# Pain Management in Individuals with Developmental Disabilities

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### Objectives

- Discuss the best approach for assessment of pain in individuals with intellectual disabilities
- Differentiate between acute and chronic pain
- Analyze the characteristics of opioids vs. nonopioids
- Clarify terminology surrounding drug abuse and addiction
- Identify at least two strategies for optimum pain management in this patient population

### Prevalence of pain in Individuals with ID

- Valkenburg, et al. state that up to 50% of children and adults with Intellectual Disabilities experience gastroesophageal reflux disease
  - Significant pain and discomfort
- Assessment in this special population can be challenging when there is no self report
  - Then what?
- Things to consider....

### **During Assessment**

- Consider biases
  - Age
  - Gender
  - Attractiveness
  - Intellectual and physical abilities

# Assessment in the Cognitively Impaired

- Direct observation or history from caregivers
  - Assessment by proxy—nursing assistants or family members or regular caregivers
- Observe during movement (walking, morning care, transfers
- Unusual behavior should trigger assessment of pain

### Pain Indicator for Communicatively Impaired Children (PICIC)

### Most common cues identified by 67 parents:

- Screwed up or distressed looking face
- Crying with or without tears
- Screaming, yelling, groaning, moaning
- Stiff or tense body
- Difficult to comfort or console
- Flinches or moves away if touched

Ref: Stallard P, et al: Pain 98(1-2):145-149, 2002.

# Common Pain Behaviors in Cognitively Impaired Elderly Persons

- Facial expressions
- Verbalizations, vocalizations
- Body movements
- Changes in interpersonal interactions
- Changes in activity patterns or routines
- Mental status changes

### CNPI—at rest and with movement

- Vocal complaints: Non-verbal (Expression of pain, not in words, moans, groans, grunts, cries, gasps, sighs)
- Facial Grimaces/Winces (Furrowed brow, narrowed eyes, tightened lip jaw drop, clenched teeth, distorted expressions).
- Bracing (Clutching or holding onto side rails, bed, tray table, or affected area during movement)
- Restlessness (Constant or intermittent shifting of position, rocking, intermittent or constant hand motions, inability to keep still)
- Rubbing: (Massaging affected area)
- (In addition, record Verbal complaints).
- Vocal complaints: Verbal (Words expressing discomfort or pain, "ouch" "that hurts"; cursing during movement, or exclamations of protest "stop"; "that's enough")
- Feldt, K. S. (1996). Treatment of pain in cognitively impaired versus cognitively intact post hip fractured elders. (Doctoral dissertation, University of Minnesota, 1996). Dissertation Abstracts International, 57-09B, 5574.
  - Feldt, K.S. (2000). Checklist of Nonverbal Pain Indicators. <u>Pain Management Nursing, 1</u> (1), 13-21.

# Other Assessment Scales for Cognitively Impaired

- Pain Indicator for Communicatively Impaired Children
  - 10-49 yrs; non-verbal; indicators—facial, activity, vocal, consolability, physiological, individual indicators (0-24 point scale)
- Non-communicating Children's Pain Checklist postoperative version
  - 3-19 years; postop pain; indicators—facial, activity, vocal, social, physiological (27 items on a 0-81 point scale—11/81: mod to severe pain)
- Paediatric Pain Profile—postoperative pain
  - 1-18 years; indicators—facial, activity, vocal, social, consolability, physiological (20 items with 0-60 point 14/60: mod or worse pain

### More scales

- Checklist Pain Behavior—postoperative pain
  - 3-19 years; indicators—facial, activity, vocal, physiological; (10 point scale)
- FLACC (Revised Face, Legs, Activity, Cry, Consolability)—postoperative pain
  - 4-19 years; indicators—facial, activity, vocal, social, consolability, physiological and individual indicators; (5 items with 0-10 point scale—4/10: mod pain

### During Assessment also consider that....

- Pain can impact
  - communication
  - Socialization
  - Cognitive function
  - Combined with fear cognitive function may even be effected more
- Greater pain caused greater dysfunction across domains
- Pain had a greater impact on individuals with more severe ID

#### Less-obvious Pain Indicators

- May be attributed to psychosis or dementia
  - Aggressive behavior
  - Fidgeting
  - Noisy breathing
  - Rapid blinking
  - Rigid, tense body posture
- Untreated pain can increase confusion
  - Patients on opioids at risk for dose being cut

