

# Striking a Balance

Understanding Pain
Management and Opioids



#### **DISCLOSURE**

Authors have no conflicts of interest to disclose

Presentation will include discussion of off label medication use

INTERMOUNTAIN HEALTHCARE AND AUTHORS HAVE NOT RESEARCHED OR INDEPENDENTLY VALIDATED ANY ASSERTIONS OR CLAIMS AS DESCRIBED IN THIS PRESENTATION AS SUCH, INTERMOUNTAIN HEALTHCARE AND AUTHORS OFFERS NO ASSURANCE AS TO THE INFORMATION DESCRIBED HEREIN, ANY SUCH RELIANCE AND LIABILITY IS SOLELY ON THE INDIVIDUAL REVIEWEING AND RELYING UPON ANY USE.





# BIO:

**Norman Bowers, PA-C** has worked in the field of pain management since 2016. Prior to this he has worked in Southeastern New Mexico providing care for ear nose and throat patients. He currently sees patients suffering from a variety of chronic pain issues, including chronic back pain, neck pain, neuropathic pain, and has a special interest in providing care to a variety of headache and migraine conditions. He uses an interdisciplinary approach to help patients live the healthiest lives possible.

Norman is from Arizona, and enjoys basketball, volleyball, golf, hiking, and camping with his family

#### DISCLOSURE:

Norman Bowers has no financial relationships with the manufacturers of goods or services discussed, or corporate supporters of this activity.



# BIO:

Mathew Romankowski, MD was raised in Salt Lake City, UT and attended Westminster College prior to completing medical school at the University of Utah School of Medicine in 2013. Completed Transitional Year in Tucson, AZ at THMEP in 2014 followed by Anesthesiology residency at Jackson Memorial Hospital/University of Miami and the University of Florida in Gainesville. Further subspecialized with ACGME fellowship in Multidisciplinary Pain Medicine at the University of Florida in 2019. I have been practicing pain medicine at Utah Valley Pain Management since October 2019 with keen focus on multimodal and interdisciplinary treatment strategies to care for our diverse chronic pain patient population. I have particular interest in interventions and neuromodulation including dorsal column, DRG, and peripheral nerve stimulation.

#### DISCLOSURE:

Dr. Romankowski has no financial relationships with the manufacturers of goods or services discussed, or corporate supporters of this activity.



# BIO:

Joshua Minori, DO is a double-boarded physician specializing in non-surgical management of neuromusculoskeletal disorders as well as acute and chronic pain conditions. He completed his fellowship in pain medicine at the University of California Davis. His residency in physical medicine and rehabilitation was completed at Schwab Rehabilitation Hospital/University of Chicago. He spent an additional year as a research fellow at the Rothman Orthopedic Institute at Thomas Jefferson University Hospital. Before attending medical school at the Philadelphia College of Osteopathic medicine, he earned a degree in athletic training and worked as a Certified Athletic Trainer in high schools, colleges and physical therapy clinics.

#### DISCLOSURE:

Dr. Minori has no financial relationships with the manufacturers of goods or services discussed, or corporate supporters of this activity.







# BIO:

Joy Hrushka, DO received her medical degree from Edward Via College of Osteopathic medicine in South Carolina, and subsequently completed her anesthesiology residency at the University of Louisville in Kentucky. She recently completed her fellowship training at the University of Utah to specialize in Pain Medicine. She has extensive experience in managing acute pain in a perioperative setting from her anesthesia background as well as having knowledge in treating chronic pain from her specialty training. She believes in approaching a patient from an interdisciplinary aspect utilizing procedural interventions, medications, behavioral health counseling focused on pain, and physical therapy through an individualized approach.

#### DISCLOSURE:

Dr. Hrushka has no financial relationships with the manufacturers of goods or services discussed, or corporate supporters of this activity.

# **ACKNOWLEDGMENTS**

Presented by the California Academy of Family Physicians, a member of the CO\*RE Collaborative, ten interdisciplinary organizations working together to improve pain management and prevent adverse outcomes. For more information about CO\*RE, visit <a href="http://core-rems.org/">http://core-rems.org/</a>.

This activity is supported by an independent educational grant from the Opioid Analgesics REMS Program Companies (RPC). This activity is intended to be fully compliant with the Opioid Analgesic (OA) 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.

Scan the QR code to go to the FDA OA REMS Blueprint



# THE CO\*RE COLLABORATIVE

This course does not advocate for or against the use of opioids.

We intend to help clinicians 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.























# **CO\*RE FACULTY ADVISORY PANEL**



Michael Cheshire, DO, FACOI, FACP WEST VIRGINIA SCHOOL OF MEDICINE



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None of the Faculty Advisors, Reviewers, or Planners for this educational activity have relevant financial relationships with ineligible companies to disclose.

This course is based on the FDA's Opioid Analgesic REMS (FDA Blueprint, Sept. 2018) and existing guidelines, including the 2022 CDC Clinical Practice Guideline for Prescribing Opioids for Pain.

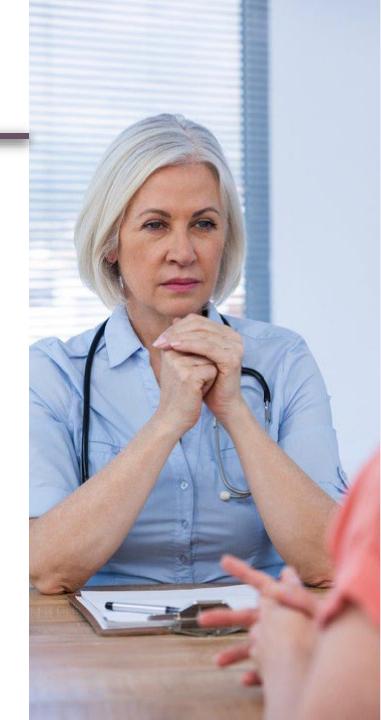




Barbara St. Marie,

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

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





# Cannabis and Pain

November 2023





# **FACULTY CONTENT DEVELOPERS/REVIEWERS**



Dr. Tauheed Zaman Director, Addiction Consult & Opioid Safety San Francisco Veterans Administration

He has no relevant financial relationships with ineligible companies to disclose.

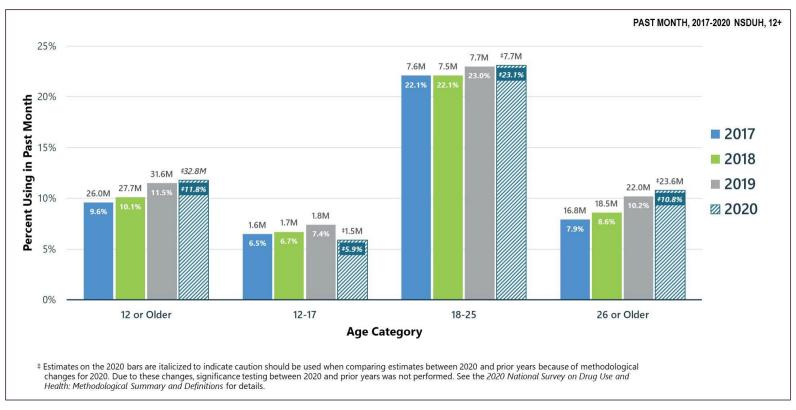


Dr. Smita Das Medical Director of Psychiatry Lyra Health

She has no relevant financial relationships with ineligible companies to disclose.



