















## C Assessing Acute Pain

"Many factors influence self-reported pain ratings including gender, social support, provider characteristics, trust." ~ *Barry S. Oken, MD, PhD* 

- Intensity
- Location
- Onset
- Duration
- Radiation
- Exacerbation
- Alleviation

Oken BS. Brain. 2008 Breivik H, et al. Br J Anaesth. 2008 Mason ST, et al. Handbook of Pain Assessment. 2011





## © Predicting Risk of Progression to Chronic Pain

### **Example: STarT Back Screening Tool**

						Disagree	Agre
ı	My back pain has	spread down my	v leg(s) at some time	e in the last 2 week	s		
2	I have had pain in	the shoulder or a	neck at some time in	n the last 2 weeks			
3	I have only walke	d short distance	s because of my bac	k pain			
4	In the last 2 weeks	s, I have dressed	more slowly than u	sual because of ba	ck pain		
5	It's not really safe	for a person with	a condition like mi	ine to be physically	active		
6	Worrying though	nts have been goin	ng through my mind	a lot of the time			
7	I feel that my bac	k pain is terrible	and it's never goin	ng to get any bette	r		
8	In general I have I	not enjoyed all th	e things I used to en	njoy			
			baak nain baan in th	ne last 2 weeks?			
9.	Overall, how <b>both</b> Not at all	slightly	Moderately	Very much	Extre	mely	

Helps identify modifiable risk factors (biomedical, psychological and social) for chronic back pain disability

Total score stratifies patients into low, medium or high risk



urgery	Over-prescription	Reference
Thoracic	71% taking half or less	Bartels et al. Plos One. 2016
C-section	83% taking half or less	Bartels et al. Plos One. 2016
Upper extremity	77% taking half or less	Rodgers et al. J. Hand Surg. 2012
Upper extremity	84% had leftover pills	Kim et al. <i>JSBS</i> . 2016
Dermatologic	86% had leftover pills	Harris et al. JAMA Derm. 2013
General surgery	71% of pills not taken	Hill et al. Annals Surgery. 2016
Urologic	42% of pills not taken	Bates et al. J. Urology. 2011





Analgesic(s)	Dose (mg)	NNT vs Placebo ≥ 50% maximum pain relief over 4-6 hours	~50,000 participants
SINGLE AGENTS:			~460 high quality studio
Ibuprofen	600	2.7	~460 high-quality studies
Naproxen	500	2.7	(mostly dental extractions)
Celecoxib	400	2.6	
Acetaminophen (APAP)	1000	3.6	
Oxycodone	15	4.6	
Codeine	60	12.0	
Gabapentin	250	11.0	
COMBINATIONS:			
Ibuprofen + APAP	400+1000	1.5	
Ibuprofen + oxycodone	400+5	2.3	
APAP + oxycodone	325+5	5.4	
APAP + codeine	300+30	6.9	





	40 year old female	
- (14)	Current Medications: Metformin 1000 mg 2x/day	
CASE STUDY	Current Pain Medications: Oxycodone 10 mg 4x/day (60 MME*) Gabapentin 300 mg 3x/day	
Kathy James	<b>Previous Pain Medications:</b>	
	NSAIDs (ibuprofen, naproxen)	Inadequate pain relief and GI upset
	Acetaminophen	Inadequate pain relief
	Tricyclic antidepressants (TCA) (amitriptyline)	Inadequate pain relief and dry mouth
	Serotonin-norepinephrine reuptake inhibitor (SNRI) (venlafaxine)	Unable to tolerate due to nausea and dizziness
	Tramadol	Inadequate pain relief
	Acetaminophen with codeine	Inadequate pain relief and nausea
	*Morphine Mg Equivalents	19



### 40 year old female

#### **Social History:**

Receptionist law office 20 hours/week Married husband manages hardware store Children 6, 8 and 12 years of age

#### **Substance Use History:**

Alcohol use 1-2 glasses of wine on some weekends and holidays Tried cannabis in high school No recent history of illicit drug use Smokes tobacco 1 pack per day for the past 20 years

