and treatment approaches for addiction are generally as successful as those for other chronic diseases.



OPIOID USE DISORDER: DSM-5-TR CRITERIA

Be alert to these factors in your patients on long-term opioid therapy

- Taking larger amounts and/or for longer periods than intended
- 2. Persistent desire or inability to cut down or control use
- 3. Increased time spent obtaining, using, or recovering
- 4. Craving/compulsion to use opioids
- Role failure at work, home, school
- 6. Social or interpersonal problems
- 7. Reducing social, work, recreational activity
- 8. Physical hazards
- 9. Physical or psychological harm
- Tolerance
- Withdrawal



- 2-3 = mild
- 4-5 = moderate
- ≥6 = severe
- Not valid if opioid is taken as prescribed



WORDS MATTER





HOW TO IDENTIFY RISK FOR MY PATIENTS

10%–26% of patients on chronic opioid therapy (COT) for chronic noncancer pain (CNCP) may develop OUD

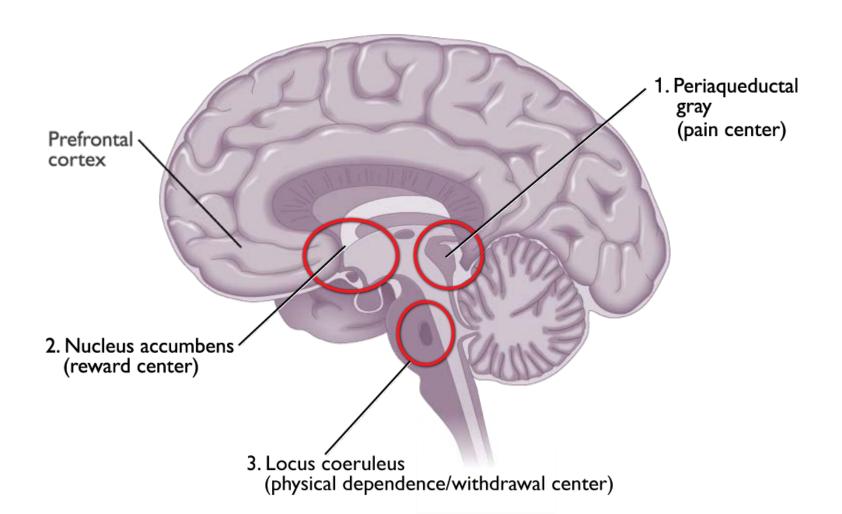
What to look for:

- High dosages
- Prolonged use
- Low hedonic tone
- Mental health disorders
- Past history of substance use disorder

Clinical judgment is key.

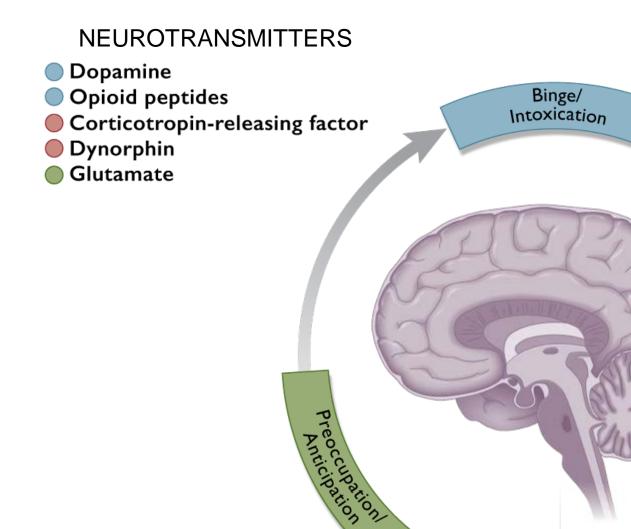


OPIOID RECEPTORS IN THE BRAIN: RELATIONSHIP TO ANALGESIA, OUD, AND WITHDRAWAL





THE CYCLE OF SUBSTANCE USE DISORDER



N. With drawall

Medication for Opioid Use Disorder (MOUD)

- Important and evidence-based medication that saves lives
- You can start from your office, as an outpatient
- Patients with OUD have decreased mortality when treated

There are three medication options:

- 1. Buprenorphine (Schedule III)
- 2. Methadone (Schedule II)
- 3. Naltrexone (not a controlled substance)

Are we just replacing one drug with another?

Myth or fact?



BUPRENORPHINE

- The most commonly prescribed pharmacotherapy for the treatment of OUD
- Partial mu-agonist with "plateau effect" for respiratory depression
- Good efficacy and safety profile
- Congress eliminated the X-waiver requirement to prescribe Bup
- All DEA-licensed HCPs can prescribe without patient number caps
- Long-acting and sublingual form indicated to treat opioid withdrawal and craving

FDA-approved bup products for pain:

- Butrans: 7-day transdermal patch
- Belbuca: buccal mucosal film; BID dosing



AVOID OTHER SUBSTANCES THAT COULD CONTRIBUTE TO AN ACCIDENTAL OVERDOSE

- Benzodiazepines (BZDs), sedatives, muscle relaxants; they are CNS depressants
- More than 30% of opioid overdoses involve benzodiazepines (BZDs)
- Use a comprehensive SUD evaluation to support recovery efforts for all substances



SOURCE: NIDA. Takaki H, et al. Am Journal Addictions. 2019;1-8.



USE A WHOLE-PERSON APPROACH WHEN TREATING A PATIENT WITH OUD FOR PAIN

- Must address both pain and opioid use disorder
- Remember that untreated pain is a trigger for return to use
- Avoid other potentially problematic medications
- Consider a multimodal pain program, including nonpharma options
- Avoid stigmatizing patients who are on long-term opioids for pain

- Consider buprenorphine for both pain and OUD
- Enlist patient's family/caregivers to secure and dispense opioids
- Recommend an active recovery program
- Remember to use PDMP
- Use screening methods (UDT, pill counts, PPA) to identify challenges and initiate discussion

SOURCE: Bailey J, et al. Pain Med 2010;11:1803-1818.



REFERRALS AND TREATMENT CENTERS

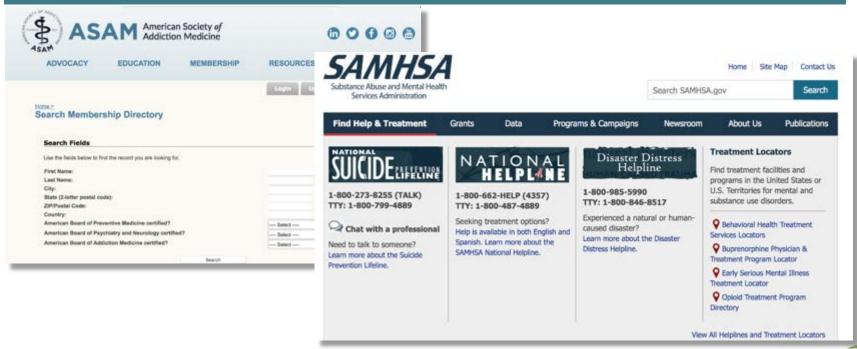
ASAM, SAMHSA, and AAAP are all helpful referral resources.

ASAM resources:

https://asam.ps.membersuite.com/directory/SearchDirectory_Criteria.aspx

SAMHSA locator:

AAAP locator:



IN SUMMARY

- There is a place for opioids, but use caution
- Use multimodal therapies as part of the pain management care plan
- Screen for OUD risk with a validated instrument
- Continually reassess patients using opioids
- Patient and family/caregiver education is essential
- If you suspect OUD, begin treatment



THANK YOU!

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