

# Opioid Abuse and Pain Management

A review of changing government requirements  
and current medical guidelines for acute and  
chronic pain  
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## Opioids

- Currently a hot topic issue
- Large push from politicians to decrease mortality from opioid use
  - But didn't they start this?
  - Don't they grade us based on patient satisfaction
  - Where do we draw the line?

## Goals

- Understand the risk of opioid prescription
- Become familiar with opioid equivalents
- Know the safety limits of appropriate opioid Rx
- Meet requirements for Proper prescribing in your home state.
- \*Updates in management of common pain complaints.
- \*Then move in to case based questions and recommendations.

## Pearls

- Increased patient satisfaction increased mortality 26% [5]
- Cost increased 9.1% on drugs[5]
- 8.8% on testing [5]
- Opioids are not recommended for headaches[4]
- Opioids should be considered a last line
- Removing pain does not fix the problem

- a recent study of patients aged 15–64 years receiving opioids for chronic noncancer pain and followed for up to 13 years revealed that one in 550 patients died from opioid-related overdose at a median of 2.6 years from their first opioid prescription, and one in 32 patients who escalated to opioid dosages >200 morphine milligram equivalents (MME) died from opioid-related overdose (25).

## Oklahoma specific

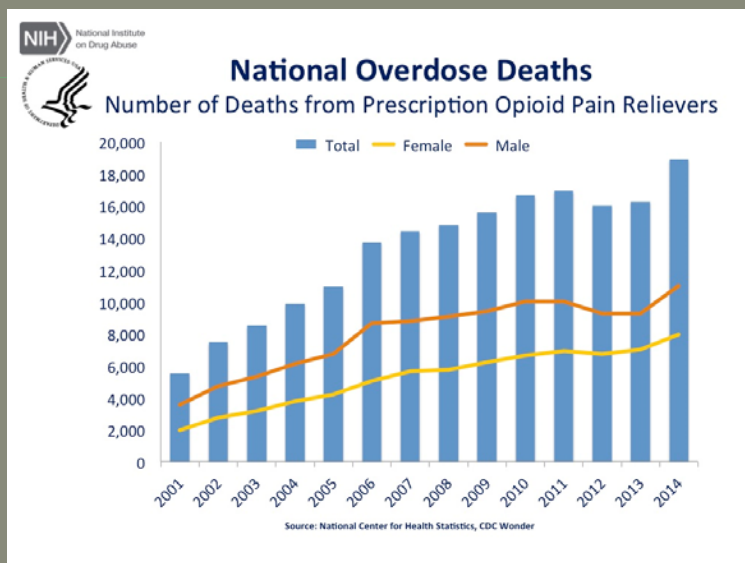
- Top 5 in prescribing
- Top 5 in deaths
- Major push for regulation and monitoring
- Required PMP checks
- Registering of pain management clinics
- Pill mills

## Governmental intervention

- CDC Guidelines
- Released March 2016
- Opioid overdoses and deaths
- Emphasis on high dose opioids
- First governmental guidelines
- Voluntary
- Reducing opioid consumption
- Access to treatment

## So How Did We Get Here?

- Pain as the 5<sup>th</sup> vital sign, started by the American Pain Society in 1995
- Big push from the VA in October 2000.
- From 1999-2014 rate of opioid overdose quadrupled
- CMS recently published a study showing that including pain management as part of quality indicators did not contribute to death,
  - However deaths HAVE trended up since the push for pain as the 5<sup>th</sup> vital