#### **Family History:**

Mother died from complications of alcoholic cirrhosis



#### $\mathbf{\overline{C}}$ Acute versus Chronic Pain Acute Pain **Chronic Pain** Can be a disease in itself Live sustaining symptom Adaptive by eliciting motivation to Maladaptive, pathologic, disorder minimize harm and allow healing of the somatosensory pain signaling pathways influenced by genetic and epigenetic factors - Nociceptive pain: somatic or visceral pain caused by ongoing activation of nociceptors to a noxious stimulus (e.g., injury, disease, inflammation) - Neuropathic pain: nerve injury or impairment causing aberrant signal processing in the \* Petrosky E, et al. Ann Intern Med. 2018 peripheral or central nervous system Ilgen MA, et al. JAMA Psychiatry. 2013 Tang NK et al. Psychol Med. 2006 Associated with higher risk of fatal Dzau VJ, Pizzo PA. JAMA. 2014 and nonfatal suicide attempts\* Walk D, Poliak-Tunis M. Med Clin N Am. 2016 Argoff CE, et al. Pain Med. 2009





























## © Opioid Analgesics

- Turn on descending inhibitory systems
- Prevent ascending transmission of pain signal
- · Inhibit terminals of C-fibers in spinal cord
- Inhibit activation of peripheral nociceptors
- Vary by response (not all patients respond to the same opioid in the same way)
  - >3,000 polymorphisms in human MOR gene
  - Single nucleotide polymorphisms (SNPs) identified that affect opioid metabolism, transport across the blood brain barrier, and activity at receptors and ion channels

#### Activate the reward pathway

McCleane G, Smith HS. *Med Clin N Am.* 2007 Smith HS. *Pain Physician*. 2008 Ren Z et al. *Pain Physician*. 2015













# © Patient-related Risk Factors: Overdose and Addiction

Patient-related Factors	R	lisk
Mental health disorder (e.g. depression, anxiet	y) Overdose	Addiction
Substance use disorder (e.g., alcohol, tobacco, illicit and prescription dr	ug) Overdose	Addiction
Family history of substance use disorder		Misuse
Adolescent		Addiction
Age <45		Misuse
Age >65	Overdose	
Sleep-disordered breathing	Overdose	
Legal history (e.g., DUI, incarceration)		Misuse
History of sexual abuse		Misuse
History of overdose	Overdose	
ves J, et al. BMC Health Serv Res. 2006 Reid MC, et	t al. J Pain Symptom Manage. 200 al. J Gen Intern Med. 2002 et al. N Engl J Med. 2016	4

Condition	Prevalence	References	
	Patients with Chronic Pain		
Depression	33 - 54%	Cheatle M, Gallagher R. 2006.	
	33 3470	Dersh J, et al. 2002.	
	16.5 - 50%	Knaster P, et al. 2012.	
Anxiety Disorders		Cheatle M, Gallagher R. 2006.	
Personality Disorders	31 - 81%	Polatin PB, et al. 1992.	
Personality Disorders		Fischer-Kern M, et al. 2011.	
	49% veterans	Otis, J, et al. 2010.	
PTSD	2% civilians	Knaster P, et al. 2012.	
whatewas Use Discussion	45 200/	Polatin PB, et al. 1992.	
Substance Use Disorders	15 - 28%	Cheatle M, Gallagher R. 2006.	

























### 2 weeks later

#### In the interim...

- Unable to contact previous PCP who retired and moved out of state
- UDT positive for oxycodone

#### **Office Visit 2**

- PEG (Pain, Enjoyment, General activity) "6 out of 10" but sometimes a "10 out of 10" before next dose
- She denied sedation
- Completed her 2 week prescription for oxycodone/APAP on schedule



www.fda.gov/downloads/Drugs/DrugSafety/InformationbyDrugClass

## **(C**) **Opioids**

## **IR/SA** Opioids

- No opioid tolerance/opioid naïve
- Intermittent or occasional pain (PRN dosing)

## **ER/LA** Opioids

- Opioid tolerance exists
- Constant, severe, around-the-clock pain (scheduled dosing)
- To stabilize pain relief when patient using multiple doses of IR/SA opioids
- Note: No adequate studies of ER/LA opioids in pregnant women; use only if benefit significantly outweighs risk.
- MUST NOT be broken, chewed or crushed

## Always start low and go slow

**CDC Recommendation 4:** When starting opioids, use immediate-release

instead of extended-release opioids.

CDC

Dowell D, et al. MMWR. 2016.