#### Assume Pain is Present

- Assume Pain is Present
- Is there a painful stimulus
  - Surgical incision
  - Fracture
  - Painful procedure
  - Any tissue damage
- If so, treat
  - Observe

### Types of Pain

- Nociceptive vs Neuropathic
- Physiologic vs pathophysiologic
- Acute vs chronic
- Malignant vs nonmalignant
- Pain syndromes

### Nociceptive Pain (Acute Pain/ Physiologic Pain)

Pain resulting from activation of primary afferent nociceptors by mechanical, thermal or chemical stimuli

### Pain Mechanisms: The "Pain Process"

• The neural mechanisms by which pain is perceived involve a process that involves four major steps:

- 1. Transduction
- 2. Transmission
- 3. Modulation
- 4. Perception



# Neuropathic pain Pathophysiologic Pain

Pain resulting from damage to peripheral nervous or central nervous system tissue or from altered processing of pain in the central nervous system

### Neuropathic—Pathophysiologic Pain

- Results in cellular changes that occur in peripheral and central nervous systems
  - Results in sensitization to the transmission of pain signals
- Neuroplasticity—ability of neurons to change their structure and function
- Peripheral and central sensitization response to stimuli is increased

# Result of Central and Peripheral Changes

- Hyperalgesia
  - Primary hyperalgesia
  - Secondary hyperalgesia
- Allodynia
- 'wind-up' of C fibers (a phenomenon of progressively increased neural response to repeated noxious stimuli)

### Chronic Pain—Subtypes

- Inflammatory
  - OA (27 million) and RA (1.5 million)
- Neuropathic
  - DN; PHN
- ▶ LBP—59 million
- Non-inflammatory, non-neuropathic pain
  - Fibromyalgia—5 million
  - CRPS

### RSD/CRPS

- Reflex Sympathetic Disorder/Complex Regional Pain Syndrome
- An extreme example of chronic severe pain
- Can occur after any type of injury-small or large; surgery; burn
- Pain is as severe as the initial injury
- Effects the nervous system and can have swelling, discoloration, sweating to the effected area
- Allodynia is a major symptom

### **CRPS**

http://www.abc.net.au/catalyst/stories/2621515.htm

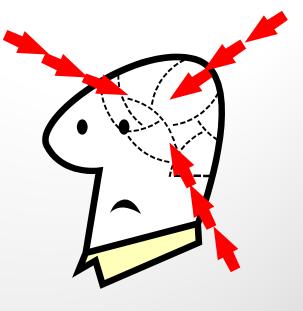
### Multiple Dimensions of Pain

### The ABCs of Pain

Affective Dimension

Behavioral Dimension

Cognitive Dimension

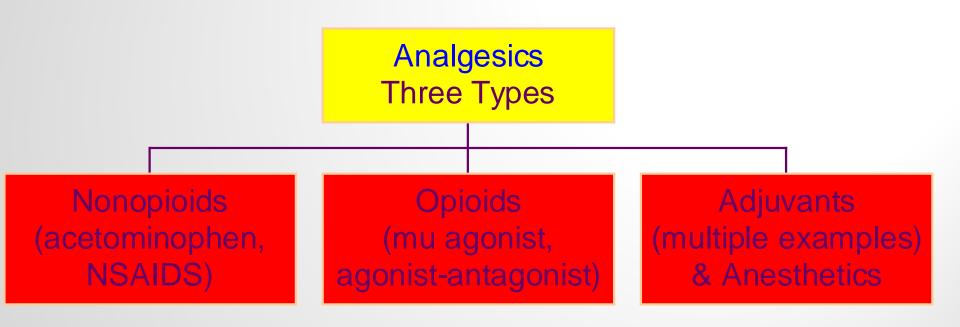


Physiological-Sensory Dimension

### **Definition of Pain**

"Unpleasant sensory and emotional experience arising from actual or potential tissue damage or described in terms of such damage (IASP)

