# Cancer Pain

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#### Prevalence of the Most Common Symptoms in Advanced Cancer (1000 Adults)

Symptom	%	Symptom	%	
Pain 82		Lack of Energy 59		
Easy Fatigue	67	Dry Mouth	55	
Weakness	64	Constipation	51	
Anorexia	64	Dyspnea	51	
>10% Wt Loss	60	Early Satiety 50		

Donnelly and Walsh Semin Oncol, 1995

# Pain definition

Pain is an emotional and sensory experience or described in those terms

### Incidence Of Cancer Pain

- 50 % of all cancer patients
- 70-80 % of all advanced cancer patients
- 50 % will have moderate to severe pain
- 30 % will have severe pain

#### Types Of Pain in Cancer

- Related to direct tumor involvement-60-65 %
- Related to cancer treatment-20-25 %
- Unrelated to cancer-10-15 %

### Pain Syndromes

#### **NOCICEPTIVE:**

#### A.Somatic:

Sharp, localized, aching, throbbing, gnawing

#### **B.Visceral**:

Dull, poorly localized, crampy, nauseous, squeezy, pressure **NEUROPATHIC**:

Burning, tingling, shooting, stabbing, itching, electric like, numb

## Types of pain

- Nociceptive pain somatic; Post-op bone pain chest pain mucositis
- Visceral; PCP, Liver mets, pleural

- Neuropathic pain:
- Peripheral neuropathy PHN Brachial Lumbosacral plexopathy
- Central: Stroke&MS,Tum

#### Pain Assessment

Detailed pain history-

- <u>Site</u>
- intensity
- Factors influencing pain
- Breakthrough Pain
- Medication history
- Associated neurological deficits
- Psychosocial history
- Assessment of other symptoms

#### Intensity Of Pain

0-10 (Verbal numeric scale) Mild, moderate, severe 0-10cm Line (Visual analogue scale) Faces Fruits Currency

Sometimes cannot quantify

#### Happy/Sad Faces

(Memorial Sloan-Kettering, © 1990)

#### Assessment Of Pain

**Questionnaires:** 

#### Wisconsin Brief Pain Inventory:(Cleeland) (BPI)

Captures pain at different times Functional interference is assessed Other symptoms are assessed

Well tested, easy to administer

#### Assessment Of Pain

**Graphic Scale:** 

- Edmonton Symptom Assessment Scale (ESAS) :
- Easy to administer, captures other symptoms on a 0-10 scale format.
- Easy follow through and can act as a flow chart.

#### ESAS

#### THE UNIVERSITY OF TEXAS MDANDERSON CANCER CENTER

#### FOLLOW-UP AND PROGRESS NOTES

Svn	notom Contr	01 & 1	Palliatis	e Car	. Su	moth	4.	earre		Cast.	
<b>Referral Date</b>					e oji	-	1. 113	32337	10714	scure	
Date: Marc	h	20	21/22	123	24	25	T	T			-
Pain	(0-10)*		0. 00	4.5	ar	20	=		=		-
Fatigue	(0-10)*						-				
Nausca	(0-10)*										
Depression	(0-10)*										
Anxiety	(0-10) *									-	-
Drowsiness	(0-10)*										
Shortness of Bre	ath (0-10)*				-				-		
Appetite	(0-10)*			-	_				-		-
Sleep	(0-10)*									-	
Feeling of Wellb	being (0-10)*								-		
Mini Mental Sta (0-30)	te Score		8	0						-	
Assessment from: (If SO or HCP -	Pt/SO/HCP use red ink)										-
Total Opioid MEDD: mg.	/day										
Staff Initials (Sig	gnature &						-	-		-	-

Death

#### Assessment Of Cancer Pain

Pain:

"A new pain or a change in character of an old pain in cancer patient is always because of cancer unless otherwise proved"--Reddy

# **Multifactorial Nature of Pain**



(Adapted from Portenoy, 1988)

# Treatment of Cancer Pain

Pharmacotherapy Opioids & Non-opioids Non-Pharmacotherapy Behavioral treatment Physical therapy Counselling Anesthetic procedures

### Pharmacotherapy of cancer pain

Non-opioid drugs:

NSAIDS Antidepressants Anticonvulsants Corticosteroids Phenothiazines Benzodiazepines **Opioid drugs:** Weak opioids Strong opioids





# Adjuvant Analgesics(Step 1 WHO)

Non-Steroidal anti-inflamatories (NSAIDS)

Antidepressants(TCA)

Anticonvulsants/Anti-epileptics(AEDs)

Corticosteroids

Bisphosphonates

Anesthetics-Lidoderm Patch

N-Methyl D-aspartate antagonists(NMDA)

Radiopharmaceuticals

Phenothiazines

Benzodiazepines

#### NSAIDS

- Acetaminophen (Paracetamol) can cause liver damage if dose exceeds 3-4 gm a day
- Liver damage risk increased in alcoholics and preexisting liver damage.
- Acetaminophen (Paracetamol) has been shown to cause renal damage

Insel PA: Goodman Gillman ed 8 1990

## NSAIDS(Ibuprufen,Naprosyn)

- Acts by inhibiting cyclooxygenase to decrease prostaglandin synthesis
- May have central action at the spinal cord level
- Exhibit ceiling effect
- Tolerance and physical dependence is not seen
- Cause end organ toxicity
- Cox -2 inhibitors cause less GI and Platelet dysfunction ( Other side-effects same)(Celecoxib)
- Vane JR: Inhibition of prostaglandin synthesis as mechanism of action of aspirin like drugs.

