

# MULTIMODAL ANALGESIA

## STRATEGIES FOR PERIOPERATIVE PAIN MANAGEMENT

JOSEPH KAY MD FRCPC  
Department of Anesthesia  
University of Toronto



**Sunnybrook**  
HEALTH SCIENCES CENTRE



# Outline

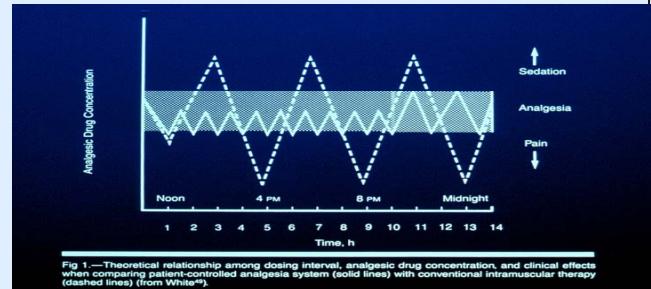
- Why treat pain aggressively?
- Pain pathways
- Components multimodal analgesia
- Clinical pathways

# Why Treat Pain Aggressively

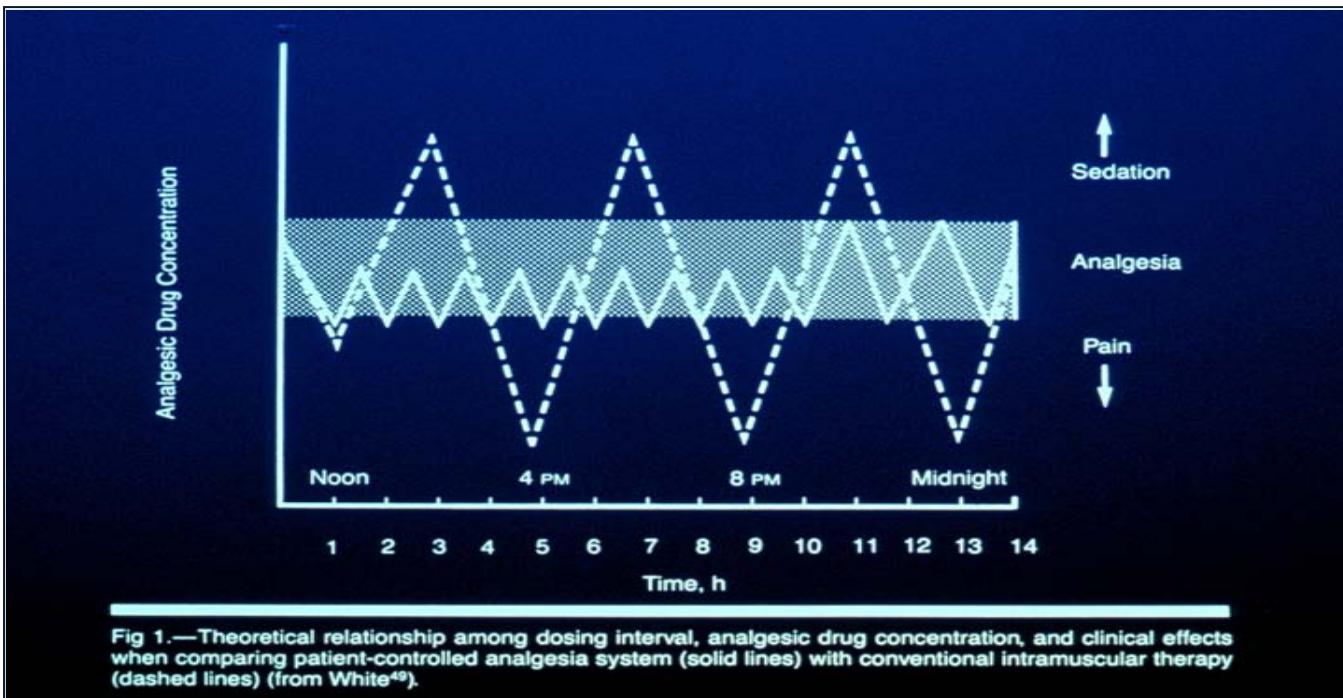
- Reduce complications
- Improve rehabilitation
- Improve outcome
- Prevent chronic pain

