




**Treatment of Pain
Opioid Choices and Considerations**


Greg Caldwell, OD, FAAO
Mid-Winter Getaway
Optometric Education Consultants
Sunday, January 28, 2024



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Disclosures- Greg Caldwell, OD, FAAO
All relevant relationships have been mitigated

- Lectured for: Alcon, B&L, BioTissue, Dompé
 - Disclosure: Receive speaker honorarium
- Advisory Board: Dompé, Immunogen, Iveric
 - Disclosure: Receive participant honorarium
- I have no direct financial or proprietary interest in any companies, products or services mentioned in this presentation
 - Disclosure: Non-salaried financial affiliation with Pharmacia
- Healthcare Registries – Chairman of Advisory Council for Diabetes and AMD
- The content of this activity was prepared independently by me - Dr. Caldwell
- The content and format of this course is presented without commercial bias and does not claim superiority of any commercial product or service
- Optometric Education Consultants – Scottsdale, AZ, Pittsburgh, PA, Sarasota, FL, Barcelona, Spain, Orlando, FL, Mackinac Island, MI, Quebec City, Canada, and Nashville, TN: Owner



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My Practice

I am a clinician first then a scientist

- Some are scientists first then clinician
- I need to simplify for patient and patient care.
- Science is great. But not good if there isn't a clinical application.
- Some lectures are science based without clinical application.
- My lecture will be a hybrid. Showing clinical applications of the science



It is wonderful to have someone who's digging so many aspects of optometry (scientific, clinical, education, teacher & lecturer). It is refreshing and very informative. Sarah

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Course Description

- This course will describe how to appropriately choose a pain medication based upon individual patient and drug factors.
- Additionally, opioid medications will be evaluated in terms of risk versus benefit, with an emphasis on pain levels and the potential for addiction.
- Case anecdotes will include management of ocular pain, with specific emphasis on oral/systemic medications and how to protect both patient and practitioner.

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Learning Objectives

- When given a patient case, choose an appropriate pain treatment plan for the management of ocular pain, in terms of drug choices based on pain level, dosing issues, and a monitoring plan for efficacy and toxicity.
- Identify and describe some of the potential signs, symptoms, and behaviors associated with opioid or substance abuse, and describe ways to respond to this issue.
- List systems available to evaluate a patient for potential opioid/substance abuse.
- Describe the treatment issues and options associated with the treatment of ocular pain in a patient with a drug abuse history.

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Two major types of pain:

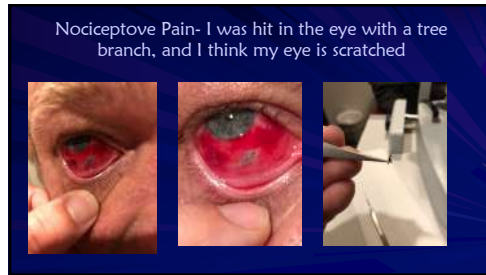
Nociceptive Pain – normal processing of stimuli that damages normal tissues; how pain becomes conscious:

- responsive to non-opioids
 - examples: NSAIDs, acetaminophen, steroids
- responsive to opioids
 - examples: codeine, hydrocodone, tramadol

Neuroathic: abnormal processing of sensory input by the peripheral or central nervous system:

- treatment includes adjuvant analgesics
 - sleep aids, nerve pain meds, muscle relaxers, anxiolytics

