Pain management, Identification of Addiction, Practices of Prescribing, and Dispensing of Opioids

Tracy Offerdahl, PharmD
Greg Caldwell, OD, FAAO
2 hour course
Biography – Tracy Offerdahl

- Dr. Offerdahl attended Temple University School of Pharmacy in Philadelphia, PA, for both her undergraduate and graduate degrees. Upon completion of her Doctor of Pharmacy degree, she completed a Residency at Temple University Hospital, where much of her time was spent in internal medicine and infectious diseases. She is currently on faculty at Salus University, Pennsylvania of Optometry, and the Philadelphia College of Osteopathic Medicine, and she has a practice site in Villanova, PA. Dr. Offerdahl is currently earning another degree in holistic medicine, a certificate in veterinary pharmacy, and she lectures extensively to the optometry community regarding systemic drugs.
Biography – Greg Caldwell

- Greg Caldwell, O.D., is a 1995 graduate of the Pennsylvania College of Optometry. He completed a one-year residency in primary care and ocular disease at The Eye Institute in Philadelphia Pennsylvania. He is a fellow of the American Academy of Optometry (AAO) and a Diplomate of the American Board of Optometry (ABO). He currently works in Duncansville, Pennsylvania as an ocular disease consultant. Dr. Caldwell’s primary focus is the diagnosis and management of anterior and posterior segment ocular disease and he has been a participant in multiple FDA investigations. Dr. Caldwell has lectured extensively throughout the county and over twelve countries internationally. In 2010 he served as President of the Pennsylvania Optometric Association (POA) and served on the AOA Board of Trustees 2013-2016. He is President of the Blair/Clearfield Association for the Blind.
Disclosures- Greg Caldwell, OD, FAAO

• Will mention many products, instruments and companies during our discussion
  • I don’t have any financial interest in any of these products, instruments or companies
• Pennsylvania Optometric Association –President 2010
  • POA Board of Directors 2006-2011
• American Optometric Association, Trustee 2013-2016
  • Thank you to the members and those who join
• I never used or will use my volunteer positions to further my lecturing career
• Lectured for: Shire, BioTissue, Optovue, Alcon, Allergan, Aerie
• Advisory Board: Allergan, Sun, Arerie
• Envolve: PA Medical Director, Credential Committee
• Optometric Education Consultants- Scottsdale, Nashville, and Quebec City, Owner
Disclosures- Tracy Offerdahl, PharmD

- Boiron: honorarium, webinar/speaker
- Has not received any assistance from any commercial interest in the development of this course
Course Objectives

- Describe the differences between nociceptive pain and neuropathic pain
- List and describe how to interpret pain scales
- Describe the commonly prescribed pain medication classes in terms of mechanisms, side effects, drug interactions, and applicability for pain management:
  - Opioids
    - Codeine-based
    - Morphine-based
    - Novel agents
  - Combinations therapy
- When given a patient case, choose an appropriate pain treatment plan for the management of ocular pain, in terms of drug(s), dosing issues, duration of treatment, 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
NIH: National Institute on Drug Abuse
As of March 2018

• Every day, more than 115 people in the United States die after overdosing on opioids
• The misuse of and addiction to opioids
  • Prescription pain relievers, heroin, and synthetic opioids such as fentanyl
• Serious national crisis that affects public health as well as social and economic welfare
• The Centers for Disease Control and Prevention estimates that the total "economic burden" of prescription opioid misuse alone in the United States is $78.5 billion a year
  • Including the costs of healthcare, lost productivity, addiction treatment, and criminal justice involvement
What do we know about the opioid crisis?
NIH: National Institute on Drug Abuse (March 2018)

• Roughly 21 to 29 percent of patients prescribed opioids for chronic pain misuse them
• Between 8 and 12 percent develop an opioid use disorder
• An estimated 4 to 6 percent who misuse prescription opioids transition to heroin.
• About 80 percent of people who use heroin first misused prescription opioids
• Opioid overdoses increased 30 percent from July 2016 through September 2017 in 52 areas in 45 states
• The Midwestern region saw opioid overdoses increase 70 percent from July 2016 through September 2017
• Opioid overdoses in large cities increase by 54 percent in 16 states
What are HHS and NIH doing about it?