# Traditional Analgesia

- Parenteral opioid
- im/sc prn
- iv PCA



# PCA



# Opioid side-effects

- Nausea /vomiting
- Sedation
- Respiratory depression
- Pruritus
- Urinary retention
- Constipation
- Hyperalgesia / Tolerance
- Immunity

# Multimodal Analgesia

## ADD NON-OPIOID ANALGESICS

- To decrease opioid requirements
- Decrease opioid related s/e
- Improve dynamic analgesia
- Improve rehabilitation /outcome
- Reduce chronic pain

# Improved Outcome

- Local infiltration / FNB: THA, TKA  
↓ LOS, better flexion
- Thoracic Epidural: Colon resection  
↓ ileus, ↓ LOS, improved QOL
- PVB/Thoracic Epidural: ↓ cancer recurrence

ANDERSEN ACTA ORTHOP 2007 78:180 CARLI ANESTHESIOLOGY 2002 97:540  
CHELLY J ARTHROPLASTY 2001 16:436 WANG REG ANES PAIN MED 2002 27:139  
WERNER ACUTE PAIN 2005 7:5 BIKI ANESTHESIOLOGY 2008 109: 180  
EXADACTYLOS ANESTHESIOLOGY 2006 105:660

# Improved Outcome

- Celecoxib     ACL, TKA, laparoscopy
- Gabapentin    TKA, ACL, Breast
- Dexamethasone   LAP CHOLE, TL
- Oxycontin    TKA

BUVANENDRAN JAMA 2003   MENIGAUX ANESTH ANALG 2005  
FASSOULAKI ANESTH ANALG 2005   BISGAARD ANN SURG 2003  
WHITE CAN J ANESTH 2007   PETERS J ARTHROPLASTY 2006   ANDERSEN  
ACTA ORTHOPEDICA 2007   TOFTDAHL ACTA ORTHOP 2007   CHEVILLE J BONE JT  
SURG 2001   WHITE CJA 2007

# Chronic Pain

## THE PREVALENCE OF CHRONIC POST SURGICAL PAIN IN CANADA

Elizabeth G VanDenKerkhof DrPH, David H Goldstein MSc FRCPC

Department of Anesthesiology, Queen's University, Kingston, ON K7L 2V7

- >30,000 new pts chronic pain 1999

Procedure	No. performed	Prevalence estimates	Estimated prevalence
Hysterectomy	55,404	16%-50%	8,865-27,702
Cholecystectomy	>50,000	21%-27%	>10,500->13,500
Hip Replace	19,853	3%-35%	596-6,949
Knee Replace	21,649	30%	6,495
Breast surgery	14,438 <sup>+</sup>	13%-49%	1,877-7,219
Thoracotomy	16,305 <sup>++</sup>	7%-67%	1,141-10,924
<b>Total</b>	<b>&gt;177,649</b>	<b>3%-67%</b>	<b>29,474-&gt;72,789</b>

# Prevent Chronic Pain

## RISK FACTORS

Anesthesiology 2000; 93:1123-33

© 2000 American Society of Anesthesiologists,

### ***Chronic Pain as an Outcome of Surgery***

#### *A Review of Predictive Factors*

Frederick M. Perkins, M.D.,\* Henrik Kehlet, M.D., Ph.D.†

- Preoperative pain present >1 mo
- Operative nerve damage
- **Severe acute postoperative pain**

# Prevent Chronic Pain

## Multimodal Analgesia with Gabapentin and Local Anesthetics Prevents Acute and Chronic Pain After Breast Surgery for Cancer