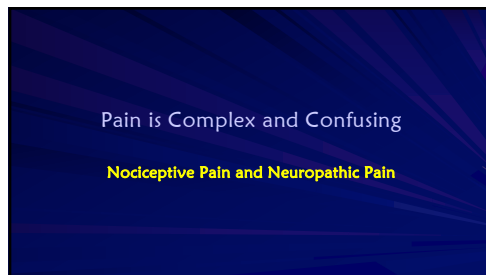
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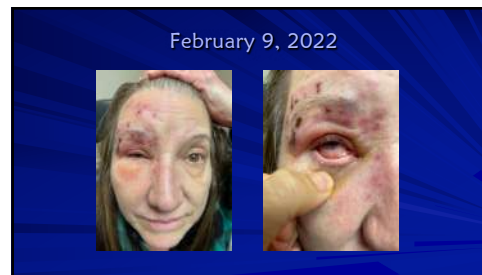
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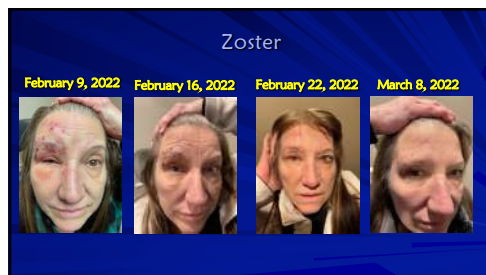
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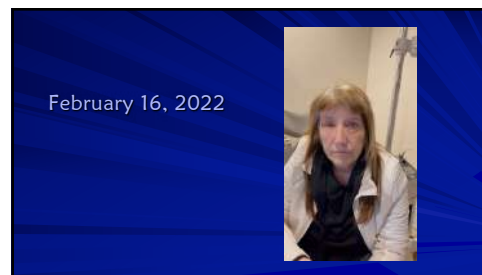
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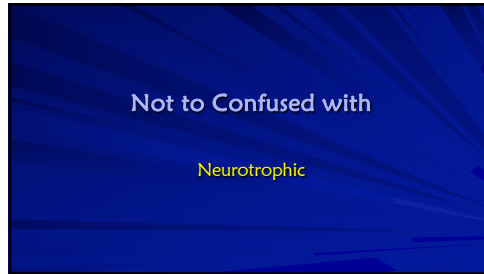
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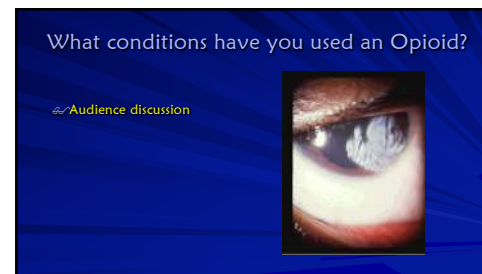
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What Opioid do you prescribe?

👉 Audience discussion



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Drug Treatment Options...Neuropathic Pain

👉 Why is this relevant?

👉 Adjuvants – means “add on” medications

- *Some of them have addiction potential
 - Anti-seizure medications that address nerve damage/inflammation
 - MOA: work on the GABA system – similar to benzodiazepines (ex. Xanax)
 - Gabapentin (Neurontin) – controlled substance in multiple states
 - Pregabalin (Lyrica) – controlled substance in all 50 states
 - Anti-anxiety and sleep medications
 - Zolpidem (Ambien)
 - Alprazolam (Xanax), Lorazepam (Ativan), Diazepam (Valium)

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Pain Assessments and Scales

👉 Adds objective data to a patient's feeling of pain

- * It is a subjective problem to assess!
- * Remember...no patient should needlessly suffer!

👉 “Does the injury or wound or diagnosis fit the patient's presentation?”

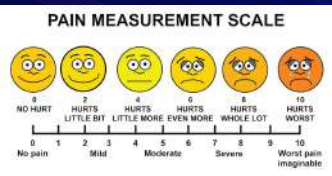
- * It is important to be able to assess the degree of pain in a patient.

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Combination Pain Scale...