### Medication Management--Analgesics



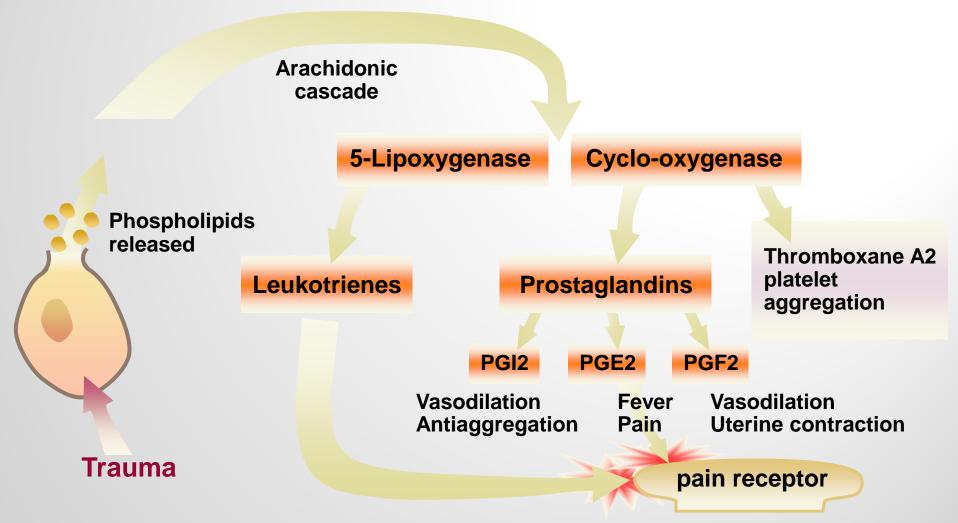
### Acetaminophen

- Mechanism of action is not certain
- Probably centrally acting—?cox-3 inhibitor
- Acetaminophen toxicity
  - Hepatotoxity
    - Toxic metabolite (NAPQI)
    - Several other mechanisms lead to hepatotoxicty
    - Mechanism not completely understood
  - Nephrotoxicity >4g/day for long periods
    - Uncertain cause
    - May be caused by activity of NAPQI in renal microsomes
    - Increase frequence to 6-8 hrs in renal failure

#### **NSAIDS**

- NSAIDS—Antiinflammatory, antipyretic, analgesic
- Mechanism of action—prostaglandin inhibition by way of COX-1
  - Prostaglandins
    - important in maintaining integrity of GI and duodenal mucosa
    - Important in modulating renal plasma flow
- NSAIDs inhibit formation of thromboxane effecting platelet aggregation
- Use with caution in pts. with history of asthma
  - Inhibits prostaglandin E—responsible for bronchodialation

# Transduction: Nociceptive Chemical Stimuli



Class	Generic name	UAD	Brand name
Proprionic acids	Naproxen	500 mg initially- followed by 250mg q6-8h	Naprosyn, Anaprox, Alleve
	Flurbiprofen		Ansaid
	Oxaprozin		Daypro
	Ibuprofen	400-800mgQ6-8h 25-75 mg Q6-8h Max 120mg/d (parenteral)	Motrin
	Ketoprofen		Orudis, Oruvail
	Ketorolac		Toradol
Indoleacetic acids	Sulindac	200mg Q12h	Clinoril
	Indomethacin	25-50mg q8h 200-40mg q6- 8h	Indocin Lodine
	Etodolac		

Class	Generic name	UAD	Brand name
Phenylacetic acids	Diclofenac	50 mg/q8h	Cataflam, Voltaren
Salicylic acids (nanacetylated)	Salsalate Choline magnesium trisalicylate	1000-1500 mg/q12h 1000-1500 mg/q12h	Disalcid Trilisate
Naphthylalkanone	Nabumetone	1000-2000 mg/day	Relafen
oxicam	Piroxicam		Feldene

#### COX-2 Inhibitors

- May have fewer GI effects than COX-1 inhibitors
- Should be avoided in patients with creatinine clearance <30ml/min</p>
  - Carry same risk as traditional NSAIDs
- Celecoxib—Celebrex
  - UAD=100-200 mg q12h max=400 mg/d

### CLARIFYING TERMINOLOGY

### ADDICTION—WHAT IS IT??

- Is it a moral deficit in which the person freely chooses...
- a criminal offense...
- or is it a physiologic disease?