# **EPIDEMIOLOGY & RECENT TRENDS**



- Most commonly-used federally illicit substance in the U.S.
- 44% of people aged 19-30 used in the last year with daily use at 11%, all time highs
- Use is increasing among those 12+ and 26 +



### THE CANNABIS PLANT

- Cannabis is a complex plant with over 500 compounds.
- Multiple species of cannabis including sativa, indica and rudelaris.
   Marijuana often refers to the dried leaves, flowers, stems and seeds.
- Cannabinoids are stored in trichomes on plant surface, extracted and put into consumable form
- Most commonly inhaled



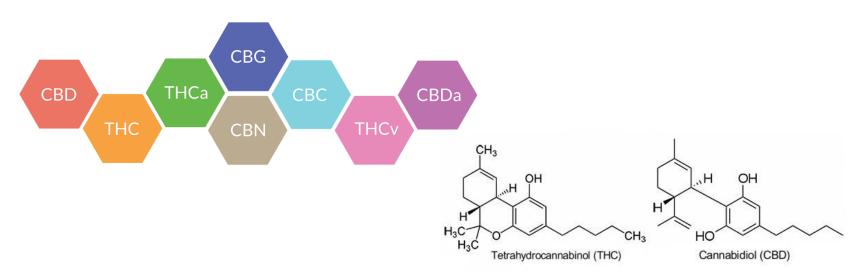


Trichomes on cannabis leaf surface



# CHEMICAL COMPOSITION

- Over 100 cannabinoids in cannabis plants, most unstudied
- THC associated with more negative effects (euphoria, addiction)
- CBD is being studied for therapeutics, but outside of synthesized CBD for rare seizure disorders, no well-established indications
- Preparations often labeled with inaccurate THC & CBD content
- Varying concentration, other cannabinoids may have health effects



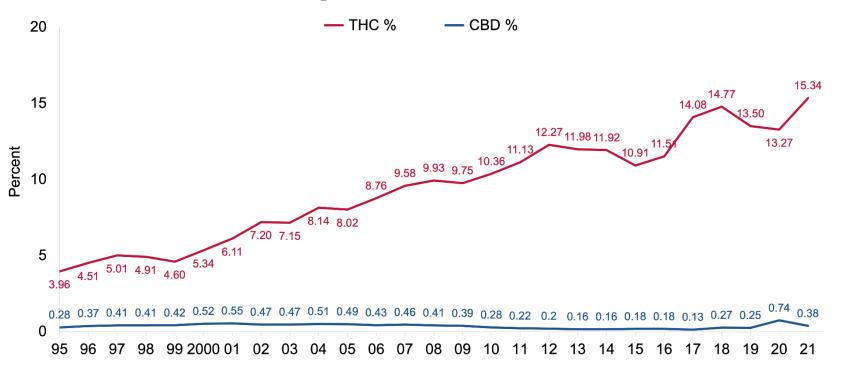
SOURCE: Hayakawa, K. et al. Therapeutic Potential of Non-Psychotropic Cannabidiol in Ischemic Stroke. Pharmaceuticals 2010, 3, 2197-2212



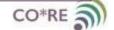
# INCREASED THC POTENCY OVER TIME

THC concentration increased  $5\% \rightarrow > 30\%$  in recent years

# Percentage of THC and CBD in Cannabis Samples Seized by the DEA, 1995-2021



SOURCE: U Miss, Potency Monitoring Project



# **NEWER CANNABINOIDS IN USE BY PATIENTS**

CANNABINOL (CBN)	CANNABIGEROL (CBG)
<ul> <li>Oxidation byproduct of THC</li> </ul>	Present in small amounts
<ul> <li>Less potent than THC</li> </ul>	Precursor to other cannabinoids
<ul> <li>Used for sleep problems, depression, anxiety, pain</li> </ul>	<ul> <li>Theoretical anti-inflammatory properties</li> </ul>
<ul> <li>No rigorous studies</li> </ul>	Used for pain management
	<ul> <li>No rigorous studies</li> </ul>

SOURCE: Corroon, Cannabis and Cannabinoid Res, 2021; Navarro et al, Cannabis and Cannabinoid Res, 2020;



# **PREPARATIONS**

PREPARATIONS	DESCRIPTION	USE
MARIJUANA	Dried plant product consisting of leaves, stems, and flowers	Smoked or vaporized
HASHISH	Concentrated resin cake	Ingested or smoked
TINCTURE	Cannabinoid liquid extracted from plant	Consumed sublingually
HASHISH OIL	Oil obtained from Cannabis plant by solvent extraction	Smoked or vaporized
INFUSION	Plant material mixed with nonvolatile solvents (e.g., butter, cooking oil)	Ingested



# **PREPARATIONS & PACKAGING**



Political Property of the Control of

SOURCE: wyomingpublicmedia.org

SOURCE: Hoodline 2017







SOURCE: Lindsay Fox/Wikimedia Commons



# "MEDICAL" CONTAINERS





SOURCE: Zaman 2016



# SYNTHETIC CANNABIONOID PRODUCTS

- Typically chemicals sprayed onto dried plant product
- Examples: K2, Spice, Joker, Black Mamba, Kush, KronicSynthetic chemicals sprayed onto dried, shredded plant product
- Mimic THC, bind strongly to same receptors → stronger effects
- Could cause changes in mood/perception, psychosis, tachycardia, vomiting, violent behavior, SI, renal impairment, seizures, death
- Often undetectable in standard urine drug tests
- Warn patients against using these products, severe adverse effects







# **MECHANISM: HOW DOES CANNABIS WORK?**

#### The Endocannabinoid System

Brain cells (neurons) communicate with each other by sending chemical messages. The chemicals (neurotransmitters) cross a gap between neighboring neurons before attaching to their specific receptors.

#### Presynaptic:

The neuron sending a message by releasing a chemical when signaled to do so

Postsynaptic: The neuron receiving the message when its receptors are activated by specific chemicals (neurotransmitters)

Neurotransmitters: The chemical messengers that travel from one brain cell to another

Receptors: Activated by neurotransmitters, receptors trigger a set of events that allows a message to be passed along to other neurons

Cannabinoid Receptor Presynaptic (sending neuron) autotrusulters Cannabinoids **Lipid Precursors** (fat cells) Postsynaptic (receiving neuron)

Cannabinoids: Natural chemicals (anandamide and 2-AG) that bind to cannabinoid receptors in the brain and the body

THC: The main active ingredient in marijuana; THC, also a cannabinoid, interferes with the normal functioning of the endocannabinoid system

- Endogenous cannabinoids originate from postsynaptic membrane
- Act on presynaptic cannabinoid receptors
- Modulate release of neurotransmitters (e.g., dopamine)
- Exogenous cannabinoids co-opt this system
- Also affects 5HT, alpha, TRPV, TRPA receptors



# THEORETICAL MECHANISMS FOR ANALGESIA

- Cells in injured tissue release endocannabinoids
- CB1-r in brain, spinal cord → mitigate sensitization, inflammation
- CB2-r in brain, spinal cord, dorsal root ganglion → reduce inflammatory hyperalgesia
- Long-term studies of exogenous cannabinoids and pain still needed; other mechanisms possible
- Caution with any drug where subjective pain improves, but has addictive properties (e.g., alcohol, opioids, benzodiazepines)



### PERCEPTIONS OF MEDICAL EFFICACY vs DATA

# **Perceptions**

- 81% of patients believe marijuana has at least one benefit
- 66% of patients believe in pain benefit

#### **Data**

- Systematic Review of RTCs: 2021: Outcomes had low or very lowquality evidence, neither supporting nor refuting efficacy
- Meta analysis 2022: Placebo contributes significantly to pain reduction in cannabis clinical trials
- Review 2022: High THC:CBD products (>98% THC) associated with 25% reduction in pain in short-term studies of variable quality

SOURCES: Steigerwald et al, J Addict Med, 2020; Fisher et al Pain 2021; Gedin et al, JAMA 2022; McDonagh Annals 2022



# **OPIOID-SPARING THEORY vs DATA**

**Theory:** If cannabis products treat pain, patient may use these products and reduce their use of opioids

### Data

- States with medical cannabis have modestly lower rates of opioid prescribing and risky opioid prescribing
- 2019 Study: Association between med cannabis and reduced opioid mortality has reversed over time
- 2021 Meta Analysis: Opioid-sparing effects remain uncertain due to very low evidence
- 2022 Meta Analysis: Preclinical/observational studies show opioid-sparing effect, but higher-quality RCTs do not
- 2023 Living Systematic Review: Cannabis impact on use of opioids remains insufficient



SOURCES: Shah et al, JGIM 2019, Noori et al BMJ Open 2021; Nielsen Neuropsychopharm 2022; Chou et al, AHRQ, 2023.