PAIN MEASUREMENT SCALE

0 NO HURT 2 HURTS LITTLE BIT 4 HURTS LITTLE MORE 6 HURTS EVEN MORE 8 HURTS WHOLE LOT 10 HURTS WORST

0 1 2 3 4 5 6 7 8 9 10

No pain Mild Moderate Severe Worst pain imaginable

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Drug Treatment Options... Nociceptive Pain

3 Groups of analgesics

- * Non-opioids
 - Acetaminophen (Tylenol)
 - NSAIDs (Ibuprofen, naproxen sodium)
 - Glucocorticosteroids (methylprednisolone, prednisone)
- * Opioids –
 - Codeine (Tylenol with codeine)
 - Hydrocodone (Vicodin)
 - Tramadol (Ultram)

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Controlled Substance Schedules

Schedule I – not considered to be medically necessary, research only

- Heroin
- “Medical” Marijuana
 - State control of marijuana and CBD
- LSD
- Mushrooms
- Ecstasy

Schedule II – more likely to be abused (as compared to Schedule III, IV, V)

- Opioids, ASX “Narcotics”**
 - Oxycodone (OxyContin)**
 - Hydrocodone (Vicodin, Lorcet, Norco)
 - Morphine (MScotin, MSIR)
 - Hydromorphone (Dilaudid)
 - Methadone
 - Fentanyl (Duragesic)
- ADD/ADHD meds:
 - Methylphenidate (Ritalin)
 - Mixed amphetamine salts (Adderall)

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Controlled Substance Schedules

Schedule III – Safer, less likely to be abused (as compared to Schedule II)

- Combination products with APAP or ASA (codeine)
- Esketamine – nasal spray for treatment resistant depression

Schedule IV – Safer, less likely to be abused (as compared to Schedule II and III)

- Tramadol (Ultram)
- Benzodiazepines (lorazepam, diazepam, oxazepam)
- Sleep agents (zolpidem, etc.)

Schedule V – safest, least likely to be abused

- Expectorants with codeine

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Opioids “narcotics”

- ~ Mainstay of therapy for the treatment of pain
- ~ NO maximum daily dose limitation
- ~ Useful for acute and chronic pain

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Morphine Products

Morphine

- * Standard for comparison of other agents
- ~ **MSIR** (IR caps) (q 3-4 hours pm)
- ~ **MS Contin** (CR tabs) (q 8-12 hours)
- ~ **Kadian** (CR caps) (q 12 – 24 hours)
- ~ **Avinza** (CR caps) (q 24 hours)

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Hydromorphone Products

- Hydromorphone (Dilaudid)** tablets – immediate release
- Hydromorphone ER (Exalgo)** tablets – extended release
- ~ Used for severe pain

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Codeine-Based

- ~ Codeine – C3; Schedule III
- ~ Hydrocodone – C2; Schedule II
- ~ Oxycodone – C2; Schedule II

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Codeine tablets

- WEAK analgesic: commonly used, so MOST have heard of it!
- Add acetaminophen/aspirin – Schedule III
 - ***Tylenol #3** = 300 mg acetaminophen & 30 mg codeine
- Add expectorant – Schedule V
 - * If you think someone won't try to get their hands on "codeine cough syrup" as a drug of abuse, you'd be surprised!!!

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Oxycodone Products

- Long-Acting, Extended-Release
 - OxyContin
- Immediate Release: short-acting tablets
 - OxyIR** (IR cap)
 - Roxicodone** solution
- with Acetaminophen:
 - Percocet** and **Endocet** (oxycodone/APAP dose)

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OxyCONTin (Controlled release tablets (q 12 hours...once in a while q 8 hours); new formulation is out to help control abuse)

Manual Crushing Followed by Dissolution



The image shows two spoons. The left spoon, labeled 'Crushed New Formulation', contains several small, irregular, yellowish-white fragments. The right spoon, labeled 'Crushed Original Formulation', contains a larger, more cohesive, white, powdery mass.

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Tampering for IV Abuse

- New formulation results in gelatinous material which cannot be drawn into a syringe for injection (the syringe is empty)



The image shows two syringes. The left syringe, labeled 'New formulation', is empty. The right syringe, labeled 'Original formulation', contains a white, gelatinous substance that has been drawn into the syringe.