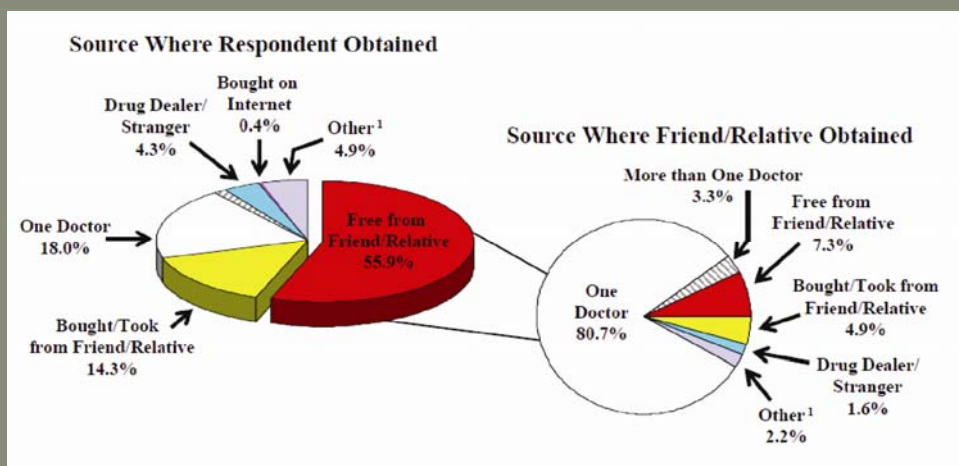


## National Center on Addiction and Substance Abuse

- 15.2 million abuse prescription drugs (2.5 X increase in 10 years)
- 20% of patients obtaining opioids for chronic pain abuse the medication
- 10-20% of these patients abuse illicit drugs
- Increased prescribing of opioids linked to misuse, abuse and deaths
- Absolute link between increased prescribing and availability for abuse

# Supply

- Explosion in the use of prescription opioids in response to the “under treatment” of pain
- Retail grams of opioids sold show significant increase
- Since 2004 risk has escalated without increased evidence of benefit
- Sources of opioids
- Number one source is from family and friends
- The medicine cabinet is our greatest threat



## Why is the Government Getting Involved

- Major reason for CDC involvement
- Significant escalation
- Diversion: most deaths are from “non-prescribed” opioids
- Lethal combinations especially with benzodiazepines
- Good data to support dose linked relationship
- Without question the number one reason for governmental intrusion

## Impact

- Physicians and nurses are being held accountable for patient death from overdose.
- Patients are suing physicians for becoming addicted to opioids.

2014

- White House recently unveiled a “multi-agency” plan to address the prescription drug epidemic
- Physician education
- Patient education
- Expanding monitoring systems
- Appropriate disposal of unused opioids
- Focus on “pill mills”
- Still only addresses Schedule II medications with emphasis on long acting opioids

## CDC Emphasis

- Directed at primary care physicians
- Opioids not recommended for routine use
- Does not include end of life, cancer pain and palliative pain care
- Management of pain is a multidisciplinary problem requiring numerous modalities to address physical and psychosocial aspects



## Goals

- Non-pharmacological approach
- Non-opioid approach
- Emphasis on Behavioral therapies
- Functional therapies
- Adjunctive medications
- Patient and provider expectation
- Opioids are a “last resort” option

## What Does that Mean to Me?

- The main goal is education.
- Tell patients why you are not writing pain medicine.
- Tell patients that pain medicine only leads to tolerance and side effects.

## Education

- Let patients know, that “good feeling” is euphoria
- Euphoria= getting high
- Goal is not to remove pain but improve function.
- Don't use pain medicine as a sleep medicine

## Do Pain Meds Ever Really Fix Anything?

- Evidence is scant
- Opioid use may be the most important factor impeding recovery of function
- Opioids do not consistently and reliably relieve pain and can decrease quality of life
- The routine use of opioids cannot be recommended
- As you can see the government is preparing a case to come down hard on us.

## Opioid Induced Hyperalgesia

- Increased sensitivity to noxious or non-noxious stimuli
- Sensitization of pro-nociceptive mechanisms
- Hypersensitivity and allodynia
- Confused with tolerance
- Caused with rapid escalation and high dose therapy
- Activity at the NMDA receptor in dorsal horn

## Last Ditch Effort

- Informed consent and discussion of risk vs. benefit
- Therapeutic trial of 4-6 weeks
- Exhaustion of other modalities
- Insufficient data on starting dose
- “ Start low go slow”
- Conversion tables
- Ongoing monitoring and assessment of benefit vs. risk, expectations and alternative modalities
- Consider a taper or wean even in functional patients

## Remember...

- Narcotics are not a life saving measure.
- You are not obligated to write.
- Patients are being put at risk
- Increased risk: Benzodiazepine use with opioids
- Significant increase in deaths and ER visits

- Providers should prescribe lowest possible dose
- Additional precautions at > 50 MED's
- Should avoid > 90 MED's

## Safety Lines

- Rate of Death 2X at >50MME
- Rate of Death 9X at >100MME

- Pay attention to a pattern of activity that suggests abuse and address
- Monitor closely through follow up and documentation
- Use available tools:
  - PMP website
  - UDS and pill counts
  - Pharmacies
- Obligated to protect yourself, your patient and society from opioid abuse and diversion