# **MEDICAL INDICATIONS**



# Psychiatric:

Not well-studied or FDA-approved for any psychiatric condition

# **Non-Psychiatric FDA Approvals:**

- Nausea, vomiting related to chemotherapy
- Anorexia/wasting related to HIV
- Rare childhood forms of epilepsy

The American Psychiatric Association has a Position Statement Against the Use of Cannabis for PTSD and a Position Statement in Opposition to Cannabis as Medicine



# **FDA-APPROVED CANNABINOIDS**

Medication	Type	Indication
<b>Dronabinol</b> (Marinol; Syndros)	Synthetic	Anorexia/wasting in AIDS patients
Nabilone (Cesamet)	Synthetic	Nausea, vomiting in chemotherapy patients
Cannabidiol (Epediolex)	Plant-derived	Lennox-Gastaut; Dravet's



# **CLINICAL CONSIDERATIONS**

- Individual risk stratification is crucial
  - Person/family history of mental health, addictions
  - Baseline psychosis risk
  - Risks related to driving, work, education, parenting
  - Medical, cognitive issues worsened by cannabis
- Counsel patients
  - Federally, cannabis is illegal (Schedule 1)
  - States vary
  - Review harm reduction strategies
- Use PPA and document conversations about risks
- Seek institutional legal counsel to reduce liability



# **CANNABIS AND HARM REDUCTION**

- Abstinence is best way to avoid health risks
- Avoid early-age initiation
- Avoid high frequency use (daily or near daily)
- Choose low-potency THC or balanced THC:CBD ratios
- Abstain from synthetic products
- Avoid combustible products, non-smoking methods preferable
- Avoid use in older adults due to risks

- Avoid deep/risky inhalation
- Abstain from cannabisimpaired driving
- High-risk populations should avoid use (e.g., psychosis, addictions)
- Track use over time, including metered dosing
- Set a time limit and goal for use, and continue to monitor for signs of Use Disorder



# **KRATOM**

- Substance in powder form, derived from tropical plant
- Prevalence 0.8% (past yr); 1.3 % (life-time use)
- Contains *mitragynine*, 7-alphahydroxymitragynine
- Not a cannabinoid
- Has opioid and stimulant effects (pain, detox, high)
- Can produce opioid withdrawal, addiction
- Does not appear on standard UDT
- Treat with supportive meds, opioid agonists/partial agonists
- Counsel patients on risk, refer for addiction eval as appropriate





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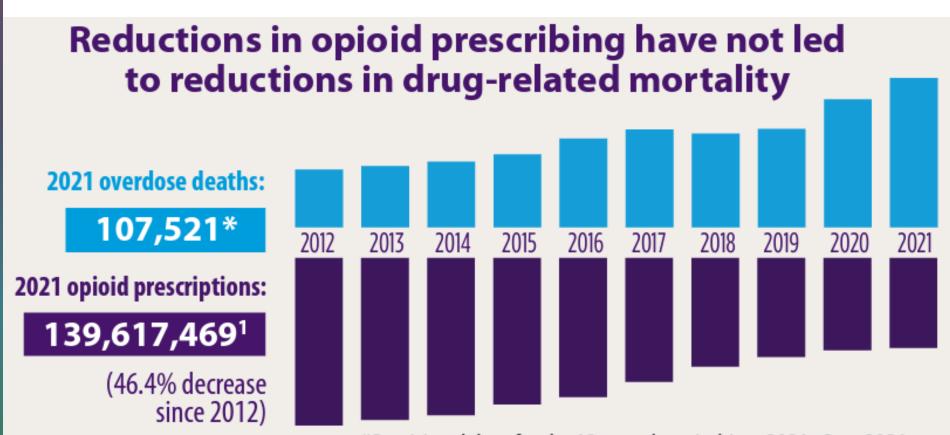


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\*Provisional data for the 12-month period Jan. 2021—Dec. 2021 https://www.cdc.gov/nchs/nvss/vsrr/drug-overdose-data.htm

Source: https://www.ama-assn.org/system/files/ama-overdose-epidemic-report.pdf



# **Opioid Prescribing Rates & Overdose Deaths**

Prescribing Rates (per 100 people)

2018
2019
57.1
51.4
48.4

Office of National Drug Control Policy
(ONDCP)

Non-Fatal Opioid Overdose Tracker

Opioid Overdose Death Rate Per 100,000 Population

UT 2022

13.6

US 2022

24.8

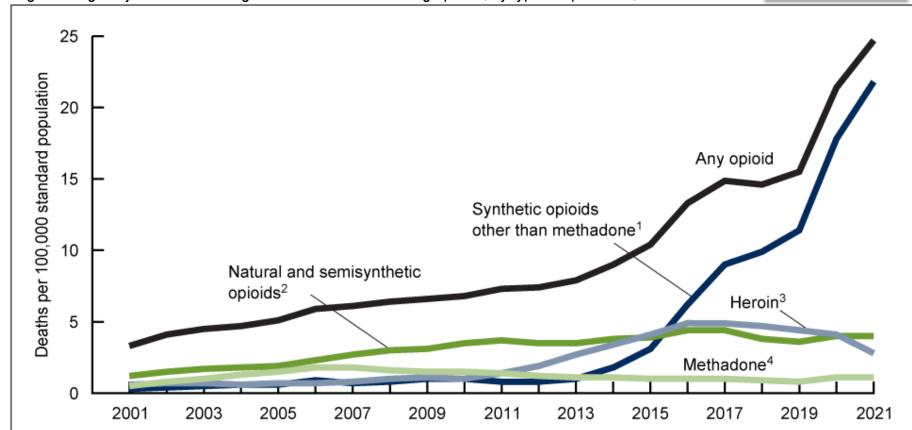
https://www.cdc.gov/drugoverdose/rxrate-maps/
https://www.kff.org/state-category/health-status/opioids/



# OPIOID OVERDOSE DEATHS BY TYPE OF OPIOID



Figure 4. Age-adjusted rate of drug overdose deaths involving opioids, by type of opioid: US, 2001-2021



Source: https://www.cdc.gov/nchs/images/databriefs/451-500/db457-fig4.png



## PDMP: Prescription Drug Monitoring Program

	<ul> <li>Utah Controlled Substance Database Program (CSD)     https://dopl.utah.gov/csd/index.html</li> </ul>
General	<ul> <li>Administered by the Division of Occupational and Professional Licensing</li> </ul>
	Schedule II-V are monitored
	<ul> <li>Dispensers and prescribers are required to register and input data</li> </ul>
	<ul> <li>Before prescribing, there is an obligation to review under certain circumstances</li> </ul>
	<ul> <li>Prescribers can authorize a registered delegate</li> </ul>
	<ul> <li>Must be entered into PDMP on a daily basis after dispensing</li> </ul>
	<ul> <li>Unsolicited reports/alerts are sent to prescriber, dispensers, and law enforcement</li> </ul>
Reporting	<ul> <li>Utah does share data with other states' PDMP</li> </ul>
	<ul> <li>Out-of-state pharmacies are not required to report to the patient's home state</li> </ul>

• 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 Paguirements Initial prescribing limits for acute pain: 7-day supply

	Physician	PA	Advanced Practice Nurse
Prescriber Status	Licensed	Schedule II-V	Schedule II-V
Education Requirements	4 hrs./2 yrs	4 hrs./2 yrs.	4 hrs./2 yrs.

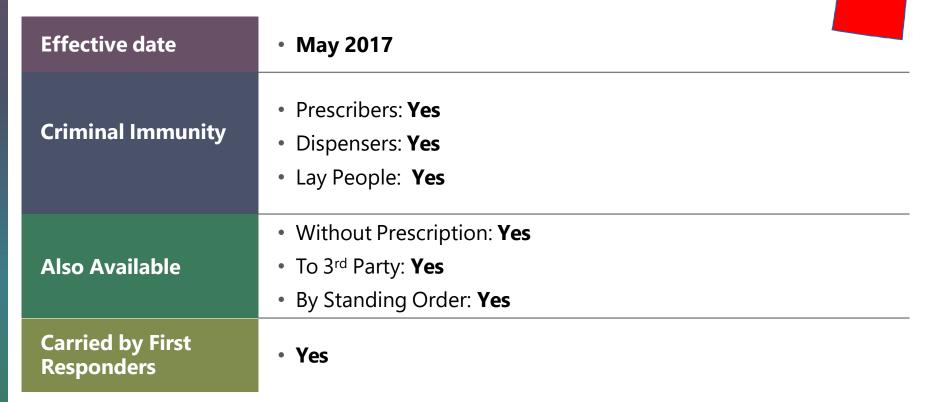
The Medication Access and Training Expansion (MATE) Act requires new or renewing Drug Enforcement Agency (DEA) registrants, as of June 27, 2023, to have completed a total of at least eight hours of training on opioid or other substance use disorders. This course meets the criteria outlined by Substance Abuse and Mental Health Services Administration (SAMHSA) to count toward this training requirement.

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/

https://www.asam.org/education/dea-education-requirements



## **Naloxone Regulation**



On March 29, 2023, FDA announced approval of Narcan (naloxone hydrochloride) Nasal Spray (NNS) for use as a nonprescription opioid overdose reversal agent. OTC NNS commercially available Sept 2023.

Other naloxone products will remain prescription drugs.

<u>State Naloxone Access Rules and Resources - SAFE Project, January 2023</u>
<a href="http://legislativeanalysis.org/wp-content/uploads/2023/02/Naloxone-Access-Summary-of-State-Laws.pdf">http://legislativeanalysis.org/wp-content/uploads/2023/02/Naloxone-Access-Summary-of-State-Laws.pdf</a>
<a href="https://www.thefdalawblog.com/2023/03/2023-is-the-year-for-otc-naloxone">https://www.thefdalawblog.com/2023/03/2023-is-the-year-for-otc-naloxone</a> 3/30/2023



## MATE ACT AND STATE REQUIREMENTS

#### **MATE Act**

As of June 27, 2023, DEA registrants are to have completed a total of at least 8 hours of training on treatment and management of patients with opioid or other substance use disorders. This activity meets the criteria outlined by SAMHSA to count toward this training requirement.