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Hydrocodone Products

- Immediate-Release Products
- AS OF AUGUST 2014, hydrocodone products are ALL CIII!
- Hydrocodone 7.5 mg + IBU 200 mg (Vicoprofen)**
- Hydrocodone + acetaminophen:**
 - ***Vicodin**® 5/300; 7.5/300; 10/300
 - ***Lortab** = 2.5/300, 5/300, 7.5/300, 10/300
 - ***Norco** = 5/325, 7.5/325, 10/325

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Miscellaneous

- Fentanyl Patch (Duragesic)**
 - * MOST potent opioid
 - * Black Box Warning against use in acute pain and in opioid naïve patients
- Methadone**
 - * Typically reserved for morphine/codeine allergic patients

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Methadone tidbits...

- Chronic pain or opioid abuse deterrent
- 2-phase elimination
 - Alpha phase = 8 hrs
 - Opioid pain control
 - Beta phase = 16+ hrs
 - Mitigates withdrawal symptoms
- Patient 1: On a short-acting pain med = likely being used to treat chronic pain
 - Twice per day dosing
- Patient 2: On methadone ONLY; lower doses
 - Once daily dosing

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Tramadol

Tramadol (Ultram) tabs
Tramadol with 325 mg APAP (Ultracet), Tramadol ER tabs

- Dual action: **mu** receptors & inhibits neuronal uptake of **serotonin** & **norepinephrine**
- Lowers seizure threshold; increases serotonin levels
 - watch drug interactions with other meds that ↑ serotonin
 - Selective serotonin reuptake inhibitors (SSRI): fluoxetine/Prozac
 - Migraine meds ("triptans"): sumatriptan/Imitrex
- AS OF AUGUST 2014, NOW A C4 (Schedule IV)
 - "tramies" = abuse potential; helps decrease withdrawal symptoms

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Opioid Allergies

⚡ If a patient states "codeine allergic", ask appropriate questions...

- *"You have indicated that you have an allergy to codeine, can you describe what happens when you take codeine?"
 - This is SIGNIFICANT, because if a patient is truly allergic to codeine, then they are most likely allergic to morphine, hydromorphone, oxycodone, hydrocodone, and tramadol
 - AND...if they had an opioid IV after surgery, then their "reaction" may have been due to histamine release...
 - NOT always an allergic reaction

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Opioid Allergies

⚡ DO YOU KNOW WHAT A PATIENT CAN TAKE?

- Fentanyl
- Methadone
- Meperidine

⚡ Assessing "allergies" appropriately helps practitioners sort through ACTUAL allergy potential and "placebo allergies"

- Fear versus drug seeking

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Specific Medications Using Numeric Pain Scale

Mild pain = 1 - 3

- Acetaminophen (APAP; Tylenol)
- Ibuprofen (Advil, Motrin)
- Naproxen sodium (Aleve)
- Tramadol (Ultram) - low dose

Moderate pain = 4 - 6

- Tramadol (Ultram) - mid to high dosing
- Tylenol with codeine (Tylenol #3)
- Acetaminophen with oxycodone (Percocet)
- Acetaminophen with hydrocodone (Vicodin, etc.)

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Specific Medications Using Numeric Pain Scale

Severe pain = 7 - 10

- Tylenol with hydrocodone (Vicodin, etc.) - higher doses
- Tylenol with oxycodone (Percocet, etc.) - higher doses
- Morphine (MSIR)
- Hydromorphone (Dilaudid)
- Fentanyl (Duragesic patch; Actiq lozenge on a stick)

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Opioid Effects/ADRs

- ~ Sedation
- ~ Euphoria – mu receptors
- ~ Dysphoria/Hallucinations
- ~ Pruritis – allergy versus normal release of histamine
- ~ Nausea/vomiting
 - * Triggers CTZ
 - * Codeine "allergy"

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Opioid Effects/ADRs

- ~ Confusion
- ~ Miosis
- ~ Respiratory depression – this is what kills a patient
 - * **Mixing opioids with other CNS depressants**
 - Alcohol
 - Benzodiazepines
 - Muscle relaxers
 - Sleep agents
 - Antihistamines
 - Anti-seizure medications