- WEAN!
- Contact law enforcement agencies
- Refer the patient for appropriate help
- Treat withdrawal if indicated
  - Do not treat withdrawal with more drugs, use clonidine and zofran
- Contact other physicians and pharmacies
- 30 day supply of opioids maximum
- Certain circumstances, consider referral
- “Under no circumstances may a physician dispense with the knowledge the drug will be abused or diverted” (DEA 2006)
  - *If a patient has abused/diverted drugs in past it is a felony to prescribe again if there is a chance at diversion*

## What the DEA considers inappropriate

- Inadequate attention
- Inadequate monitoring
- Inadequate patient education and consent
- Unjustified dose escalations
- Excessive opioid dosing
- Not using tools for risk mitigation

## What Have We Done?

- Removal of morphine from routine orders decreased the rate of falls by 50%
- Decreased the number of rapid responses by 33%
- Decreased the number of code blues associated with rapid responses another 15%

## What Are We Doing

- Removing Demerol
- Pharmacy to start calling when safety limits are reached.
- If a patient is not NPO or post surgery there is not a reason for them to be getting IV pain medicine (see attached chart)
- Movement towards opiod free ER
  - Hoping to remove Dilaudid formulary and order things in morphine equivalentents

## Results

- HCAHPS (Patient Satisfaction rates) increased 2.89% (p=0.025)
- Length of Hospital Stay reduction decreased 15.09% (p=0.0023)
- Falls decreased 17.31%
- Adverse Drug Events per 1,000 patient days down 20.69%
- No significant change in mortality
- Saved \$3.1 million

## Next question

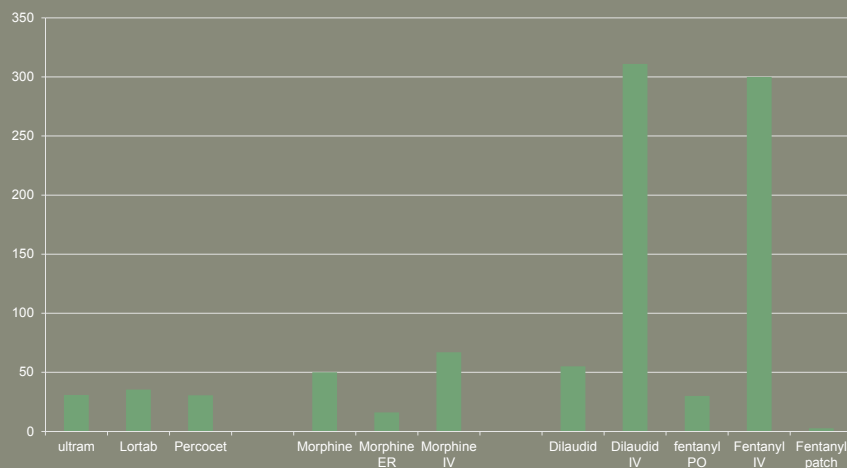
- Why the different doses and routes?



## Peak Evaluation

- This is how high and how fast it hits
- Think about it as the higher the number the greater the risk of side effects like respiratory arrest.
- Think Vanc trough, past a point the number is too high and you risk toxicity
- Or think alcohol
- 3 Beers vs a shot of Everclear
  - You end up with the same BAC at an hour but it is a function of how quick it hits you and the risk of side effects

## IV vs PO



## Studies have shown these to be equivalent doses

Drug	PO	IV
Ultram	75mg	n/a
Lortab	7.5mg	n/a
Percocet	5mg	n/a
Morphine	7.5mg	1.875mg
Dilaudid	.75mg	.1875mg
Fentanyl	75mcg	18.75 mcg

- People saying that an opiod equivalent dose doesn't work as well for them are falling victim to secondary events.
- "But IV makes me feel so much more relaxed"

## What Does a Safe Dose Look Like?

- 50MME/day Oral Meds=2x (100mme =9X)
- Ultram 100 mg q4-6      t  $\frac{1}{2}$  7 hours
- Lortab 10mg q4-6      t  $\frac{1}{2}$  4 hours
- Percocet 5mg q4      t  $\frac{1}{2}$  4 hours
- Percocet 10mg q 8      t  $\frac{1}{2}$  4 hours
  - Not recc dose bc t  $\frac{1}{2}$  < dosing frequency
- Morphine 10mg q 4-6      t  $\frac{1}{2}$  5 hours
- Dilaudid 2mg q 12      t  $\frac{1}{2}$  2.5 hours
  - So this is a never drug bc safe timing is much less than t  $\frac{1}{2}$
- Fentanyl 25mcg q 12      t  $\frac{1}{2}$  4 hours
  - Another never dose