### **State Requirements**

This course also meets many states' requirements for pain education.





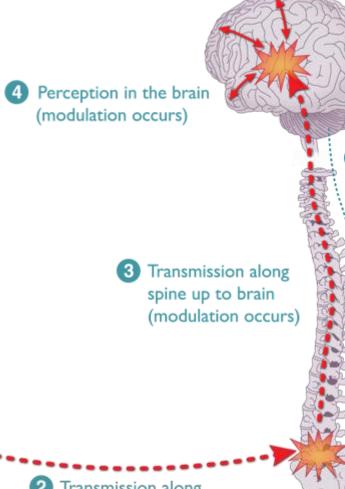
#### THE NEUROMECHANISMS OF PAIN

## Peripheral Pain Modulators:

- Histamines
- Prostaglandins
- Cytokines
- Bradykinin
- Substance P

Injury

Others



Descending

Descending pathway

(down regulation)

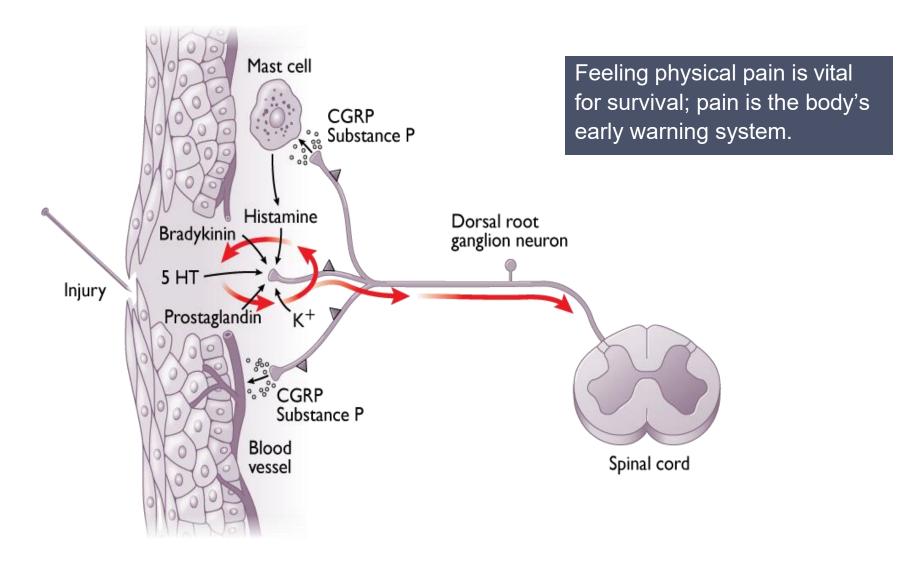
## Neurotransmitters:

- Serotonin
- Norepinephrine
- Endogenous opiates
- Others



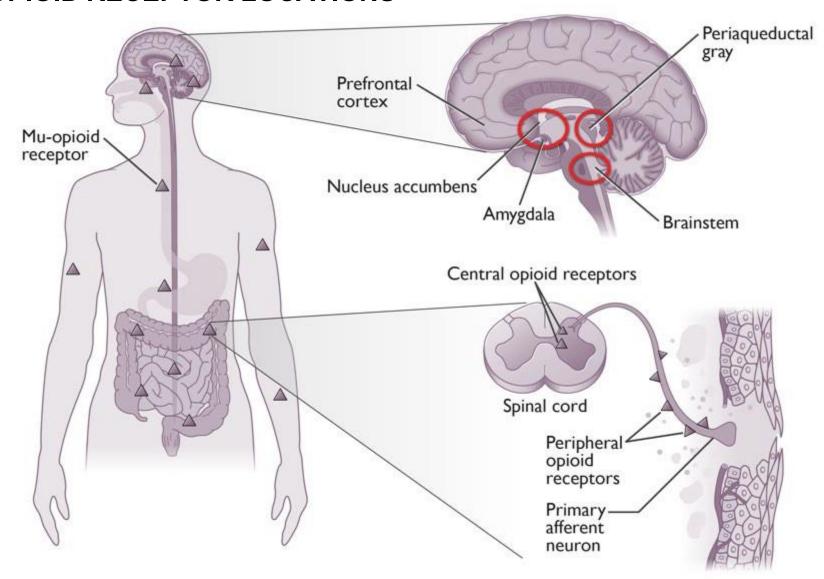


#### **MEDIATORS OF PERIPHERAL NOCICEPTION**





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

#### **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

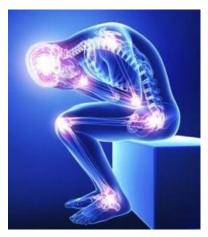


Post-herpetic neuralgia, trigeminal neuralgia, distal polyneuropathy, CRPS, neuropathic low back pain