#### Prevalence of Addiction

- 6-10% of the general population has an addiction to illicit drugs, prescribed opioids and alcohol
- ▶ In chronic pain populations—6–10%
  - Those with a history of previous addiction
- Chronic pain alone does not add to the risk of addiction
- Rate of addiction in patients without a previous history of addiction when taking opioids for pain remains to be ~1%

### Clarifying Definitions (AAPM, APS, ASAM 2001)

- Physical Dependence: Adaptation that is manifested by a drug class specific withdrawal syndrome that can be produced by abrupt cessation, rapid dose reduction, decreasing blood level of drug and/or administration of an antagonist
- "Dependence" is used by addiction specialists referring to addiction

### **Definitions**

- Tolerance: A state of adaptation in which exposure to a drug induces changes that result in a diminution of one or more of the drug's effects over time
- Physical dependence, tolerance and addiction are separate phenomena but may co-exist

#### Addiction

- Addiction is defined as a chronic, relapsing brain disease that is characterized by compulsive drug seeking and use, despite harmful consequences
  - National Institute of Health 2010

#### Pseudoaddiction

- Behaviors that mimic drug abuse
  - Drug seeking
  - Clock watching
  - Anticipate the next dose
  - Demand more pain meds
- Due to the undertreatment of pain
  - Patients may become deceptive
  - May even resort to the purchase of illicit drugs
- Distinguished from addiction analgesia demonstrates improved function and use the med as prescribed not for sedation or euphoria

#### Clarification of Terms

- Substance misuse
  - Use other than intended purpose
- Substance abuse
  - Use that is unlawful or detrimental
- Diversion
  - given, sold, or traded to someone other than the patient for whom it was intended
- Nonmedical use
  - Taking the drug for the feeling it gives

## "Opiophobia"

- Fear that opioids will cause addiction
  - Up to 90% of the US population above age 12 has experimented with illicit drugs or alcohol
  - Very small percentage go on to develop substance abuse
  - Treating pts with a hx of addiction will cause relapse
    - In reality, pain is more likely to cause relapse

### DSM-IV describes "Substance Dependence" as

- A maladaptive pattern of substance use manifested by at least 3 of the following occurring any time over a 12-month period
  - Tolerance
  - Withdrawal
  - Taking larger amounts over a longer period than was intended
  - Unsuccessful efforts to cut down
  - A lot of time spent in efforts to obtain the substance
  - Important activities given up because of use
  - Continuation of substance despite knowing that is causing problems

#### Characteristics of Opioids

- No ceiling effect
- Usually no end organ damage with chronic use
- Metabolized by the liver
  - Metabolite toxicity
    - Avoid using meperedine and propoxyphene
- Excreted by the kidney
- Cause tolerance and physical dependence
- Reversible with an antagonist
- Bind to opiate receptors— $\mu$ ,  $\kappa$ , δ
- Tolerance to side effects except constipation

#### **Pharmacokinetics**

- Absorption
  - Drug solubility—lipophilic vs hydrophilic
- Bioavailability
- First pass Effect
- Solubility
- Metabolism metabolites, active or inactive
  - Prodrugs, e.g. codeine metabolized by CYP450 enzyme CYP2D6
- Half-life, clearance, steady state and accumulation

### **Pharmacodynamics**

- Opioid responsiveness
  - Efficacy—extent to which a drug "works" (as compared to others)
  - Potency—the dose of a drug required to produce a specified effect, e.g. hydromorphone > potency than morphine
  - Opioid responsiveness—affected by age, organ dysfunction
- Tolerance—rule out disease progression; compliance to tx
  - OIH—rare
  - Incomplete Cross-tolerance—due to receptor subtypes—reduce new opioid 25% - 50% calculated equianalgesic dose (methadone dec. by as much as 90% then titrate as needed
- Physical dependence

#### PRN

- What does "PRN" mean?
- If pain is ongoing give opioids ATC
- Half-life
- Steady state
- Time to peak effect

# Opioids Mu-agonists

Bind to mu opiate receptors blocking transmission of pain

- Morphine
- Fentanyl
- hydromorphone
- oxycodone
- hydrocodone

- Codeine
- \*Methadone
- \*meperedine
- \*tramadol

## Morphine

- Hydrophilic—delayed onset and longer duration
- Two metabolites but only one active at opioid receptor—morphine-6-glucuronide (M6G)—analgesic
- Patients with renal impairment should start at ¼ dose and titrate as needed
  - Accumulation results in neurologic side effects as well
  - Removed with dialysis

### Hydromorphone (Dilaudid)

- Hydrophyllic—similar to morphine
  - IV—1.5 mg:10 mg morphine/PO—7.5 mg:30 mg morphine
    - Onset 5 min; peak in 8-20 min. duration ~ 4 hrs
  - Oral
    - 60% bioavailable; onset 30 min. duration 3-4 hrs
  - Metabolized in the liver
  - Several metabolites
- Use decreased amounts in renal impairment due to possible sensitivity to hydromorphone-3-glucuronide→possible neuroexcitation
  - there is no 6-glucuronide so may have fewer SEs
- May be safer than morphine in renal insufficiency