• In the summer of 2017, NIH met with pharmaceutical companies and academic research centers to discuss:
  • Safe, effective, non-addictive strategies to manage chronic pain
  • New, innovative medications and technologies to treat opioid use disorders
  • Improved overdose prevention and reversal interventions to save lives and support recovery
Two major types of pain:

- **Nociceptive Pain** – normal processing of stimuli that damages normal tissues; how pain becomes conscious;
  * Responsive to non-opioids and opioids

- **Neuropathic**: abnormal processing of sensory input by the peripheral or central nervous system;
  * Treatment includes adjuvant analgesics
  * Sometimes much harder to treat
Drugs Treatment Options... Neuropathic Pain

• Not the focus of today’s discussion...
• Adjuvants – multipurpose & specific to type of pain
  • Anti-seizure medications that address nerve damage/inflammation
    • Gabapentin (Neurontin)
    • Pregabalin (Lyrica)
    • Topiramate (Topamax)
  • Sleep, depression, anxiety, muscle aches/spasms
Goals of Pain DO Differ...

- The goal for managing **acute pain** is to keep the patient as comfortable as possible while minimizing the **adverse drug reactions (ADRs)** from the pain meds.

- The goals for managing **chronic pain** are to keep the patient as comfortable as possible (this may not mean the patient is pain free), and integrating the patient back into a “normal life” and activities of daily living, while minimizing the ADRs from the pain meds.
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!

- **“Eyeball” the patient!**
  - Does the injury or wound or diagnosis fit the patient’s presentation?
    - BP
    - HR
    - Additional autonomic nervous system tidbits
      - Sweating
      - Nausea/vomiting
      - Pupil size
      - ACUTE vs CHRONIC
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: No pain
1: Mild
2: Moderate
3: Severe
4: Worst pain imaginable
Combination Pain Scale...

- **Verbal Descriptor Scale**
  - **0** No pain
  - **1-3** Mild pain
  - **4-6** Moderate pain
  - **7-9** Severe pain
  - **10** Very severe pain
  - **Worst pain possible**

- **Wong-Baker Facial Grimace Scale**
  - **0** No pain
  - **1-3** You feel some pain or discomfort but you can still complete most activities.
  - **4-6** The pain makes it difficult to concentrate and may interfere with your ability to do certain normal activities, such as reading, watching TV, having a phone conversation, etc.
  - **7-9** The pain is quite intense and is causing you to avoid or limit physical activity. Cannot concentrate on anything except pain.
  - **10** Worst pain imaginable.
Drug Treatment Options...
Nociceptive Pain

- **3 Groups of analgesics**
  - Non-opioids – acetaminophen & NSAIDs
  - Opioids – $\mu$ agonists & mixed agonist-antagonists
  - Adjuvants – multipurpose & specific to type of pain
    - Sleep, depression, anxiety, muscle aches/spasms
Opioids ("narcotics")

- Mainstay of therapy for the treatment of pain
- NO maximum daily dose limitation
- Useful for acute and chronic pain
- They mimic the actions of endogenous opioid compounds:
  - enkephalins, dynorphins, endorphins
Controlled Substance Schedules

- **Schedule I** – not considered to be medically necessary, research only
  - Heroin; “Medical” Marijuana

- **Schedule II** - More likely to be abused
  - **“Narcotics”:** Morphine, fentanyl, meperidine, hydromorphone, oxycodone, methadone, hydrocodone
  - **ADD/ADHD meds:** Methylphenidate, dexmethamphetamine, amphetamine salts

- **Schedule III** - Safer, less likely to be abused
  - Combination products with APAP or ASA (codeine)

- **Schedule IV** – Safer, less likely to be abused
  - Tramadol (Ultram)
  - Benzodiazepines (lorazepam, diazepam, oxazepam)
  - Sleep agents (zolpidem, etc.)

- **Schedule V** – safest, least likely to be abused
  - Expectorants with codeine
State-By-State Restrictions...

- **Marijuana**
  - Still considered to be “C1” or “Schedule I”
  - Federal government “ignores” it

- **Hydrocodone products**
  - C3 to C2 as of 2014
  - “hydrocodone exception”
    - NJ, etc
Mechanisms of Action:

- Relieve pain and induce euphoria by binding to the opioid receptors (mu, kappa, delta) in the brain and spinal cord
  - Gamma is not an opioid receptor subtype

- **Mu, kappa, delta** receptors in other places = ADRs
  - Mu: analgesia, euphoria, miosis, sedation, constipation, respiratory depression, addiction
  - Kappa: analgesia, diuresis, sedation, miosis, dysphoria, psychomimetic effects, respiratory depression, constipation
  - Delta: analgesia
Formulations...