**(C**) **IR/SA vs ER/LA Opioid Uncertainties** Insufficient Individualize Debate Evidence Treatment to determine whether whether bolus dosing **Choose options** ER/LA opioids are (IR/SA) or continuous that best meet more effective or safer exposure (ER/LA) is patient's needs than IR/SA opioids more likely to result in analgesic tolerance, hyperalgesia or addiction

Chou R, et al. J Pain Symptom Manage. 2003 Argoff CE, Silvershein DI. Mayo Clin Proc. 2009

## © Transdermal Preparations

- Convenient dosing
- Slow peak onset (>24-72h)
- Delayed offset (serum t½ life >17-26h)
- Sustained release requires predictable blood flow and adequate subcutaneous fat
- Absorption is increased with fever or broken skin
- · Absorption is decreased with edema
- Some with metal foil backing and not compatible with MRI

#### Fentanyl

- Every 72 hours
- Dosages available (mcg/hr): 12, 25, 37.5, 50, 62.5, 75, 87.5, 100

#### Buprenorphine

Every 7 days
Dosages available (mcg/hr): 5, 7.5, 10, 15, 20

#### 61

## © Methadone is Different

- The problem...potentially the most dangerous opioid
- Long, variable, unpredictable half-life
  - Analgesia 6-8 hours
  - Serum t½ 20-120 hours
- QTc prolongation, risk of torsades de pointes

#### Some possible advantages:

- NMDA receptor antagonist
  - Potentially less analgesic tolerance, better efficacy in neuropathic pain
- No active metabolites
- Inexpensive, small dosage units (5mg tablets)

Fredheim OM, et al. Acta Anaesthesiol Scand. 2008 Chou R, et al. J Pain. 2014

62















## **Special Populations – Liver Disease**

- Opioid clearance is decreased with hepatic insufficiency
  - Increased risk of hepatic encephalopathy
- Fentanyl likely safe with modest hepatic dysfunction
- Morphine, oxycodone, and hydromorphone require reduced doses and prolonged dosing intervals
- Codeine effects are difficult to predict, and alternatives should be considered
- Tramadol may be safe but experience is limited
- Nonopioids in advanced liver disease
  - Acetaminophen: limit to 2 g/d
  - NSAIDs: increased risk of variceal bleed, impaired renal function, and diuretic-resistant ascites
  - Gabapentin, pregabalin, and nortriptyline can be used but lower doses

Hamilton JP, et al. UpToDate. Dec 2018



Davison SN. UpToDate. Dec 2017




# © Common Universal Precautions

- Comprehensive pain assessment
- Formulation of pain diagnosis(es)
- Opioid misuse risk assessment
- Opioid prescriptions should be considered a test or trial; continued, modified or discontinued based on assessment and reassessment of risks and benefits (e.g. every 1-3 months)
- Prescription should include maximum number of tablets that can be taken per day
- Regular face-to-face visits including periodic review of all medications
- Co-prescribe naloxone
- Clear documentation of your assessment and plan

Federation of State Medical Boards *Model Policy* April 2017. Chou R, et al. *J Pain*. 2009 Gourlay DL, et al. *Pain Med.* 2005 Franklin GM. *Neurology*. 2014



# © PPA Informed Consent

### **Realistic Goals**

- Reduce pain, not eliminate
- Increase function (individualized and SMART goals)
  - Specific
  - Measureable
  - Action-oriented
  - Realistic
  - Time-sensitive

#### CDC Recommendation 2 and 3:

Before starting opioids establish realistic goals. Continue opioids only if meaningful improvements outweighs risks. Discuss risks and benefits of opioids. Dowell D, et al. MMWR. 2016

## **Potential Risks**

- Side effects, physical dependence
- Drug interactions
- Over-sedation and potential for impairment (esp. during dose adjustments); e.g. driving risk
- Misuse, overdose, death
- Pregnancy and risk of Neonatal Opioid Withdrawal Syndrome
- Possible hyperalgesia (increased pain)
- Victimization by others seeking opioids

 Nicolaidis C. Pain Med. 2011
 Cheatle MD, Savage SR. J Pain Symptom Manage. 2012
 Mailis-Gagnon A, et al. Clin J Pain. 2012

 Paterick TJ, et al. Mayo Clin Proc. 2008
 Tolia VN, et al. N Engl J Med. 2015
 Schumacher MB, et al. Pschopharm. 2017
 76