# Fentanyl

- ▶ Lipophillic→Short half-life, short duration of action
  - UNLESS given regularly—then half-life is extended
- No active metabolites
- Safer in renal and liver failure
- Fewer side effects
- Half-life extends with continuous use
- Multiple formulations
  - transdermal, oral transmucosal, buccal

### oxycodone

- Available in combination or single-entity
  - Short and Long–acting
- More potent than morphine
- Metabolized in the liver by cytochrome CYP2D6
  - Multiple metabolites
- Binds at µ and κ receptors—may be better in chronic pain states
  - Half-life and bioavailability slightly longer than MS
  - One active metabolite—oxymorphone
  - Women may have a greater effect
  - Excretion impaired in uremic patients and
    - Elimination half-life is severely impaired in these patients
  - May cause CNS toxicity and sedation in renal failure

# Hydrocodone (Vicodin)

- Only available in combination with acetaminophen, ibuprofen, aspirin
  - Onset 20 min. peak by 60 min; half-life 3.8 hrs
- Metabolized in the liver
  - Several metabolites
- Significant renal excretion of active forms
- Should be avoided in patients with renal failure
- Adverse effect -hearing loss

### Demerol (meperedine)

- Half-life is 2-3 hrs (parenterally)
- Bioavailability from p.o. is ¼ that of parenteral
- Onset 10 minutes; peak effect 30 min. duration up to 4 hrs
- More likely than other opioid drugs to cause delirium in postop pts of all ages
  - More nausea and vomiting
- Limit use to 600 mg/d and no more than 48 hours due to metabolite—normeperedine
- Observe for signs of neuroexcitation restlessness, shakiness, tremors, twitching and jerking
- Misconception—produces less biliary spasm than other opioids—all opioids can produce this

#### Normeperidine

- Normeperedine—only active metabolite of meperedine
  - toxic metabolite
  - half-life 15-20 hrs
  - causes neuroexcitation—hyperreflexia, myoclonus, agitation and grand mal seizures
  - half analgesic potency but twice the toxicity
  - Is not reversed with naloxone and may increase risk of seizures if naloxone given
- Use extreme caution in patients with seizure disorder
- Use caution in patients with renal insufficiency
- Contraindicated with MAOI (monoamine oxidase inhibitors)—can cause serotonin syndrome or death

#### Codeine

- ▶ 60mg = 600 mg of aspirin
- Not appropriate for moderate to severe pain
- Usually more constipating
- Has more psychotomimetic effects
- Metabolized in the liver to morphine
  - Several metabolites
  - Metabolism is necessary for analgesia
  - Poor metabolizers may show absence of analgesia
- Reduced renal clearance in advanced renal failure
  - Reports of serious adverse effects in renal failure

## Methadone—good news

- Inexpensive
- Adverse effects similar to other opioids
- Rapid onset—30-60 minutes; duration 4-6 hrs; peak effect 2.5 hrs
- ~ 80% bioavailability
- No active metabolites
- Long duration with continued use
- No ceiling dose other than side effects
- Has some SSRI and NMDA antagonist activity
- ▶ For opioid naïve patients → start at 2.5mg Q8H
- Excreted in feces—considered safe in renal insufficiency

#### Methadone—not so good news

- Long half-life—15-60 hours-
  - Unpredictable
  - difficult to titrate
  - Difficult to convert from other opioids to methadone
- Duration initially is 3-6 hrs→8-12 hr with repeated dosing
- Varied inter-individual effects
- Efficacy is greater with repeated dosing
- Multiple drug-drug interactions that can induce or inhibit effect by other drug or be effected
  - Close observation is required

### Propoxyphene (Darvocet)

- REMOVED FROM THE MARKET IN 2011—YAY!
  - Was removed from the market in the UK many years ago

#### Dual-mechanism Analgesics

- Tramadol—for mod to moderately severe pain
  - Weak mu-agonist and norepeniphrine and serotonin reuptake inhibitory activity similar to TCAs
  - Peak effect in ~ 2 hrs of 100mg dose
  - Potency equivalent to codeine and five times less potent than morphine
  - Ceiling effect
  - Max dose is 400mg/24h
  - Use with caution in pts w seizures or on SSRIs
- Tapentadol Nucynta
  - Agonist at mu and blocks reuptake of norepinephrine
  - Schedule II drug
  - Indicated for mod-severe pain
  - Avoid combining with SSRIs