## What Does a Safe Dose Look Like? 50 MME/day (and double for 9X risk)

- Morphine 2.5mg q 4 hours
- Morphine 5 mg q 8 hours
- Dilaudid 0.25mg q 4hours
- Dilaudid .5 mg q 8 hours
- Fentanyl 50mcg x1
  
- This is why it is imperative to switch to Oral as soon as possible
- There is no unlearning this information

## Change is coming

- We are going to review some common pain complaints from the:
  - Clinic
  - ER
  - Hospital
  - Palliative
- And current guidelines with secondary therapies.

## Regardless of the Place

- Start with a thorough history and physical.
- Make your documentation “Bullet proof”
  - PQRSTA on all, this is now a legal requirement at initiation and changes
  - Neuropathic: burning/numb/heavy/tingling
  - Nociceptive: Sharp/aching/throbbing
- Do what is best for the patient.
- Pain scale is the hardest.
  - Document appearance and rating, discuss discrepancies.
- Focus on education, no can be pain free all the time.
  - You should be striving for improved function, not complete pain relief.
  - Is there an underlying psychiatric component?

## Risk Mitigation

- Medical Necessity
  - Even if just increasing dose
- Risk factors for abuse
- Co-morbidity
- Failure of conservative therapies
- Informed consent
  - Big new one, does patient understand risk
- Primary Endpoint
  - Quality of life activities
- Indications for initial screening?

## Changes and Follow up

- CDC suggest follow up at least every 6 months [9]
- Many states are requiring this for licensure.

## New Patients, Weaning, Withdrawl

- A previous prescription for narcotics is not an indication on its own.
- Weaning is not indicated if
  - Opioids not present in UDS
  - Concerns for diversion (lost/stolen/shopping)
  - Risk factors greater than side effects of withdrawal
- Clonidine/Guanfacine and Promethazine at usual dosing to treat withdrawl symptoms
  - Replacement therapy is not indicated [9]

## Suggested Weaning schedule

- 2 to 3 week tapering regimen should be adequate in most cases
- daily dose by 10%
- 20% every 3-5 days
- 25% per week
- Avoid reducing the daily dose by > 50% at any given interval

## Terminating the physician-patient relationship :

- 1. Giving the patient or patient's representative written notice, which may be by certified mail, return receipt requested, or other reasonable proof. A copy of the letter should be included in the medical record.
- 2. Providing the patient with a brief and valid reason for terminating the relationship.
- 3. Agreeing to continue to provide care for a reasonable period of time (at least 30 days) in order to allow the patient to obtain care from another physician.
- 4. Providing recommendations to help the patient locate another physician of like specialty.
- 5. Offer to transfer records to the new physician upon signed authorization and include an authorization form with the letter.
- 6. A physician assistant or nurse practitioner may not independently terminate the physician-patient relationship.

## Screening tools

- Brief Risk Interview (BRI)
- Diagnosis, Intractability, Risk, Efficacy (DIRE)
- Opioid Risk Tool (ORT)
- Current Opioid Misuse Measure (COMM)
- Pain Medication Questionnaire (PMQ)
- Screener and Opioid Assessment for Patients with Pain-revised (SOAPP)
  - Preferred screener, less risk for deception

**A. Numeric Rating Scale**

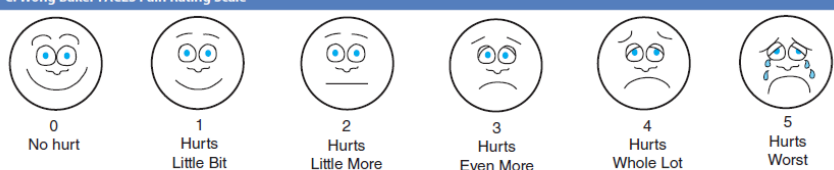
No pain Worst pain

0 1 2 3 4 5 6 7 8 9 10

**B. Numeric Rating Scale Translated Into Word and Behavior Scales**

Pain Intensity	Word Scale	Nonverbal Behaviors
0	No pain	Relaxed, calm expression
1-2	Least pain	Stressed, tense expression
3-4	Mild pain	Guarded movement, grimacing
5-6	Moderate pain	Moaning, restless
7-8	Severe pain	Crying out
9-10	Excruciating pain	Increased intensity of above