- **Immediate release:**
  - AKA short-acting; breakthrough pain
  - “breakthrough pain” = acute pain that occurs after a chronic pain patient has taken their long-acting product
    - Ex: Cancer patient takes OxyContin at 7am. Their next dose is due at 7pm, however they have an acute pain problem at 12 noon...they would take a dose of their short-acting (“breakthrough pain”) opioid to treat the acute pain.
  - Uses: acute pain; breakthrough pain
    - Ex: Percocet, Tylenol w/ codeine, tramadol, Vicodin, etc.

- **Controlled release:**
  - AKA long-acting; sustained release; extended release
  - Uses: basal control of chronic pain; typically NOT for acute pain nor in opioid naïve patients!
    - “basal control” = basic or low-level pain control that lasts for 24 hours
    - Ex: OxyContin, MS Contin, Duragesic patch, etc.
Morphine Products

- **Morphine**
  - Standard for comparison of other agents
- Used for severe pain
- Multiple BRAND/TRADE names for long-acting morphine products, with very diverse delivery and release systems
Morphine Products

- **MSIR** (IR caps) (q 3-4 hours prn)
- **MS Contin** (CR tabs) (q 8–12 hours)
- **Kadian** (CR caps) (q 12 – 24 hours)
- **Avinza** (CR caps) (q 24 hours)

**Clinical pearls**
- ANY long-acting products (MS Contin, OxyContin, etc.) should NEVER be chewed, crushed, or cut in half, as this results in the ENTIRE dose being released at once and this may result in death due to overdose!
  - Death is typically due to respiratory depression
  - Morphine sulfate (Avinza) would be the most dangerous medication in the class if crushed or chewed
Hydromorphone Products

- **Hydromorphone (Dilaudid) tablets**
  - Take 1 – 2 tablets every 4 to 6 hours as needed for pain

- **Hydromorphone ER (Exalgo) tablets**
  - Take once per day

- Used for severe pain

- Very potent
  - Equianalgesic
    - 30 mg PO morphine = 8 mg PO hydromorphone
Codeine-Based

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

- **The WEAKEST opioid analgesic (codeine)**
  - Equianalgesic
    - 30mg PO morphine = 200 mg PO codeine
- **Add acetaminophen/aspirin – Schedule III**
  - Tylenol #2 = 300 mg acetaminophen & 15 mg codeine
  - **Tylenol #3** = 300 mg acetaminophen & 30 mg codeine
  - Tylenol #4 = 300 mg acetaminophen & 60 mg codeine
- 1 – 2 tablets every 4 – 6 hours as needed for pain
  - Not to exceed **3 grams** of APAP per day
- **Add expectorant – Schedule V**
Oxycodone Products...
Oxycodone (OxyCONtin)

- Controlled release tablets
  - q 12 hours...once in a while q 8 hours
- Generally not appropriate for acute pain due to the fact that it is long acting
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

Crushed New Formulation
Crushed Original Formulation
Tampering for IV Abuse

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

New formulation  Original formulation
Oxycodone Products

- **Immediate Release; short-acting tablets**
  - **OxyIR (IR cap)** 5 mg
  - **Roxicodone** solution 5 mg/5 mL

- With APAP:
  - **Percocet** and Endocet (oxycodone/APAP dose)
    - 2.5/325, **5/325**, 7.5/325, 7.5/500, 10/325, 10/500, 10/650 tablets

- Take 1 – 2 tablets by mouth every 4 to 6 hours as needed for pain
  - Not to exceed 3 grams of APAP per day
Oxycodone Products

- **Roxicet** solution—oxycodone 5 mg + 325 mg APAP/ 5 mL
- **Percodan** (oxy + asa) – no one uses this product
- Beware of combination with acetaminophen (Percocet), various strengths
- **Equianalgesic**
  - 30 mg PO morphine = 20 mg PO oxycodone
Hydrocodone Products

- **Hysingla ER, Zohydro ER** – long-acting, single ingredient hydrocodone

- **Immediate-Release Products:**
  - Hydrocodone + APAP (Norco, Vicodin, Lortab)
  - Hydrocodone (7.5 mg)+ IBU 200 mg (Vicoprofen)