### **Titration of Opioids**

- Based on effect
  - Increase dose 25%–100%
    - Ask patient how much pain was relieved by last dose
- Estimate 24 hr total and change to longacting formula...for example
  - 2 tabs 5/325 Percocet Q4H→20 mg OxyContin Q8H

#### Equianalgesic Dosing Guidelines

- Equianalgesic means approximately the same pain relief
- The chart is a guideline. Titrate meds according to pt's response
- Chart is helpful when switching from one drug to another or when switching to another route
- Dosages are not necessarily starting doses
- Consider incomplete cross-tolerance

Drug	Oral Dose	IV Dose	Duration
Morphine	30 mg	10 mg	3-5 hours
Fentanyl	Breakthrough only (OTFC)	100mcg (0.1mg) 100 mcg/h TD $\approx$ 4 mg/h IV MS; 1mcg/h TD $\approx$ 2 mg/24 h oral MS	0.5-1 hour
Hydromorphone	7.5 mg	1.5 mg	2-4 hours
Meperidine	300 mg NR	75-100 mg	2-4 hours
Codeine	200 mg NR	130 mg	3-4 hours
Methadone			
Oxycodone	20-30 mg		3-4 hours
Hydrocodone	30 mg		3-4 hours
Nalbuphine		10 mg	3-6 hours

# Opioid Side Effects— Are all self-limiting except....

- Constipation
- Nausea and vomiting
- Pruritus
- Urinary retention
- Mental status changes
- Sedation
- Respiratory depression

# PREDISPOSING FACTORS TO RESPIRATORY DEPRESSION

- Sedation
- Large doses of opioids
- Concomitant use of opioids and CNS depressants
- High-risk patients
  - obese
  - hx of pulmonary disease
  - hx of sleep apnea
  - advanced age (>65 years)
- Rarely seen in chronic pain management

#### **Adjuvant Analgesics**

- Medications that are typically used for another purpose
- Two classes
  - Multipurpose acute and chronic pain
  - For specific types of pain

# First-line Drugs in Chronic Pain

- gabapentin (Neurontin)—start w/100-300 mg/day Usual Effective Dose (UED) 300-3600 q8h
- pregabalin (Lyrica)—start with 100-150 mg/day; UED 150-600 q12h
- SNRI
  - Duloxetine (Cymbalta)—start w/ 30 mg/day; UED 60 mg q12h

# ADJUVANT ANALGESICS: MAJOR CLASSES

- Anticonvulsants
- Antidepressants
- Psychostimulants
- Muscle relaxants
- Sedatives

# Opioid Side Effects

- Nausea and vomiting
- Pruritus
- Urinary retention
- Mental status changes
- Sedation
- Respiratory depression

#### **Opioid Induced Constipation**

- The hand that writes the prescription for an opioid and
- Fails to write the order for a laxative should be
- The hand that removes the impaction

### Adjuvant Examples

- Antidepressants
  - SSRIs, SSNRIs, TCAs
- Anti-convulsants
- Corticosteroids
- Alpha-2 adrenoceptor agonists
- Anti-histamines

- Anti-spastics
- Muscle Relaxants
- NMDA receptor antagonists

# **Balanced Analgesia**

- Inter-disciplinary approach
  - Medication management
  - Physical activity
  - Maximize nutritional contributions
  - Mental health
  - Support group
  - Spirituality

#### Non-Pharmacologic Interventions

- Increase activity
- Individualize interventions
  - music
  - artwork
  - humor
- Address constipating effects of opioids

# **Goal Setting**

- Once assessment complete, discuss pain level and related goals with patient & family
- Should be based on functionality
- Decrease suffering
- Be realistic
- Be patient
- Patient and family education—why pain management
  - Minimize risk of complications
  - Myths about addiction

#### Documentation

- Analgesia
- Adverse effects
- ADLs
- Aberrhant behavior

## In Summary

- Treat pain initially aggressively—
  - \*\*\*Titrate to Effect
- Adequate analgesia results in:
  - Early participation in activity
  - Prevention of complications
  - Decrease risk of chronic pain
  - Early return to individual level of functioning
- Use assessment tool specific to population
- Always combine modalities— opioids with nonopioids and pharmacologic with nonpharmacologic