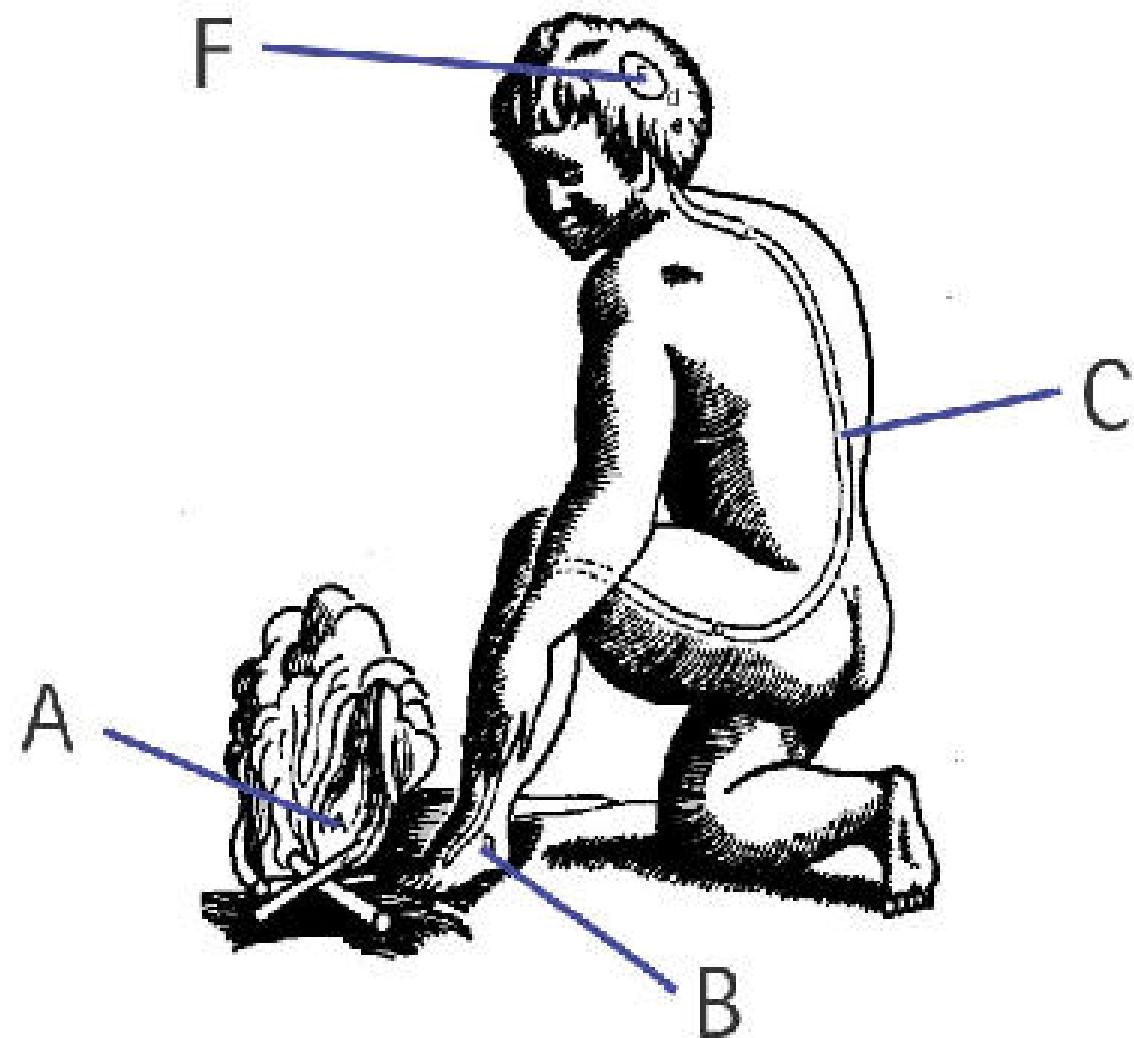
Argyro Fassoulaki, MD, PhD, DEAA\*, Argyro Triga, MD†, Aikaterini Melemeni, MD\*, and Constantine Sarantopoulos, MD, PhD, DEAA†