## **MIXED TYPES** (NOCICEPTIVE / **NEUROPATHIC)**

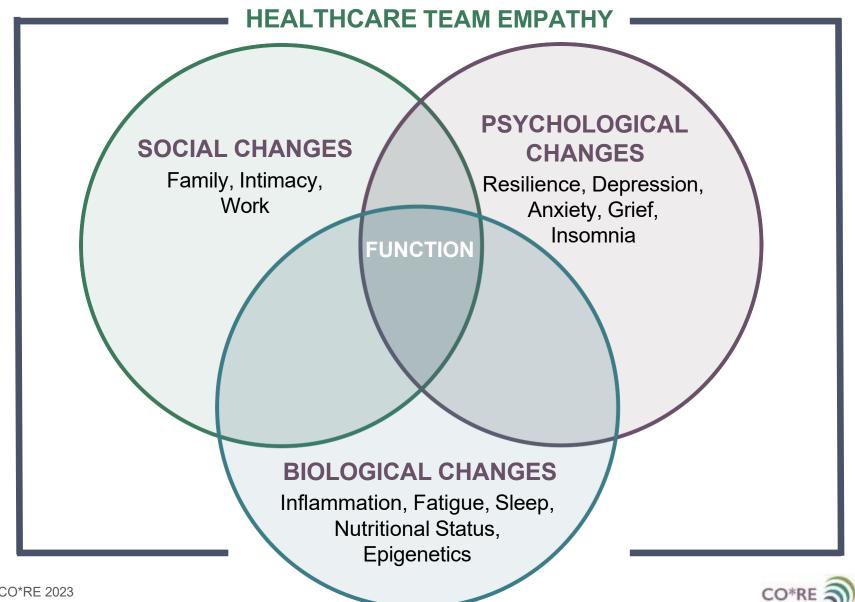


Primary injury and secondary effects

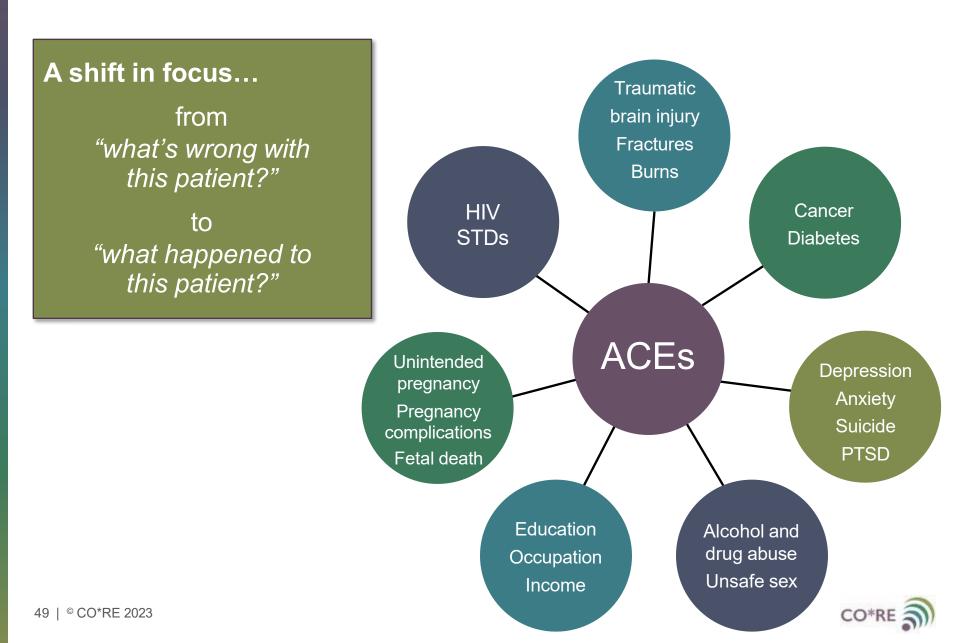


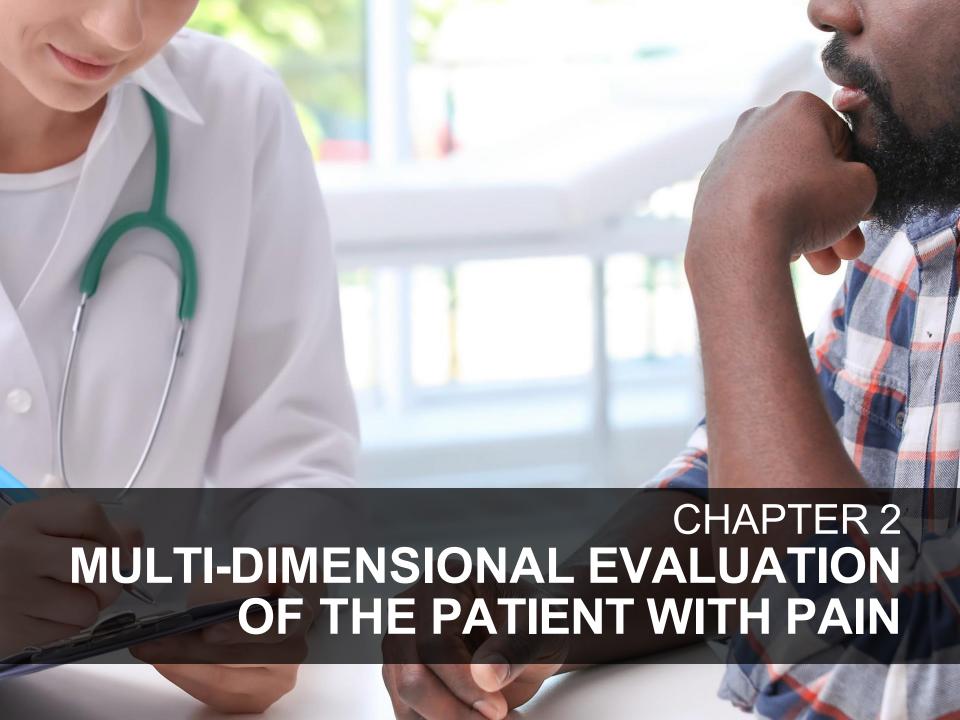


#### THE EXPERIENCE OF PAIN: A BIOPSYCHOSOCIAL MODEL



## **ADVERSE CHILDHOOD EXPERIENCES (ACEs)**





#### **HOW DO WE INITIATE DISCUSSION WITH A PATIENT?**

Ask permission: "Is it okay if I ask you about alcohol or drugs?" Reframe your approach to avoid use of stigmatizing terms:

TERMS TO AVOID	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



#### HISTORY OF PRESENT ILLNESS

Scan to view CO\*RE Tools



PRE-SCREENERS COLLECTED IN ADVANCE (PHQ-2/9, BPI)

#### **DESCRIPTION OF PAIN**







Quality





Variations/
patterns/rhythms

WHAT RELIEVES THE PAIN?

WHAT CAUSES OR INCREASES THE PAIN?

PATIENT'S LEVEL OF PAIN AND THE EFFECT OF THE PAIN ON PHYSICAL, EMOTIONAL, AND PSYCHOSOCIAL FUNCTION (eg, PEG, BPI, MPI)

CO\*RE

#### 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 clinicians 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)

#### A NON-PUNITIVE APPROACH TO PRESCRIBING ANALGESIC AGENTS

- Check when initiating opioid therapy regularly when continuing therapy
- Improves patient communication, education, and safety
  - Confirm PDMP information with patient; do not dismiss from care
  - Identify drugs that increase overdose risk when taken together
  - Provide potentially life-saving information and interventions (safety concerns, provide naloxone)
- Discuss safety concerns with other clinicians
- Lowers rates of prescription opioid-related hospitalization and ED visits
- Most PDMPs allow you to appoint a delegate

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

Source: https://www.cdc.gov/opioids/healthcare-professionals/pdmps.html



#### OBTAIN A COMPLETE PSYCHOSOCIAL HISTORY

#### PSYCHOLOGICAL HISTORY

#### Screen for:

- Mental health diagnoses, depression, anxiety, PTSD, current treatments (using PHQ-2, PHQ-9, GAD-7, etc.)
- Alcohol, tobacco, and other drug use
- History of Adverse Childhood Experiences (ACEs) using ACE Questionnaire



Scan to view CO\*RE Tools

Family history of substance use disorder and psychiatric disorders

#### Depression and anxiety can be predictors of chronic pain

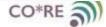
## SOCIAL DETERMINANTS OF HEALTH (SDOH)

### SDOH relate to pain in terms of

- **Economic stability**
- Education access & quality
- Health care access & quality
- Neighborhood & built environment
- Social & community context







Source: https://health.gov/healthypeople/priority-areas/social-determinants-health

#### 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, NY, NY: Oxford Univ. 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 Experiences Questionnaire

ACE

Assessment in Patients Unable to Self-Report

Hierarchy of Pain Assessment or PAINAD



Scan to view CO\*RE Tools

Psychological Measurement Tools (PHQ-2, PHQ-9, GAD-7, etc.)















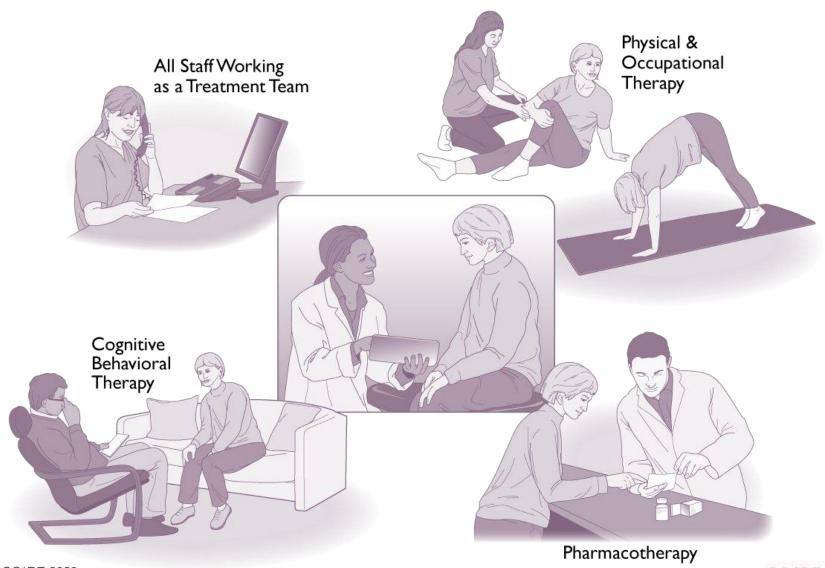




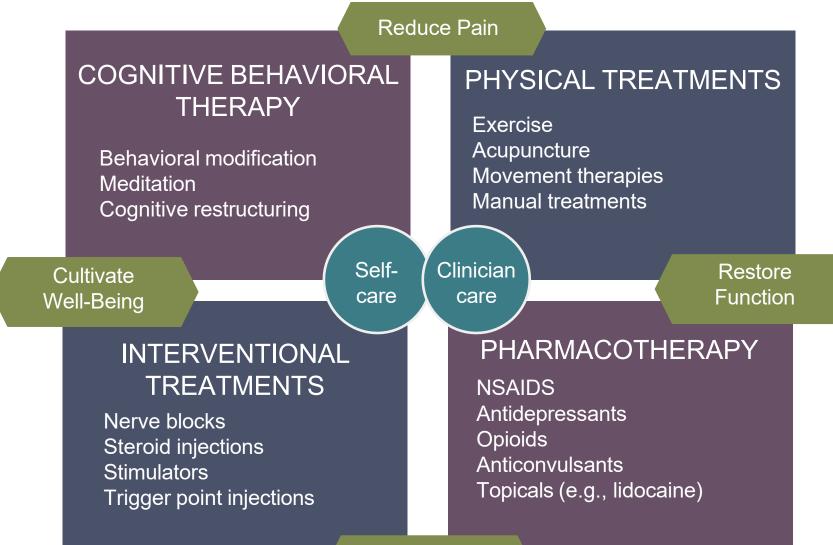
HOW IS PAIN MANAGED?