- "Vicodin" = 5/500, Vicodin ES = 7.5/750, Vicodin HP = 10/660
  - **GENERIC/Brand new doses** = 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
Hydrocodone Products

• “Take 1 – 2 tabs/caps every 4 – 6 hours as needed for pain
  • Not to exceed 3 grams of APAP per day

• CIII - for moderate/severe pain – works well
  • As of August 2014, hydrocodone products (with and without acetaminophen) are ALL CII

• Equianalgesic dosing:
  • 30 mg PO morphine = 20 mg PO hydrocodone
**Miscellaneous**

- **Fentanyl Patch** *(Duragesic)*
  - MOST potent opioid *(Fentanyl)*
  - Black Box Warning against use in acute pain and in opioid naïve patients

- **Meperidine** *(Demerol)*
  - ACTIVE metabolites = undesirable effects that include drug-seeking, hallucinations, anxiety, CNS toxicity

- **Methadone**
  - Typically reserved for morphine/codeine allergic patients
Methadone tidbits...

- Chronic pain or opioid abuse deterrent
- 2-phase elimination
  - Alpha phase = 8 hrs
    - Offers 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
Methadone tidbits...

• Don’t think only used for heroin addicts
  • Helps prevent the symptoms of withdrawal
  • May be used to treat chronic pain
  • May be used to treat opioid addiction
Acute Ocular Pain

• Based on the classes of drugs covered to this point, hydrocodone and acetaminophen would be the best choice to manage ocular pain
• Better than morphine, oxycodone, and fentanyl
Tramadol – an Optometrist’s best friend

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

- Tramadol 50 – 100 mg q 4 – 6 hours
  - Do not exceed 400 mg/day
  - *Dual action:* mu receptors & inhibits neuronal uptake of serotonin & norepinephrine
  - Lowers seizure threshold; increases serotonin levels
    - Clinical pearl: Watch drug interactions with other meds that ↑ serotonin
      - As this may cause dangerous levels of serotonin!
    - Examples of serotonergic agents:
      - Selective serotonin reuptake inhibitors (SSRIs): fluoxetine/Prozac
      - Migraine meds (“triptans”): sumatriptan/Imitrex
  - Not controlled (was once non-controlled)
    - As of AUGUST 2014, now a CIV (Schedule IV)
    - Addicts call tramadol “tramies” = abuse potential; addicts use it because it helps decrease withdrawal symptoms
Specific Medications Using Numeric Pain Scale

- **Mild pain = 1 – 3**
  - Acetaminophen, Ibuprofen, Tramadol

- **Moderate pain = 4 – 6**
  - Tramadol
  - Tylenol with codeine
  - Acetaminophen with oxycodone (Percocet)
  - Acetaminophen with hydrocodone (Vicodin, etc.)

- **Severe pain = 7 – 10**
  - Oxycodone (Oxycontin)
  - Tylenol with hydrocodone (Vicodin, etc.)
  - Tylenol with oxycodone (Percocet, etc.)
  - Morphine, Hydromorphone, Fentanyl Patch
Opioid Effects/ADRs:

- **CONSTIPATION** - anticipate it! (note it’s in capital letters)
  - **All patients** should be treated for this problem and should receive a stool softener + stimulant combo
    - docusate + senna/Senna+S
    - Clinical pearl: Side effect of opioids, treat immediately, it occurs in all patients taking opioids

- **Pruritus** – normal histamine release – PARTICULARLY problematic with **parenteral (IV)** formulations more than with oral formulations
  - Clinical pearl: IV delivery causes more pruritus than oral

- **Nausea/vomiting**
  - All opioids trigger CTZ (chemoreceptor trigger zone) = central “vomiting center”
  - Codeine has the highest affinity for the CTZ! Which is why many patients state that they are “allergic to codeine”
    - Codeine “allergy” – MANY with “codeine allergy” state that their “allergy” is actually nausea/vomiting...not TRUE allergy

- **Sedation** – patients will become tolerant to this after a few weeks of consistent opioid use
Opioid Effects/ADRs:

- Inhibition of cough reflex
- Confusion
- Euphoria – due to mu receptor agonist portion of the mechanism
- Dysphoria/Hallucinations – due to kappa receptor agonist portion of the mechanism
- Miosis
- Respiratory depression – this is what kills a patient
Opioid Allergies

- If a patient is allergic to morphine and/or codeine, then they may only be able to safely take:
  - Methadone
  - Meperidine
  - Fentanyl
  - Tramadol

- **ASK appropriate questions…**
  - “what happens when you take _______?”
Opioid Antagonists

Naloxone (Narcan) & Naltrexone (ReVia)

* Used to treat opioid overdose
Naloxone (Narcan) & Naltrexone (ReVia)

- Rapidly reverse effects of morphine & other opioid agonists
  - States are offering these to patients, friends, and family members for patients ON opioids for pain management OR for addicts!