**Table 2.** Patients with Chest, Axillary, Upper Arm, and Overall Chronic Pain, Absent or Decreased Sensation, and Patients who Required Analgesics at Home 3 and 6 mo After Surgery

No. of patients	3 mo			6 mo		
	Control n (%)	Treatment n (%)	P-value	Control n (%)	Treatment n (%)	P-value
Chest pain	7/22 (32)	7/22 (32)	1.00	5/21 (24)	3/20 (15)	0.697
Axilla pain	10/22 (45)	3/22 (14)	0.045	6/21 (29)	3/20 (15)	0.454
Arm pain	13/22 (59)	5/22 (23)	0.038	7/21 (33)	3/20 (15)	0.277
Chronic pain (total)	18/22 (82)	10/22 (45)	0.028	12/21 (57)	6/20 (30)	0.151
Absent or decreased sensation	17/22 (77)	16/22 (73)	1.00	17/21 (81)	13/20 (65)	0.424
No. of patients who needed analgesics	5/22 (23)	0/22 (0)	0.048	4/21 (19)	0/20 (0)	0.107

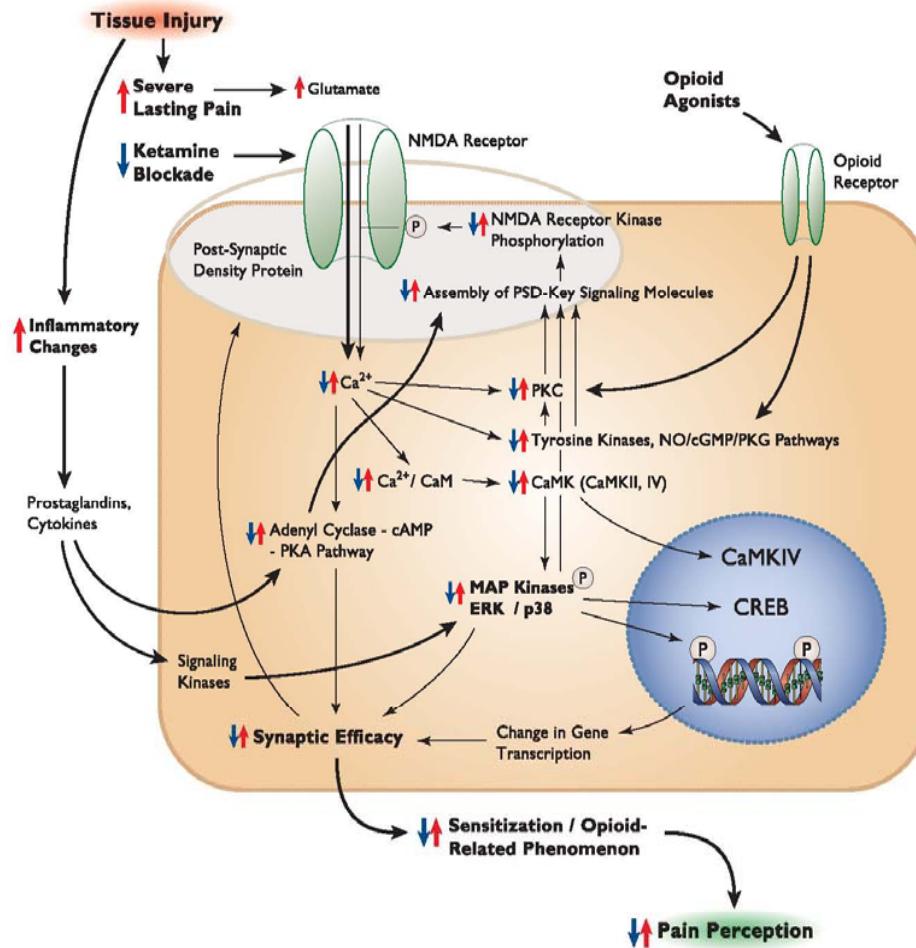
- EMLA cream x 3d, ropivacaine irrigation