## 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

Source: https://effectivehealthcare.ahrq.gov/products/noninvasive-nonpharm-pain-update/research



### PHARMACOLOGIC TREATMENTS BY TYPE OF PAIN

## Continue Effective Nonpharmacologic Options First

NOCICEPTIVE / INFLAMMATORY



IR opioids
Nerve blocks
NSAIDs
Topicals and patches

**NOCIPLASTIC** 



Anticholinergic
Anticonvulsants
TCAs and SNRIs
Other serotonin agents



**NEUROPATHIC** 



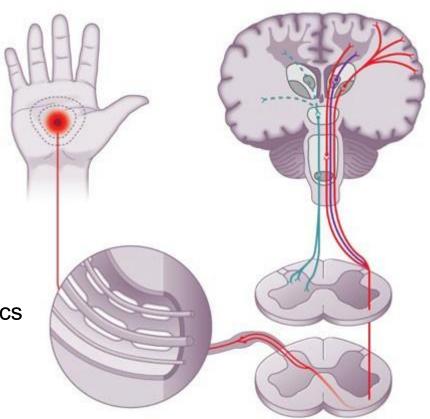
Anticonvulsants
IR and ER/LA opioids
Gabapentinoids
Nerve blocks
TCAs and SNRIs
Transdermal opioids



### POTENTIAL SITES OF ACTION FOR ANALGESIC AGENTS

## Peripherally Mediated Pain:

- Acetaminophen
- Anticonvulsants
- NSAIDs
- Opioids
- Topical anesthetics



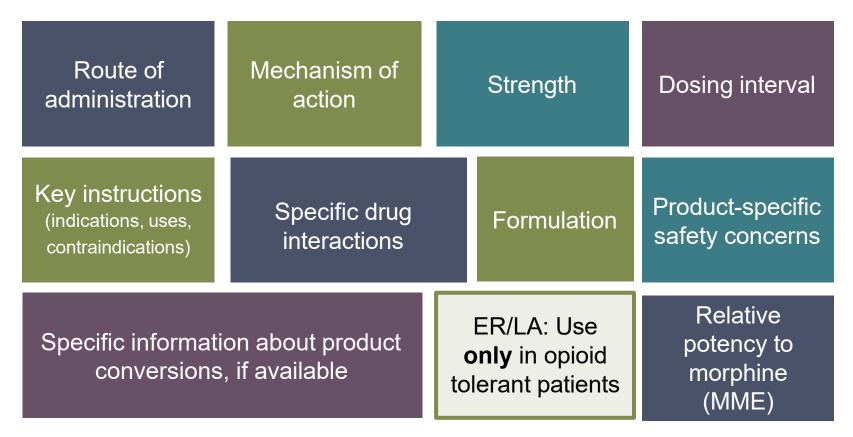
## Centrally Mediated Pain:

- Alpha-2 agonists
- Anticonvulsants
- Ca<sup>+</sup> 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



Opioid product information available at <a href="https://opioidanalgesicrems.com/products.html">https://opioidanalgesicrems.com/products.html</a>

- Immediate Release (IR): rapid onset of analgesia, relatively short duration of effect
- Extended Release/Long-Acting (ER/LA): potentially longer onset of action, longer duration of effect; formulation allows for QD or BID dosing; less frequent dosing

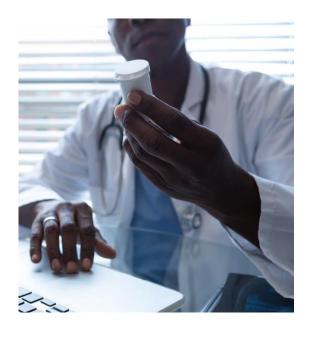


#### WHEN TO CONSIDER A THERAPEUTIC TRIAL OF IR OPIOID

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

Patient has moderate to severe nociceptive or neuropathic pain

Potential benefits are likely to outweigh risks



 CDC Guideline recommendations do not apply to pain related to sickle cell disease or cancer or to patients receiving palliative or end-of-life care (separate guidelines apply to some). There are differences in benefits, risks, and expected outcomes for these patients compared to other patients with chronic pain.

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 & CDC Guideline: https://www.cdc.gov/mmwr/volumes/71/rr/rr7103a1.htm



#### RISKS VERSUS BENEFITS OF PRESCRIBED OPIOIDS

## **POTENTIAL RISKS**

- Life-threatening respiratory depression/overdose, death
- SUD/OUD (assess using ORT-OUD or other validated tool)
- 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 nonopioid analgesics
- Relieves suffering
- May improve function and quality of life



#### **ASSESS RISK FOR OPIOID USE DISORDER**



#### TOOLS FOR PATIENTS CONSIDERED FOR OPIOID THERAPY

**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

**DAST** Drug Abuse Screening Test

**CTQ** Childhood Trauma Questionnaire

**ACEs** Adverse Childhood Experiences

Scan to view CO\*RE Tools





#### A CLOSER LOOK AT THE ORT-OUD

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

Scan to view
ORT-OUD
Video



Source: Cheatle, M., Compton, P.A., et al. J Pain 2019; Jan 26.



### **CATEGORIZATION OF OPIOIDS**

Scan to view DEA Drug Scheduling



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 Methadone Morphine Oxycodone	Buprenorphine Nalbuphine	Naloxone Nalmefene Methylnaltrexone* Naloxogel*

<sup>\*</sup>These represent PAMORA: peripherally-acting mu opioid receptor antagonist



### **OPIOID SIDE EFFECTS AND ADVERSE EVENTS**

SIDE EFFECTS	ADVERSE EVENTS
Respiratory depression	Death
GI effects: dry mouth, nausea/vomiting, opioid-induced constipation (most common; mitigate!)	Addiction
Myoclonus (twitching or jerking)	Overdose
Sedation, cognitive impairment	Hospitalization
Sweating, miosis, urinary retention	Disability or permanent damage
Allergic reactions	Falls or fractures
Hypogonadism	Opioid-induced hyperalgesia
Tolerance, physical dependence	

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 older, cachectic, or debilitated patients
- If given concomitantly with other drugs that depress respiration (such as benzodiazepines\*)
- In patients who are opioidnaïve or have just had a dose increase
- In patients with conditions causing respiratory compromise (eg, obstructive sleep apnea)
- In patients with organ dysfunction

#### **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



#### 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



<sup>\*</sup>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 w/ 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

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 clinician who will ensure appropriate treatment for the baby

Perform universal screening to avoid neonatal opioid withdrawal syndrome (NOWS)

For women using opioids daily, 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





#### PEDIATRIC CONSIDERATIONS

Scan to view

AAP resources



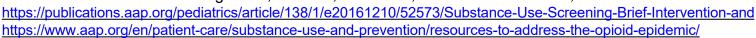
- ❖ HANDLE WITH CARE: JUDICIOUS AND LOW-DOSE USE OF IR FOR BRIEF THERAPY
- ❖ THE SAFETY AND EFFECTIVENESS OF MOST OPIOIDS ARE UNESTABLISHED
  - Pediatric analgesic trials pose challenges
  - Transdermal fentanyl approved in children ≥2 years
  - Oxycodone ER dosing changes for children ≥11 years
- **❖ 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
- ❖ ADOLESCENTS ages 12-21: Identify and treat for OUD (use SBIRT)



Scan to view
SBIRT resource

SBIRT-Screening, Brief Intervention, Referral to Treatment.

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



#### **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 <a href="https://learningipma.org">https://learningipma.org</a>

#### **PATIENT PROVIDER AGREEMENT (PPA)**

#### Reinforce Expectations For Appropriate And Safe Opioid Use

- Clarify treatment plans & 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) & 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 further inquiry: proceed with caution



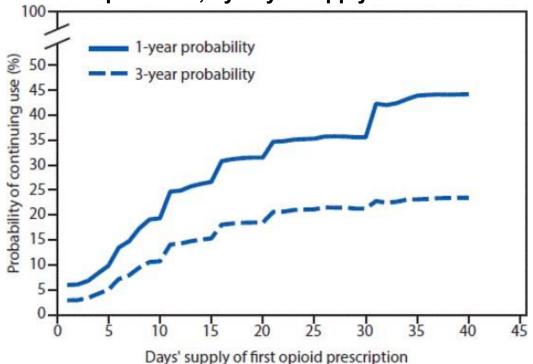


## MANAGING PATIENTS ON OPIOID ANALGESICS

#### **INITIATING IR OPIOIDS**

- Prescribe the lowest effective dose for the shortest period of time based on the individual patient's condition
- 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 stimulant laxative