# © PPA Plan of Care

- · Engagement in other recommended treatments
- Polices and procedures monitoring, refills
- Medication management
- Use exactly as directed in pharmacy Medication Guide (no adulteration of ER/LA pills, patches, buccal film; guidance on missed doses)
- No illegal drug use, avoid sedative use
- Notifying provider of all other medications and drugs, and permission to communicate with key others
- Notifying provider of worsening pain or medication side effects
- Discuss birth control, periodic monitoring for pregnancy
- Safe storage, safe disposal, no sharing or selling

Fishman SM, Kreis PG. Clin J Pain. 2002 Arnold RM, et al. Am J Med. 2006

# Monitoring for Benefits and Harm

Benefit

**(C**)

- Assess and document PEG scores
- Assess and document achievement of SMART goals

#### CDC Recommendation 7:

Evaluate benefits and harms within 4 weeks of starting and at least every 3 months thereafter. Clinically meaningful improvement defined as a "30% improvement" in pain and function.

Dowell D, et al. MMWR. 2016. Ostelo RW et al. Spine 2008

- Harm
  - Current Opioid Misuse Measure (COMM-9)\* self-administered
  - Pill counts (scheduled vs. random)
  - Urine drug tests (scheduled vs. random)
  - Prescription Drug Monitoring Program data
  - History from "reliable" family members
    - Beware of family members with secondary gain for giving inaccurate information

\*McCaffrey SA, et al. Pain Med. 2019

© Face-to-Face Office Visits						
Monitoring for: Six A's Analgesia Activities Adverse effects Aberrant behaviors Affect Adherence	<b>Also review</b> Opioid use using a 24-hour inventory "Tell me how you are taking your medications."					
Passik SD, et al. Clin Ther. 2004	79					











)n	e example from TOPCARE* (Transforming	g Opioid Prescrib	oing in Pri	mary Care) p	orogram:
• ( •   • (	<b>isk Level</b> ORT: Opioid Risk Tool DIRE: Diagnosis, Intractability, Risk, Efficacy SOAPP: Screener & Opioid Assess for Pts w/ F STAR: Screening Tool for Addiction Risk	Face-to- Face Visits/ Pain year	UDT/ year	Pill Counts/ year	PDMP*/ year
•	Low	4	2	2	2
•	Moderate	4	4	4	4
•	High	6	6	6	6

















<ul> <li>CASE STUDY Kathy James</li> <li>40 year old female</li> <li>12DM</li> <li>Chronic, painful, diabetic neuropathy and hip pain</li> <li>ER/LA oxycodone and gabapentin</li> <li>Mow will you</li> <li>Manage this</li> </ul>	<ul> <li>Presents to the ED 1 day after twisting ankle while running to catch a bus</li> <li>Pain severe especially with weight-bearing</li> <li>Took husband's hydrocodone tablet which helped</li> <li>Requesting "narcotics" for ankle pain</li> <li>Ankle exam: swelling, ecchymosis, tenderness over lateral malleolus and decreased range of motion with evidence of mild joint instability</li> <li>X-ray negative for fracture</li> <li>Dx: grade II/III lateral ankle sprain</li> <li>PDMP reviewed: monthly ER/LA oxycodone 20 mg bid #56, last filled</li> </ul>
patient's acute pain?	2 weeks ago











- She then went to the ED of her local hospital,
   requesting early refill of her oxycodone
- ED physician noted that she was in moderate opioid withdrawal and gave her enough ER/LA oxycodone to last until her next PCP appointment in one week
- ED physician left a message with the PCP office regarding patient visit and follow-up plan

#### Post ED Follow-up **History since last visit** Foot pain worse in past month "10 out of 10 most days" Started taking an extra ER/LA oxycodone in the afternoon SCENARIO 1 and ran out early **CASE STUDY** Concerned "body has become used to current dose"; **Kathy James** doesn't seem to work all day anymore Husband says she has become "addicted" Difficult to go to work due to severe pain Trouble sleeping as sheets touching her feet now cause pain 40 year old female Requests increase in her dose T2DM • Chronic, painful, diabetic She is re-educated about the serious risks including death of neuropathy and hip pain self-escalating her opioid dose ER/LA oxycodone and gabapentin









# © Discussing Possible Addiction (OUD)

- Give specific and timely feedback why patient's behaviors raise your concern for possible use disorder (e.g. loss of control, compulsive use, continued use despite harm)
- Remember patients may suffer from both chronic pain and use disorder
- May need to "agree to disagree" with the patient
- Benefits no longer outweighing risks
- "I cannot responsibly continue prescribing opioids as I feel it would cause you more harm than good."
- Always offer referral to addiction treatment