**C. Wong Baker FACES Pain Rating Scale<sup>1</sup>**



0 No hurt      1 Hurts Little Bit      2 Hurts Little More      3 Hurts Even More      4 Hurts Whole Lot      5 Hurts Worst

<sup>1</sup>Especially useful for patients who cannot read English and for pediatric patients.  
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## Nociceptive-less than 6 weeks

- Applies to most of these but is still a step wise approach [6]
  - Non-pharmacologic interventions
    - RICE, Manipulation, Rehab, offload, brace/splint
  - Nsaids (acetaminophen, ibuprofen, ketorlac)
    - Don't forget topicals such as voltaren gel.
    - Consider adjunct such as muscle relaxants.
  - Opioids
    - Only after escalation and a true effort
    - Consider CBT, trigger point injections, blocks (surgery as we move in to chronic)



## Neuropathic Pain [7]

- Antidepressants
  - Nortriptyline, Desipramine
- Calcium-Channel  $\alpha_2$  delta ligands
  - Gabapentin
  - Pregabalin
- SSNRI
  - Duloxetine
  - Venlafaxine
- Lidocaine topical or Capsaicin
- Anticonvulsants
  - Phenytoin, Carbamazepine, Oxcarbazepine
- Then Opioids
  - But should be followed by referral for intervention

## Chronic pain-any source

- Set your goals: focus on function and life not pain relief
- Multidisciplinary: medication, counseling physical therapy, blocks, surgery
- Actually follows the same path as neuropathic pain

- **The beneficial effect of opioids for chronic non-cancer pain is modest at best, and no measures have been identified to predict a good response.** [7]
- We can all think of patients who do better but most will ultimately have complications.
- When was the last time some was cured with a long term regimen of narcotics?

## Addiction Therapy

- Not the focus of this lecture, but just and FYI
- Patients adhere to Suboxone just as well as methadone.
- It has a higher rate of relapse once discontinued.
- Patients who stay on it longer tend to have less relapse.
- Both have the same rate of diversion/illicit use in the first six months (48.1%) [8]

## Breakthrough Pain

- Occasional break through pain is to be expected
  - If a patient is using breakthrough medicine more than 50% of the time than you need to increase their long acting

## Caveat

- These suggestions are based on appropriate History, Physical, and diagnosis.
- If the regimen isn't working re-evaluate.
- Key point that recurs in pain management is opioids are the last line treatment.

## Previous Paradigm [11]

- Nonopioid analgesic agents (eg, aspirin, acetaminophen, nonsteroidal anti-inflammatory drugs [NSAIDs], COX-2 Inhibitors)
- •Tramadol
- •Opioids
- •Alpha 2 adrenergic agonists
- •Antidepressants (tricyclics and serotonin-norepinephrine reuptake inhibitors [SNRIs])
- •Antiepileptic drugs (gabapentin, pregabalin, and other anticonvulsants)
- •Muscle relaxants
- •N-methyl-d-aspartate (NMDA) receptor antagonists
- •Topical analgesic agents

## New Paradigm<sup>[11]</sup>

- Nonopioid analgesic agents (eg, aspirin, acetaminophen, nonsteroidal anti-inflammatory drugs [NSAIDs], COX-2 Inhibitors)
- Topical analgesic agents
- Alpha 2 adrenergic agonists
- Antidepressants (tricyclics and serotonin-norepinephrine reuptake inhibitors [SNRIs])
- Antiepileptic drugs (gabapentin, pregabalin, and other anticonvulsants)
- Muscle relaxants
- Tramadol
- Opioids
- N-methyl-d-aspartate (NMDA) receptor antagonists

## Start low and go slow

- Should be one of the last options
  - Informed consent
  - Greater than 3-5 days increases chance of addiction
- Rates of addiction are 8-12% of those started on narcotic prescription past this point
- 1:100 will die, 1:100 will develop a heroin addiction, 40% will misuse in some manner[10]

## Palliative Care

- Focus on the patients goals [5], sedation is acceptable if that is consistent with end of life goals.
- **tolerance** (requiring increasing dosage to achieve the same analgesic effect)
- **dependence** (requiring continued dosing to prevent symptoms of medication withdrawal)
- **pseudo-addiction**, defined as exhibiting behaviors associated with addiction but only because their pain is inadequately treated.