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Opioid Effects/ADRs


- ~ Withdrawal symptoms:
 - * Short half-life agents are more likely to cause abrupt withdrawal symptoms
 - * Sweating
 - * High sympathetic tone: increase in heart rate and blood pressure, mydriasis
 - * Agitation
 - * Irritation
 - * Irrational behavior
 - * Symptoms disappear with (immediate) use of an opioid

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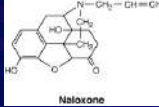
Opioid Antagonists

Naloxone (Narcan) & Naltrexone (ReVia)

- * Used to treat opioid overdose



Morphine



Naloxone

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Mixed Opioid Agonist-Antagonist

- ~ Exhibit partial agonist or antagonist activity at the opioid receptors
- ~ Agonist/Antagonist combinations for the TREATMENT of chronic pain
 - * NOT appropriate for the treatment of acute pain
 - * Morphine/Naltrexone (Embeda)
 - * Oxycodone/Naltrexone (Troxyc ER)
- ~ Schedule II controlled substance

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Mixed Opioid Agonist-Antagonist

- ~ Exhibit partial agonist or antagonist activity at the opioid receptors
- ~ Agonist/Antagonist combinations for the TREATMENT of opioid abuse/addiction
 - ~ Buprenorphine (Buprenex)
 - ~ Buprenorphine/Naloxone (Suboxone)
- ~ Schedule III
 - ~ Adverse effects
 - * Less respiratory depression & less abuse potential?
 - ~ Precipitate withdrawal in an opioid-dependent patient

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Painful Ocular Problems – things to consider...

- ~ Acute or chronic?
 - * YOU are in charge!
 - * Legal and ethical issues – do not allow yourself to be bullied by the patient!
- ~ Work with other practitioners!
- ~ Only a pain specialist should write RXs for CII medications for chronic pain issues
 - * If something looks suspicious, then make inquiries! Especially before you write an RX for a drug that can be abused and/or sold.

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Painful Ocular Problems – things to consider...

- ~ Use the tools that are available!
 - * State databases
 - PDMP = Prescription Drug Monitoring Program
 - * Pharmacists

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Tolerance

- ~ Escalation of dose to maintain effect (analgesia or euphoria)
 - * Happens to everyone
- ~ Regarding euphoria = may be life threatening because respiratory depression does not show much tolerance

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"True Addiction" (formerly "psychological dependence")

- ~ Compulsive use despite harm
- ~ Many times triggered by cravings in response to specific cues
 - * Lifestyle is geared to the acquisition of the drugs
 - * Borrowing from others, injecting oral formulations, prescription "loss", requesting specific drugs (not always a sign...at some drugs just work better)
- ~ Quality of life is not improved by the medication and eventually it becomes compulsive ("wanting without liking")
- ~ Relapse is very common even after "successful" withdrawal...it is a relapsing disease that is incredibly hard to treat

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Identifying Behaviors of Abuse/Addiction

- ~ New patients that don't seem to "fit"
- ~ "fast talkers"
- ~ Strange allergies
- ~ Excuses for "loss" of meds or why they need "a strong pain medication"

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Ways to respond

- ~ Avoid getting "bullied"
- ~ Avoid acting like you are judging the patient
- ~ State data bases
 - * Call your local pharmacy/pharmacist
- ~ Legal/ethical issues
 - * If you didn't write it down, then it didn't happen!
 - * If you accidentally give an addict a script for a pain medication, you won't get into "trouble"...

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Substance abuse history...

- ~ Avoid all opioids in a patient with a history of heroin use
 - * This includes tramadol
 - * May trigger dopamine reward and the drug "need"
 - * Stick with higher doses of a NSAID +/- acetaminophen
- ~ Patients with abuse history for other substances (ex. Benzodiazepines, alcohol, amphetamines)?
 - * It is a judgement call
 - * Some evidence to suggest that all addictive meds should be avoided!


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Conditions Which May Require Pain Management

- ~ Large cornea abrasions
 - * Cornea burn
 - * PRK/PTK
- ~ Orbital trauma
- ~ Orbital blowout fractures
- ~ Scleritis




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