## C Lack or Loss of Benefit

#### What are the next steps?

- Reassess factors affecting pain
- Re-attempt to treat underlying disease and co-morbidities

#### Consider...

- Adding or increasing non-pharmacologic treatment (e.g. acupuncture, CBT)
- Adding or increasing adjuvant medications for synergy
- Adding breakthrough medications
- Opioid rotation





## © Opioid Conversion Tables

- Derived from relative potency ratios using single-dose analgesic studies in opioid-naïve patients
- Based on limited doses or range of doses
- Does not reflect clinical realities of chronic opioid administration
- Are not reliable due to individual pharmacogenetic differences
- Most tables do NOT adjust for incomplete cross-tolerance

Treillet E, et al. *J Pain Res.* 2018 Webster LR, Fine PG. *Pain Med.* 2012 Pereira J, et al. *J Pain Symptom Manage.* 2001











#### $\mathbf{C}$ **Discontinuing Opioids** Do not have to prove addiction or diversion, only assess and reassess the risk-benefit ratio If patient is unable to take opioids safely or is non-adherent with monitoring, then discontinuing opioids is appropriate, even in setting of benefits Need to determine how urgent the discontinuation You are NOT should be based on the severity of the risks and harms abandoning Document rationale for discontinuing opioids the patient, • you are Determine if the opioid needs to be tapered • ABANDONING due to physical dependence THE OPIOID









Patient Themes	Threats to trustworthiness and latrogenic suffering	Clinicians demonstrated lack of care, empathy which caused further suffering
nemes	Communicating the invisible and subjective condition of chronic pain	Clinicians did not accept reports of pain at face value
	Motive, honesty, and testimony	Perceived as untrustworthy by clinicians
	Stigmatized identities	Having chronic pain influences the patients' perceived trustworthiness
linician hemes	Challenges of the practice context	Recalled difficult interactions and the impact these had on their approach to care
	Complicated clinical relationships	Did not see their role as a collaborative partner saw themselves in a defensive role of interrogator

# <section-header><list-item><list-item><list-item><list-item>





	Over the next 12 months
Nº .	Over 6 months she successfully tapered off oxycodone
SCENARIO 3 CASE STUDY	<ul> <li>For the following 6 months, her neuropathic pain was moderately well controlled on combination of nortriptyline 25 mg at night, gabapentin 800 mg tid and capsaicin cream 3-4 times per day</li> </ul>
Kathy James	<ul> <li>Joined a monthly chronic pain support group</li> </ul>
	<ul> <li>PEG scores remained between 4-5/10 (patient states she is surprised her pain is better off oxycodone)</li> </ul>
	Remained employed
	Remained adherent with the treatment plan and monitoring
	Continued with regularly scheduled follow up visits
	125





# © Discussing Possible Diversion

- Prescription drug diversion is one form of opioid misuse and is defined as the giving, selling, or trading prescription medications
- Discuss why you are concerned about diversion
   e.g. UDT negative for prescribed opioid, nonadherence with pill counts
- Discuss your inability to continue to prescribe opioids if the opioids are being diverted to others













## © Patient on MOUD and Pain Management

#### Patient with OUD and Chronic Pain

- Certain buprenorphine formulations (e.g. sublingual) can be prescribed in officebased settings for treating both chronic pain (off label) and OUD (requires special qualifications under DATA 2000)
- Buprenorphine's analgesic half-life is 8-12 h versus 24 h for treating OUD
- Systematic review of 10 studies: all studies (*low quality*) reported effectiveness of buprenorphine in treating chronic pain<sup>1</sup>

#### Patient on MOUD and Acute Pain<sup>2</sup>

- Patients who are physically dependent on opioids (e.g. methadone or buprenorphine) must be maintained on a daily equivalence before ANY analgesic effect is realized with opioids used for acute pain management
- Opioid analgesic requirements are often higher due to increased pain sensitivity and opioid cross tolerance

1. Cotes J, Montgomery L. Pain Med. 2014

2. Alford DP, et al. Ann Intern Med. 2016

135



SUMMARY: Part 2
Employ universal precautions but individualize care based on risk
Continue or modify opioid treatment based on clinical indication and response
Optimize office systems to involve the entire healthcare team
Document benefits, risks and harms and rationale for the plan of care