- Causes “antagonist-precipitated withdrawal”
  - Within 3 minutes after injection the s/sx of withdrawal appear; peak in about 10-20 minutes and subside in about 1 hour

- Adverse effects: insomnia, headache, nervousness, low energy, agitation, diarrhea, vomiting
Mixed Opioid Agonist-Antagonist

- Exhibit partial agonist or antagonist activity at the opioid receptors

- **Morphine/Naltrexone (Embeda), Oxycodone/Naltrexone (Troxyca ER)** – TREATMENT of chronic pain; C2

- **Buprenorphine (Buprenex), Buprenorphine/Naloxone (Suboxone)** – TREATMENT of opioid abuse; C3

- **Adverse effects**
  - Less respiratory depression & less abuse potential?

- Precipitate withdrawal in an opioid-dependent patient
Alternatives for Pain Control

- **Acetaminophen (APAP)**
  - Mild to low-Moderate pain
  - ADRs: liver, kidney?

- **Traditional NSAIDs and COX-2 Inhibitors**
  - Ibuprofen, naproxen sodium (Aleve), celecoxib (Celebrex), meloxicam (Mobic)
  - Mild to low,mid-moderate pain
  - ADRs: GI, CV, acute kidney failure, BP

- **Corticosteroids**
  - Inflammatory pain
  - ADRs: cataracts, BP, fluid retention, GI
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
Painful Ocular Problems – Things to consider...

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

• The end result of the undertreatment of pain
• Patients will know exactly how many pills they have left
• They may try to get appointments earlier than needed
  • They may seem anxious...
• Typically “cured” by changing/improving pain meds
“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...as 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
Addiction

- Remember, this is compulsive use despite harm!
  - Fast talkers
  - New patients
  - Unequal diagnosis and pain response
  - Vitals
  - Specific requests to agents
  - Strange “allergies”
  - Excuses
    - “I got robbed”; “I lost it”; “The pharmacy didn’t give me enough”…

Ways to respond

• Avoid getting “bullied”
• Avoid acting like you are judging the patient

• State databases
  • 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”…
Withdrawal from opioids...

- Time of onset, intensity, and duration of abstinence syndrome depends on the drug previously used (related to the half-life/“t ½”)

- Rhinorrhea, lacrimation, **yawning**, chills, gooseflesh, hyperventilation, hyperthermia, **mydriasis**, **muscular aches**, vomiting, diarrhea, anxiety, hostility

  - Number and intensity of signs and symptoms are largely dependent on the degree of physical dependence that has developed

- Administration of an opioid at the time of s/sx of withdrawal = suppression of abstinence signs and symptoms almost immediately
Pain Management in Eye Care
Conditions Which May Require Pain Management

- Large corneal abrasions
  - Cornea burn
  - PRK/PTK
- Orbital trauma
- Orbital blowout fractures
- Scleritis

Corneal burn with Curling iron
Pain Reliever Help

- Know your maximum daily allowances
- APAP 3000 mg (4000 mg*)
- ASA 6000 mg
- Ibuprofen 3200 mg
- Naproxen Sodium 1650 mg (Aleve/Anaprox)
- Naproxen 1500 mg (Naprosyn)
- Propoxyphene HCl 600 mg
- Codeine 240 mg
- Hydrocodone 60 mg
New Initiatives to Help Encourage Appropriate Use of Acetaminophen

New Dosing Instructions: Beginning Fall 2011
Acetaminophen, the active ingredient in TYLENOL®, can be found in more than 600 over-the-counter (OTC) and prescription medications, such as TYLENOL®, SUDAFED® Triple Action™, NyQuil®, Percocet® and Vicadin®. Acetaminophen is safe when used as directed, but when too much is taken (overdose), it can cause liver damage. Some people accidentally exceed the recommended dose when taking multiple products at the same time, often without realizing they contain acetaminophen or by not reading and following the dosing instructions.