## Specific palliative pain

- Metastatic Bone:
  - Bisphosphonates, RANKL inhibitors, and nerve blocks.
- Spinal compression/Brain Mets:
  - Dexamethasone, prednisone

## Route Pearls

- Pills, oral solutions, compounded (suckers), buccal patches
- Transdermal patches
  - Realize that the fentanyl patch is mcg per hour
  - Takes 20 hours to reach a steady state

## Clinic

- Headache (tension type/migraine)
  - Nsaids->NSAIDS w/ caffeine -> Indomethacin/prochlorperazine/caffeine-> Triptans ->adjuncts (injections, biofeedback)
  - Not opioids [12], high rate of rebound
  - Prevention:Beta-blockers/CCB, TCA (amitryptiline), SSNRI (venalfaxine), anticonvulsants ( topamax, valproate, gabapentin)

- Osteoarthritis
  - Physical therapy for strengthening
  - Bracing/Assistive devices
  - Acetaminophen is first line (up to 3 g per day)[13]
  - Intra-articular injection (hyaluronic acid q 6 months or steroids )
  - Surgery
  - Opioids should be reserve for those who are not a surgical candidate [13]

### • Low back pain

- Start with non-pharmacologic (manipulation, heat, alt)
  - Aren't you glad you're a DO?
- Initiate an anti-inflammatory or muscle relaxant
- Once chronic: Exercise , multidisciplinary approach, stretching, biofeedback , CBT
- Second line: Duloxetine, tramadol
- Opioids: Only if the benefits outweigh the risk and even then it is a weak recommendation [14]

### • Postherpetic neuralgia

- TCA (caution in elderly) -> gabapentin/lyrica -> opioids
- Can use topical Lidocaine/Capsaicin as an adjunct



- Trigeminal neuralgia

- Carbamazepine/oxcarbazepine/baclofen ->add lamotrigine and titrate to 400mg (topamax,tizanidine, or previous combinations)
- Botox
- Surgical decompression
- Rhizotomy/neurectomy/blocks

- Neuropathy

- Amitryptiline/desipramine/nortriptyline (less cholinergic effects) or or gabapentin/pregablin
- Second line: Duloxetine/venlafaxine (SSNRI)
- Third Line: Anticonvulsants,mexiletine, other antidepressants
- Adjunct: Capsaicin
- Phantom Limb: as above , not testable but mirror-box therapy, nmda, and calcitonin receptor agonist work great

- Ischemic[13]

- Underlying cause, cilostazol (pletal)->revascularization

## Clinic

- Sports/exercise injuries

- RICE, Nsaids/muscle relaxers, follows same algorithm as low back pain
- Pain persisting more than 6 weeks warrants a thorough work up for things such as stress fractures

- Gout

- Nsaids/Indomethacin->Colchicine->Steroids
- Allopurinol and probenecid for control
  - No allopurinol in acute phase

## Hospital/ER Setting

- Chronic pain is the same:
  - Emphasis on reason for admission not opportunity to fix old problems, don't start IV narcotics
- Chest Pain
  - Nitro/Nitro drip, opioids for chest pain is now a class IIb level of evidence and actually interferes with antiplatelet drugs
- Chest Pain-non cardiac
  - Nsaids
- a

## Hospital

- Pancreatitis:make sure you have the diagnosis
  - Acute:Hydration, NPO/clears/low fat diet, Morphine is preferred [15], \*new to me, avoid demerol for toxic buildup
  - Chronic: PPI, Pancreatic enzymes, Octreotide[10], offending agent abstinence, treat as neuropathic pain, tramadol
- Surgical
  - Surgery!
- Perioperative pain management should focus on lowest acceptable dose without excessive sedation and rapid transition to orals and de-escalation
  - Many areas are limiting post op to 3 days of medication
  - Not testable, just a thought

## Hospital/ER

- Sickle cell Crisis
  - Hydration, Oxygenation, Opioids
  - \*Address underlying cause, don't miss scary things
- Headache
  - If refractory to the clinical approach
  - Parenteral Ketorolac ->Metoclopramide + diphenhydramine

## Hospital/ER

- Migraine
  - Clinic treatment
  - Ketorolac -> prochlorperazine/metoclopramide/Droperidol/Chlorpromazine(give all with diphenhydramine)
  - Dexamethasone to reduce recurrence
  - DHE if refractory (pre-treat with antiemetics)