# Pain Pathways

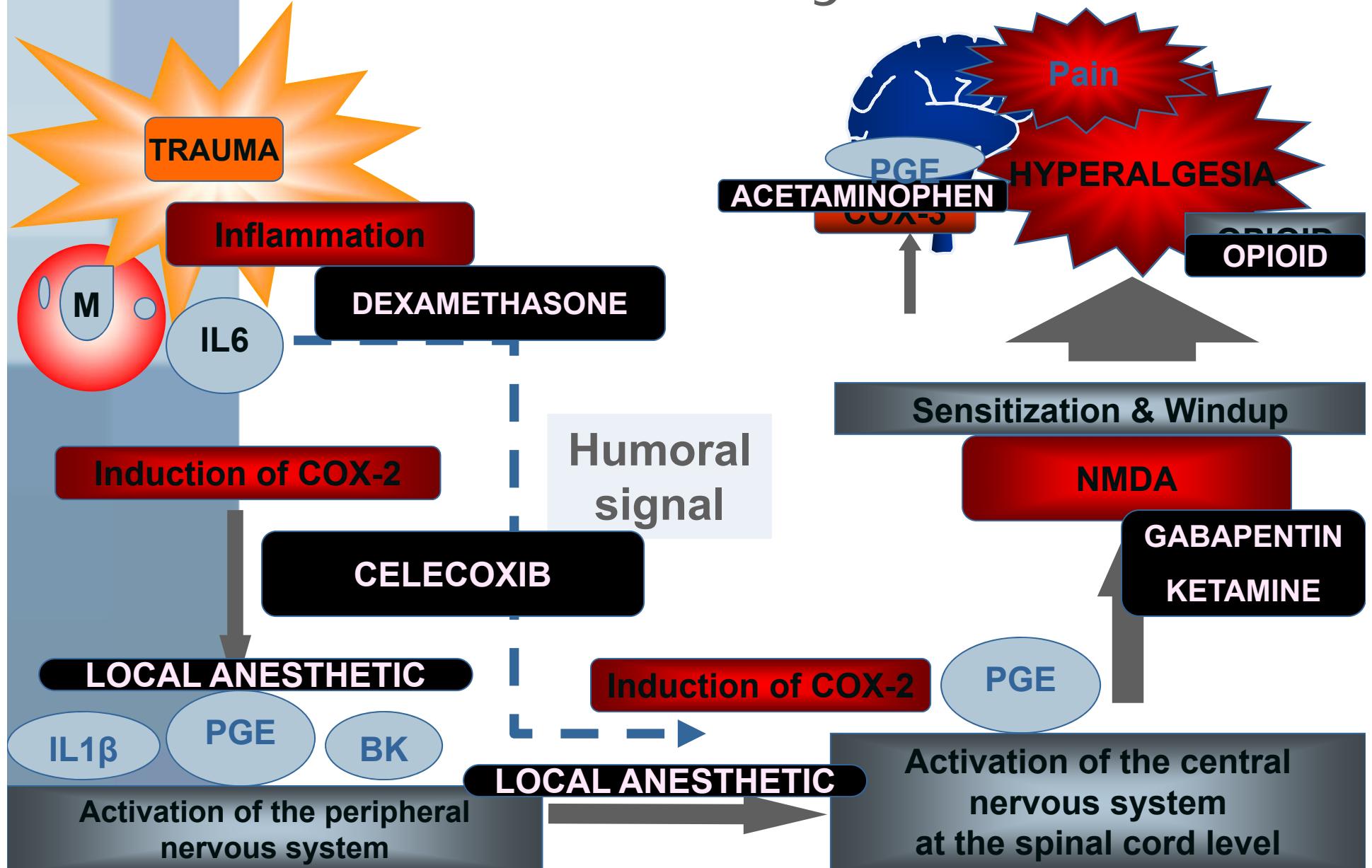


*Figure 1 Descartes' model of pain*

# Pain Pathways



# Pain Pathways



# Multimodal Analgesia

Using more than one drug, acting at a different place or with a different mechanism, each with a lower dose than if used alone, thus providing better analgesia with less side effects.