To help encourage appropriate acetaminophen use, the makers of Extra Strength TYLENOL® have implemented new dosing instructions lowering the maximum daily dose for single ingredient Extra Strength TYLENOL® (acetaminophen) products sold in the U.S. from 8 pills per day (4,000 mg) to 6 pills per day (3,000 mg). The dosing interval has also changed from 2 pills every 4 – 6 hours to 2 pills every 6 hours.

Look for the new dosing instructions on Extra Strength TYLENOL® product packages in the U.S. beginning in the fall of 2011. Consumers can continue to use their existing TYLENOL® and other adult acetaminophen-containing products as currently labeled. The company has been working closely with other manufacturers of acetaminophen products to help ensure consistency in dosing instructions.

Education Campaign: Get Relief Responsibly™
In addition to the new dosing instructions on the OTC label, the makers of TYLENOL® launched Get Relief Responsibly™, an initiative designed to educate consumers about the appropriate use of OTC and prescription medications, particularly those containing acetaminophen, and the importance of reading and following medication labels. As a part of this initiative, the makers of TYLENOL® have created a new website www.getrelieffresponsibly.com. The site includes an interactive Acetaminophen Finder tool to help consumers identify products that contain acetaminophen and build a personal acetaminophen medication list to share with their healthcare provider or pharmacist.

Acetaminophen is used by more than 50 million Americans each week to treat conditions such as pain, fever and aches and pains associated with cold and flu symptoms. The safety of TYLENOL® has been well established for more than 50 years and TYLENOL® remains the brand of pain reliever that doctors recommend more than any other and that hospitals use most. If you have any questions, we encourage you to talk to your healthcare professional, or contact our Consumer Call Center at 1-877-414-7711.

*The third-party trademarks used herein are registered trademarks of their respective owners and not McNEIL-PPC, Inc.
**The maximum daily dose will also be lowered for Regular Strength TYLENOL® and other McNeil Consumer Healthcare adult acetaminophen-containing products beginning in 2012.
Maximum Tylenol Dose Lowered To Prevent Overdoses

Trenton, N.J. — Johnson & Johnson said Thursday that it’s reducing the maximum daily dose of its Extra Strength Tylenol pain reliever to lower risk of accidental overdose from acetaminophen, its active ingredient and the top cause of liver failure.

The company’s McNeil Consumer Healthcare Division said the change affects Extra Strength Tylenol sold in the U.S. — one of many products in short supply in stores due to a string of recalls.

Starting sometime this fall, labels on Extra Strength Tylenol packages will now list the maximum daily dose as six pills, or a total of 3,000 milligrams, down from eight pills a day, or 4,000 milligrams. Beginning next year, McNeil will also reduce the maximum daily dose for its Regular Strength Tylenol and other adult pain relievers containing acetaminophen, the most widely used pain killer in the country.

Besides Tylenol, acetaminophen is the active ingredient in the prescription painkillers Percocet and Vicodin and in some nonprescription pain relievers, including NyQuil and some Sudafed products. It’s found in thousands of medicines taken for headaches, fever, sore throats and chronic pain.

But people taking multiple medicines at once don’t always realize how much acetaminophen they are ingesting, partly because prescription drug labels often list it under the abbreviation “APAP.”

Two years ago, a panel of advisers to the Food and Drug Administration called for sweeping restrictions to prevent accidental fatal overdoses of acetaminophen.

Then in January, the F.D.A. said it would cap the amount of acetaminophen in Vicodin, Percocet and
Two and Two
Analgesic
- Ibuprofen
  - 200 mg x 2 = 400 mg
  - 400 mg TID/QID = 1200 mg
- Acetaminophen
  - 500 mg x 2 = 1000 mg
  - 1000 mg TID/QID = 3000 mg

Four and Two
Analgesic and Anti-inflammatory
- Ibuprofen
  - 200 mg x 4 = 800 mg
  - 800 mg TID/QID = 3200 mg
- Acetaminophen
  - 500 mg x 2 = 1000 mg
  - 1000 mg TID/QID = 3000 mg

Can replace ibuprofen with Aleve if desired
Crystalline Retinopathies

- Talc – intravenous drug use
- Tamoxifen - used in treating breast cancer
  - Daily 10-20 grams
  - > 1 year of therapy with a total of > 100 grams
- Canthaxanthin- used as an oral tanning agent
- Treatment
  - Stop the drug
  - No additional systemic testing is necessary if a history of medication or drug use is clear
Thank you!
Tracy and Greg