### One- and 3-year probabilities of continued opioid use, by days' supply of first Rx



- 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

Source: https://www.cdc.gov/mmwr/volumes/66/wr/mm6610a1.htm



#### ONGOING MANAGEMENT OF PATIENTS ON OPIOID ANALGESICS

#### PERIODIC, CONTINUAL ASSESSMENT

- Is the patient making progress toward functional goals?
- Reassess to identify the underlying source of pain
- Reset goals if required or indicated; develop reasonable expectations
- Ask if patient is willing to engage with other modalities
- Monitor for breakthrough pain or comorbid conditions that may arise
- Review adverse events/side effects at each visit
  - Evaluate bowel function
  - Screen for endocrine function as needed
  - Implement opioid rotation, as indicated



## ONGOING MANAGEMENT OF PATIENTS ON OPIOID ANALGESICS (cont.)

#### 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

#### CONSIDERATIONS FOR TREATMENT MODIFICATION

- Continue IR
- Taper and discontinue (when opioid therapy is no longer necessary)
- Transition to ER/LA



#### TRANSITIONING FROM IR TO ER/LA OPIOID OPTIONS

#### **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 SELECTION IS CRITICAL

Some ER/LA opioids or dosage forms are only recommended for opioid tolerant patients (ER/LA in opioid-naïve patients is controversial)

- ANY strength of transdermal fentanyl
- Certain strengths/doses of other ER/LA products (check drug prescribing information)
- Consider transition to buprenorphine (patch, film)

## 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



#### **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
- 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



**Practice** Transition an 80 y/o patient from morphine 180 mg/day to oxycodone

	EQUIANALGESIC DOSE		USUAL STARTING DOSE	
DRUG	SC/IV	PO	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

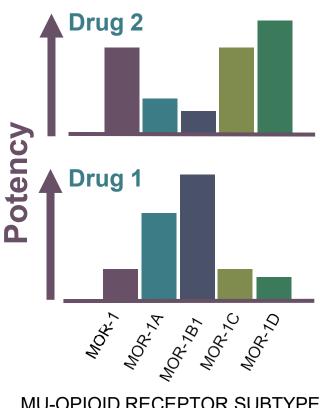
#### MANY MU RECEPTOR SUBTYPES

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

#### **MAY HELP EXPLAIN:**

Interpatient variability in response to muopioids

Incomplete cross tolerance among mu-opioids



MU-OPIOID RECEPTOR SUBTYPE



#### **GUIDELINES FOR OPIOID ROTATION**



Transition an 80 y/o patient **Example:** from morphine 180 mg/day to oxycodone

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 an older adult 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:

- Do not give methadone to opioid-naïve patients
- 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; EKG monitoring controversial

## IF SWITCHING TO **BUPRENORPHINE**:

Consider cross-taper with buccal film or transdermal patch; see guidelines for switch to higher dose

## IF SWITCHING TO TRANSDERMAL FENTANYL:

Calculate dose conversion based on equianalgesic dose ratios included in the drug package insert

SOURCES: <a href="https://pubmed.ncbi.nlm.nih.gov/31917418/">https://www.pbm.va.gov/PBM/AcademicDetailingService/</a>
Documents/Academic\_Detailing\_Educational\_Material\_Catalog/IB\_1497\_Provider\_BupChronicPain.pdf
<a href="https://accpjournals.onlinelibrary.wiley.com/doi/full/10.1002/phar.2676">https://accpjournals.onlinelibrary.wiley.com/doi/full/10.1002/phar.2676</a>, CDC 2022 Guideline for Prescribing Opioids for Pain,
<a href="https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4078896/">https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4078896/</a>





#### **GUIDELINES FOR OPIOID ROTATION: SUMMARY**





Transition an 80 y/o patient **Example!** from morphine 180 mg/day to oxycodone

**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 OPTIMIZING**

- 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

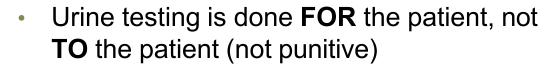
Overdose is still possible if taken orally in excessive amounts

These products are expensive with no generic equivalents



#### **URINE DRUG TESTING (UDT)**







Scan to view
Urine Drug
Screen Video

- 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**





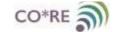
	SCREENING (Office-based)	CONFIRMATORY (Send to lab)
Analysis technique	Immunoassay	GC-MS or HPLC
Sensitivity (power to detect a class of drugs)	Low or none when testing for semi-synthetic or synthetic opioids	High
Specificity (power to detect an individual drug)	Varies (can result in false positives 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



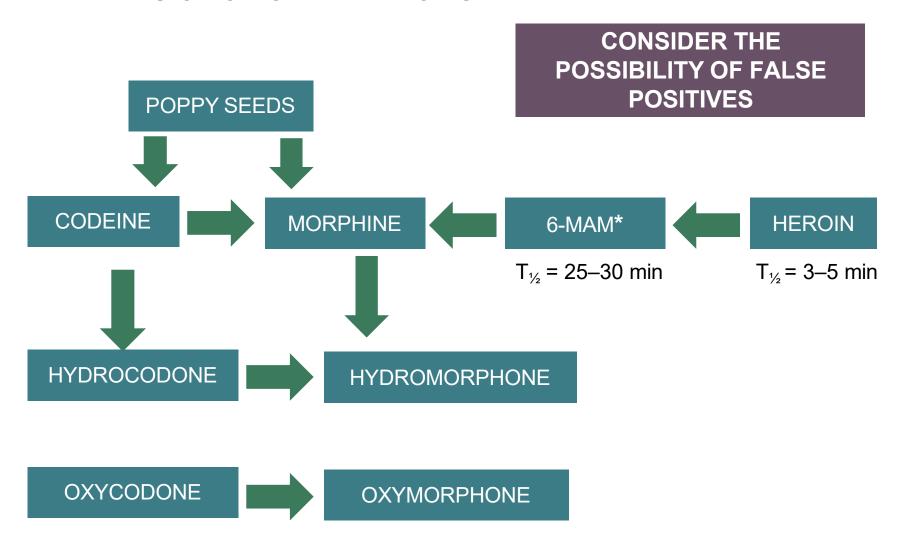
#### WINDOWS OF SPECIFIC DRUG DETECTION

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**



\*6-MAM = 6-Monoacetylmorphine



#### **CONSIDERATIONS FOR RE-EVALUATING OPIOID USE**

THERAPEUTIC GOALS ARE ACHIEVED 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



Scan to view
CO\*RE Tools



#### PATIENT-CENTERED APPROACH TO TAPERING

#### No single approach is appropriate for all patients

- Discontinue through a taper schedule
- If OUD suspected:
  - Begin treatment: Medications for Opioid Use Disorder (MOUD)
  - Consider referral to an addiction or OUD specialist if appropriate
- Consider rotation to partial agonist (e.g., buprenorphine)
- 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**

## **Proper Use**

- Take opioid as prescribed
- If a dose is missed: do not take extra, contact HCP
- Use least amount of medication necessary for shortest time
- Long-term opioid use: avoid abrupt discontinuation, taper safely to avoid withdrawal symptoms

## Monitoring/Side Effects

- Notify HCP if pain is uncontrolled
- Go over all side effects (previous chapter)

## Safety

- Inform HCP of ALL side effects, other meds/supplements taken
- Use caution when operating heavy machinery and driving

Scan to view
Patient Counseling
Guide





## **Storage**

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



## **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
- Sedation, cognitive impairment
- Falls and fractures
- Never share medications with others



## OPIOID-INDUCED RESPIRATORY DEPRESSION

Distribute, dispense, or prescribe naloxone to patient or caregiver. Teach proper administration.