# Preventative Analgesia

## **PREVENT ACUTE PAIN**

- Initiation
- Amplification
- Transmission

**PREVENT ACUTE→CHRONIC PAIN**

# Preventative Analgesia

## PERIOPERATIVE ANALGESIA

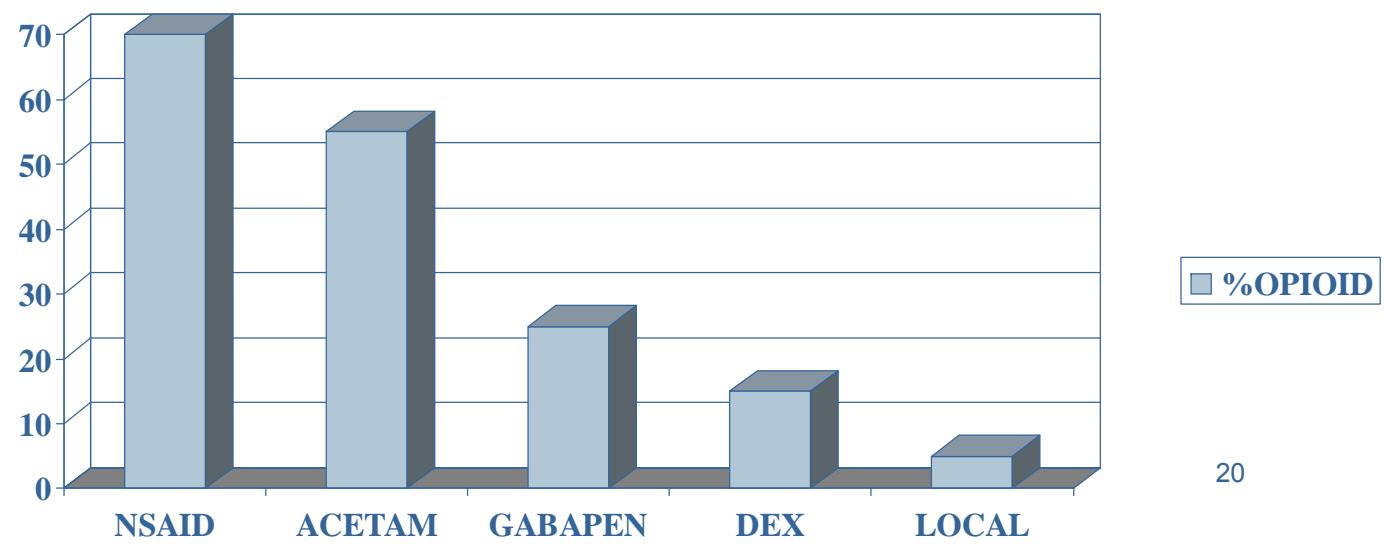
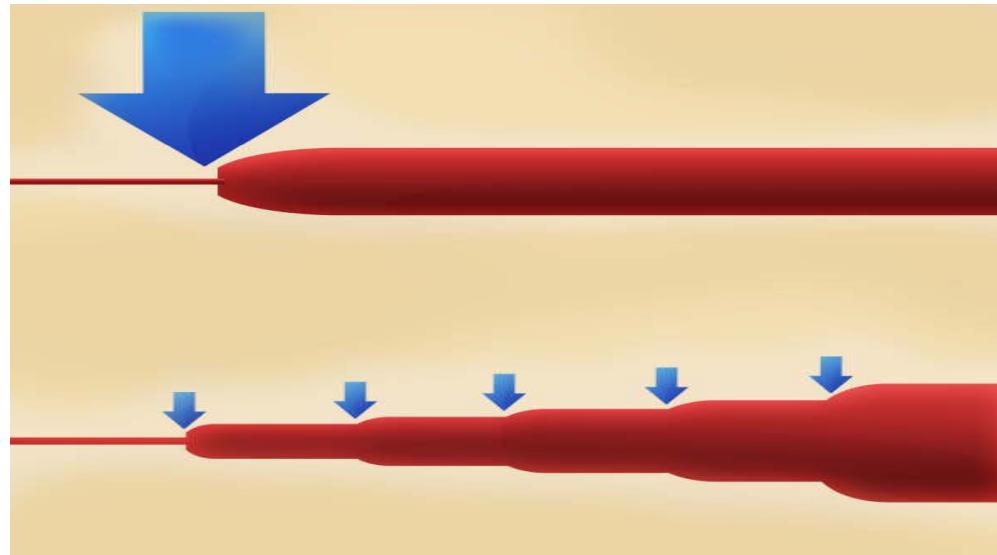
- Home
- Preoperative
- Intraoperative
- Postoperative
- Post discharge