# Insert Practice questions here

1. Edwards JE, McQuay HJ, Moore RA: Single dose dihydrocodeine for acute postoperative pain. *Cochrane Database Syst Rev* CD002760. 2000. [PubMed: 11034754]
4. McHale FM, Lovecchio F: Narcotic analgesia in the acute abdomen—a review of prospective trials. *Eur J Emerg Med* 8: 131, 2001. [PubMed: 11436909]
5. Rabow MW, Pantilat SZ. Palliative Care & Pain Management. In: Papadakis MA, McPhee SJ, Rabow MW, eds. *Current Medical Diagnosis & Treatment 2017* New York, NY: McGraw-Hill; . <http://accessmedicine.mhmedical.com/content.aspx?bookid=1843&sectionid=135698406>. Accessed July 17, 2017.
6. Rabow MW, Pantilat SZ. Palliative Care & Pain Management. In: Papadakis MA, McPhee SJ, Rabow MW, eds. *Current Medical Diagnosis & Treatment 2017* New York, NY: McGraw-Hill; . <http://accessmedicine.mhmedical.com/content.aspx?bookid=1843&sectionid=135698406>. Accessed July 18, 2017.
7. Pain and Its Management. In: Kasper DL, Fauci AS, Hauser SL, Longo DL, Jameson J, Loscalzo J, eds. *Harrison's Manual of Medicine, 19e* New York, NY: McGraw-Hill; . <http://accessmedicine.mhmedical.com/content.aspx?bookid=1820&sectionid=127653738>. Accessed July 18, 2017.
8. <https://www.ncbi.nlm.nih.gov/books/NBK195158/>
9. <http://www.ncbi.nlm.nih.gov/pmc/articles/PMC2724222/>
10. Conwell DL, Banks PA. Acute Pancreatitis. In: Greenberger NJ, Blumberg RS, Burakoff R, eds. *CURRENT Diagnosis & Treatment: Gastroenterology, Hepatology, & Endoscopy, 3e* New York, NY: McGraw-Hill; . <http://accessmedicine.mhmedical.com/content.aspx?bookid=1621&sectionid=105184675>. Accessed July 18, 2017.
11. Philip Chang, MD, University of Kentucky research ongoing, as presented 7/1/2017 for joint webinar with US Dept of Health
12. PubMed TI Tension-type headache. AU Kaniecki RG SO Continuum (Minneapolis Minn) . 2012 Aug;18(4):823-34.
13. Johnson BE, Chapter 23. Arthritis: Osteoarthritis, Gout, & Rheumatoid Arthritis. In: South-Paul JE, Matheny SC, Lewis EL, eds. *CURRENT Diagnosis & Treatment in Family Medicine, 3e* New York, NY: McGraw-Hill; 2011. <http://accessmedicine.mhmedical.com/content.aspx?bookid=377&sectionid=40349415>. Accessed July 18, 2017.
14. <http://annals.org/aim/article/2003223/noninvasive-treatments-acute-subacute-chronic-low-back-pain-clinical-practice>
15. PubMedTI Peripheral arterial disease detection, awareness, and treatment in primary care. AU Hirsch AT, Criqui MH, Treat-Jacobson D, Regensteiner JG, Creager MA, Olin JW, Krook SH, Hunninghake DB, Comerota AJ, Walsh ME, McDermott MM, Hiatt WR SO JAMA. 2001;286(11):1317.
- 75 Thomas SH, Silen W: Effect on diagnostic efficiency of analgesia for undifferentiated abdominal pain. *Br J Surg* 90: 5, 2003. [PubMed: 12520567]
76. Tomas SH, Silen W, Cheema F et al.: Effects of morphine analgesia on diagnostic accuracy in emergency department patients with abdominal pain: a prospective, randomized trial. *J Am Coll Surg* 196: 18, 2003. [PubMed: 12517545]
77. Gallagher EJ, Esses D, Lee C, Lahn M, Bijur PE: Randomized clinical trial of morphine in acute abdominal pain. *Ann Emerg Med* 48: 150, 2006. [PubMed: 16953529]
78. Silen W: *Cope's Early Diagnosis of the Acute Abdomen*, 20th ed. New York: Oxford University Press, 2000: 5.
79. Johnston CC, Gagnon AJ, Pepler CJ, Bourgault P: Pain in the emergency department with one-week follow-up of pain resolution. *Pain Res Manag* 10: 67, 2005.

<http://www.cdc.gov/mmwr/volumes/65/rr/rr6501e1.htm>