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:**

EMERGENCY

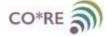
DIAL 911

- 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







### 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 and Nalmefene





## WHERE AND HOW TO DISPOSE OF UNUSED OPIOIDS





## **Authorized Collection Sites**

 Use the DEA disposal locator website to find sites near you (QR code to right) or search Google Maps for "drug disposal nearby"



Scan to view disposal locator

## **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

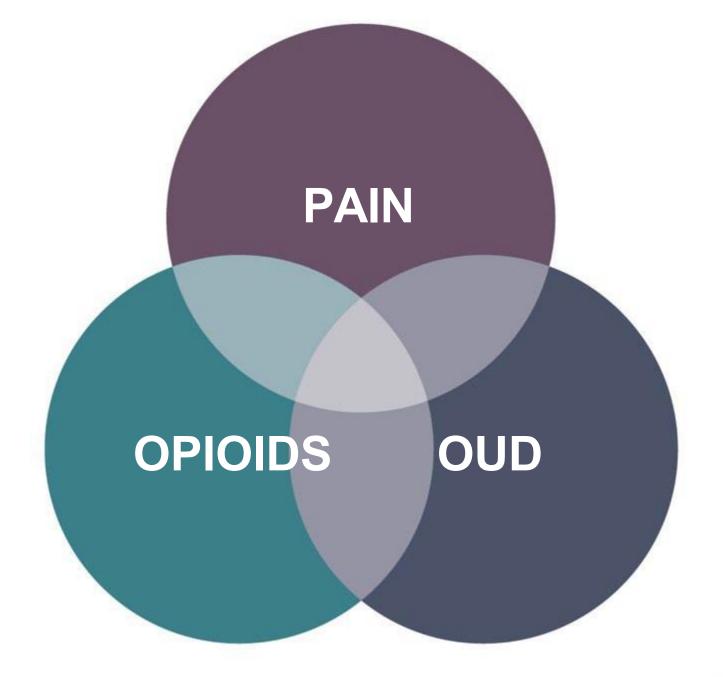
Obtain from authorized collectors

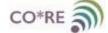


Sources: FDA. Where and How to Dispose of Unused Medicines. <a href="https://www.fda.gov/consumers/consumer-updates/where-and-how-dispose-unused-medicines">https://www.fda.gov/consumers/consumer-updates/where-and-how-dispose-unused-medicines</a>; EPA. How to Dispose of Medicines Properly. <a href="https://archive.epa.gov/region02/capp/web/pdf/ppcpflyer.pdf">https://archive.epa.gov/region02/capp/web/pdf/ppcpflyer.pdf</a>









## WHAT IS ADDICTION?



## **Practical Definition:**

Addiction is the continued use of drugs or activities, despite knowledge of continued **harm** to oneself or others.

## Official ASAM Definition:

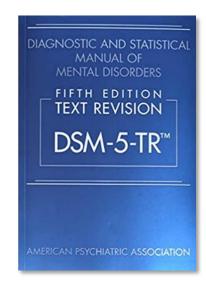
Addiction is a treatable, chronic medical disease involving complex interactions among brain circuits, genetics, the environment, and an individual's life experiences. People with addiction use substances or engage in behaviors that become compulsive and often continue despite harmful consequences. Prevention 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
- 5. 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  $\geq 6$  = severe

Not valid if opioid is taken as prescribed



## **WORDS MATTER – PEOPLE MATTER**

Physical dependence or tolerance

Doesn't necessarily equal tolerance

OUD/ Doesn't necessarily equal addiction

OUD/ Doesn't necessarily equal behavior





## **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:

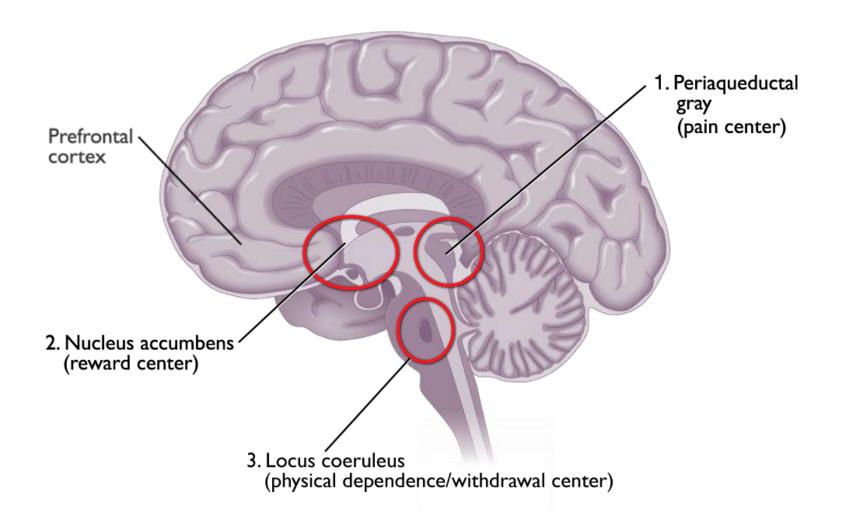
- High dosages
- Prolonged use
- Low hedonic tone
- Mental health disorders
- Past history of substance use disorder

Clinical judgment is key.

Source: Chou R, et al. Ann Intern Med. 2015;162:276-86

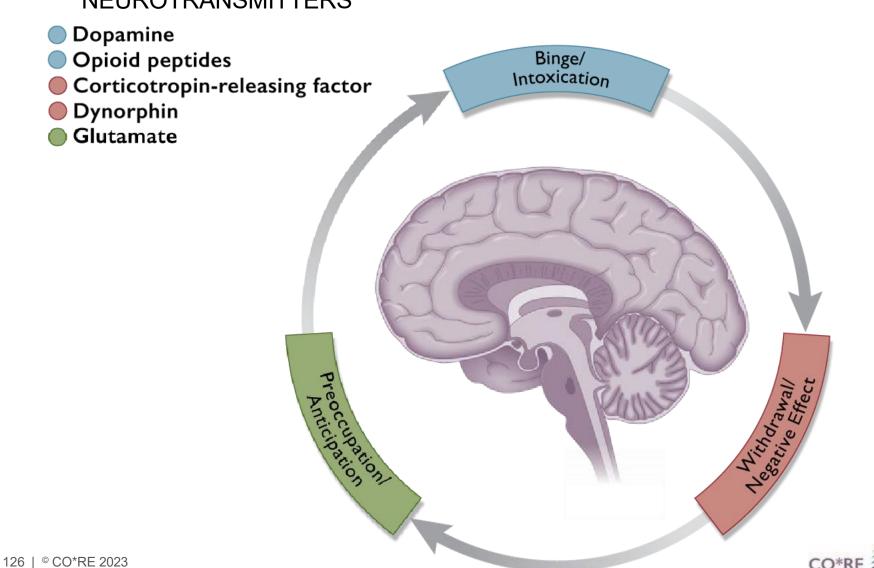


## OPIOID RECEPTORS IN THE BRAIN: RELATIONSHIP TO ANALGESIA, OUD, AND WITHDRAWAL



## THE CYCLE OF SUBSTANCE USE DISORDER

## **NEUROTRANSMITTERS**



## 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 you can save a life!

## There are three medication options:

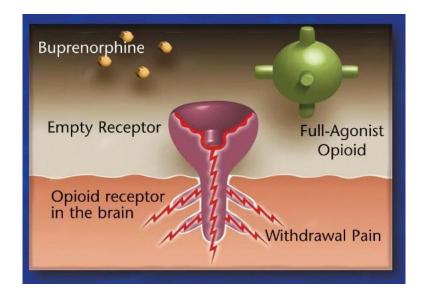
- 1. Buprenorphine (Schedule III)
- 2. Methadone (Schedule II)
- Naltrexone (not a controlled substance)

Are we just replacing one drug with another?

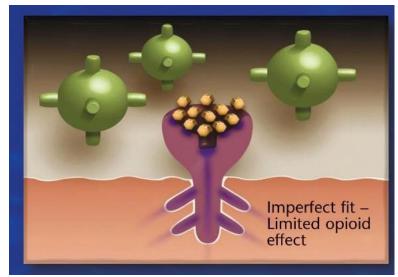
Myth or fact?

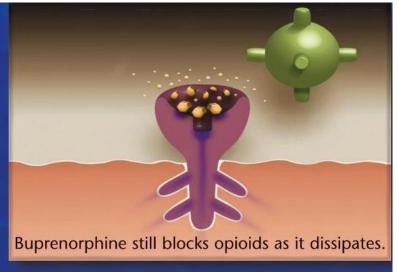


## **HOW BUPRENORPHINE WORKS**









Source: https://www.naabt.org/education/images/Receptors\_HiRes.jpg, https://pubmed.ncbi.nlm.nih.gov/16547090/



## BUPRENORPHINE

- Most commonly prescribed pharmacotherapy for treatment of OUD
- Good efficacy and safety profile
- "Plateau effect" for respiratory depression
- Congress eliminated the X-waiver requirement to prescribe Bup.
- All DEA-licensed HCPs can prescribe without patient number caps
- Long-acting injectable 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

Source: https://pubmed.ncbi.nlm.nih.gov/16547090/



## 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)
- Evaluate for SUD 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.



## **RESOURCES TO HELP YOU TREAT OR TO REFER:**

## TREATMENT SUPPORT

## SAMHSA -

Training Materials & Resources



## REFERRAL SUPPORT

ASAM -

Physician Finder



### NIDA -

**Treatment Resources** 



### SAMHSA -

**Find Treatment** 



## PCSS -

Providers Clinical Support System



### AAAP -

Specialist Finder





## **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
- a If you suspect OUD, begin treatment



## Please complete your post-test 🤗



Complete the brief post-test for CE/CME credit Your participation helps the FDA reach its goals for REMS education





## THANK YOU! WWW.CORE-REMS.ORG





FULL LIST OF SOURCES AVAILABLE UPON REQUEST