## Weak opioids/Step2 WHO/C3

- Codeine
- Hydrocodone
- (Propoxyphene)
- Tramadol

- Used for mild to moderate pain
- Not available as a single agents
- Dose limited by acetaminophen dose



### Agonists-Antagonists

- Pentazocine
- Nalbuphine
- Butorphanol
- Buprenorphine

- Demonstrate ceiling effect
- Precipitate withdrawal symptoms
- Dose-dependent psychotomimetic effects

## Strong opioids(Step 3 WHO)

- Used for all pain types
- Available as short acting and long acting
- Opioids with long half-life not the 1st choice
- Morphine -drug of first choice, different formulations available
- SRM-15, 30, 60,100 and 200mg.
- M.S. -10,15,30,60mg.

# WHO Ladder

#### Step 1:

Pain syndrome: Any or specific type of pain

Pain intensity:mild,0-4/10

Medications used: acetaminophen, anti-inflamatory, /TCA/AED

Response: somatic/neuropathic pain syndromes respond mildly.

WHO Ladder Step 2: Pain syndrome: Any or specific type Pain intensity:moderate,4-7/10 **Medications: Mild** opioids/NSAID's /TCA/AEDs **Response:Varies.** 

# WHO Ladder

#### Step 3:

Pain syndrome-Any or Specific

Pain intensity:moderate to severe,7-10/10

**Medications:Strong** 

opioid(Morphine

class)+NSAIDs,AEDs,TCAetc

Response:Good,80-90 %

#### Incidence Of Cancer Pain

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WHO Ladder Principles The five essential concepts in the WHO approach to drug therapy of cancer pain are:

By the mouth By the clock By the ladder For the individual With attention to detail

# Morphine

Gold standard for opioids

Acts as mu agonist in spinal cord, brain, and periphery

Metabolized to M6G, and M3G(May be responsible for excitatory side-effects

Metabolites accumulate in renal dysfunction-Hence adjust the dose

Available in all formulations and all routes

Start with 10-15 mg q 4 hrs initially and titrate the dose. Introduce sustained release form once pain is stable

Use IV for acute titration

**Transdermal Fentanyl** Fentanyl in a patch form Forms depot under the skin and releases fentanyl slowly into the system Takes 18 hrs to form depot Changed every 72 hrs, in some every 48 hrs-convenient **Useful for stable pain** Difficult to titrate in acute situations **?causes less constipation** 

## Selecting The Route

- Oral is the preferred route
- Transdermal route useful in patients with stable pain and cannot swallow, due to oral pathology or intractable nausea / vomiting
- Rectal route used for short term if N/V is due to GI pathology. M.S. can be given rectally
- IV Route for acute pain situations

#### WHO Analgesic Ladder



Step 1

## PRINCIPLES OF PHARMACOTHERAPY

- Match drug to pain syndrome
- Have low threshold to prescribe opioids
- Add adjunct medications where appropriate
- Oral route should be the route of choice
- Use IV route for acute titration
- Treat side-effects before switching opioids



### Treatment of somatic pain

- NSAIDs
- Mild opioids
- Physical modalities
- Psychotherapy
- Stronger opioids
- Interventions

### Treatment of Neuropathic pain

- Adjuvants:TCA, Gabapentin, Steroids, NSAID
- Stronger opioids
- Methadone
- IV Lidocaine/Mexilitene
- NMDA receptor antagonist-Ketamine/Dextromethorphan
- Interventions:Regional,Sympathetic blocks,
- Neuro-axial medications:opioids, clonidine, local anesthetic

# Difficult Pain Syndrome/Intractable Pain

Intractable pain syndrome is defined as persistent pain despite reasonable efforts to treat it.

## Adverse-effects of opioids

#### Common:

- Constipation
- Nausea/vomiting
- Somnolence
- Cognitive effects
- Dysphoria
- Myoclonus
- Itching
- Urinary retention

#### Rare:

- Histamine release
- Chest wall rigidity
- Decreased immunity
- Headaches
- Blurring of vision
- Respiratory depression
- Seizures

#### Practice

- Pharmacotherapy treats majority of pain(90%).
- Occasionally need anesthetic intervention either as an adjunct or prime reliever of pain(10%).
- Define pain syndrome before block.
- Never deny a patient of alternative pain therapy.

Principles of anesthetic procedures in cancer pain

- Usually reserved for patients who fail extensive pharmacological trials
- Should fail to show any dose response or develop dose limiting side-effects, which are resistant to treatment
- Neurolytic or destructive procedures are usually done in terminal patients due to adverse effects

# Anesthetic Procedures

The most useful Procedures:

- **1.Celiac plexus block**
- 2. Epidural infusions
- 3. Vertebroplasty
- 4.Intra-thecal neurolytic blocks
- 5. Intrathecal pumps

Issues in patients on chronic opioid medications

- 1. Tolerance
- 2. Physiological dependence
- 3. Psychological dependence

# **Multifactorial Nature of Pain**



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