# Preventative Analgesia

# MULTIMODAL ANALGESIA

- Acetaminophen REMY BR J ANAES 2005
  - NSAIDs /coxib REUBEN J BONE JT SURG 2005
  - GBP/pregabalin TIIPPANA ANESTH ANALG 2007
  - Ketamine HIMMELSEHER ANESTHESIOLOGY 2005
  - Dexamethasone SALERNO J BONE JT SURG 2006
  - Local Anesthetic LIU J AM COL SURG 2006
  - Controlled release opioid -oxycontin

# Multimodal Analgesia



# Acetaminophen

- Reduce opioid consumption 20-30%
- Reduce opioid related side effects
- Additive effect with NSAIDs
- No adverse effects

# NSAID / COXIB

- Reduce opioid consumption 30-50%
- Reduce opioid related side effects
- Improve pain scores up to 2/10
- CELECOXIB: among the most potent analgesics, no platelet dysfunction, no effect on bone fusion, no increase in thrombosis rates

# Celecoxib

The Journal of Arthroplasty Vol. 20 No. 7 Suppl. 3 2005

## **Celecoxib Does Not Affect Osteointegration of Cementless Total Hip Stems**

David R. Lionberger, MD, and Philip C. Noble, PhD

## **Effects of Celecoxib, a Novel Cyclooxygenase-2 Inhibitor, on Platelet Function in Healthy Adults: A Randomized, Controlled Trial**

*Philip T. Leese, MD, Richard C. Hubbard, MD, Aziz Karim, PhD, FCP,  
Peter C. Isakson, PhD, Shawn S. Yu, PhD, and G. Steven Geis, PhD, MD*

**No effect on bone growth or bleeding**

# Celecoxib

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## **Reports of Original Investigations**

### **Effect of short-term postoperative celecoxib administration on patient outcome after outpatient laparoscopic surgery**

*[Effet de l'administration postopératoire à court terme de célecoxib sur l'évolution des patients après une chirurgie par laparoscopie sans hospitalisation]*

Paul F. White PhD MD FANZCA, Ozlem Sacan MD, Burcu Tufanogullari MD, Matthew Eng, Nina Nuangchamnong, Babatunde Ogunnaike MD

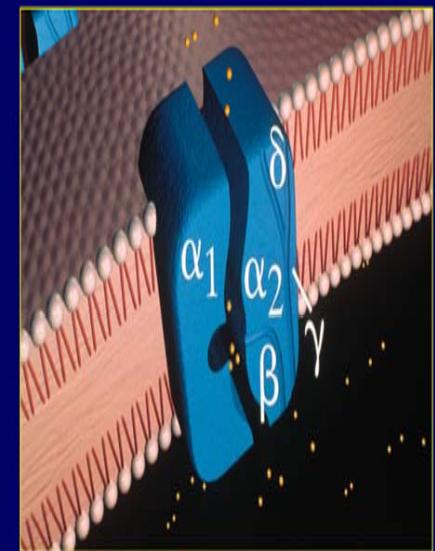
#### **Celecoxib x 3 days postop:**

- ↓ opioid use, pain, ileus
- ↑ satisfaction, recovery
- Resumed ADL 2 days earlier

# Gabapentinoids

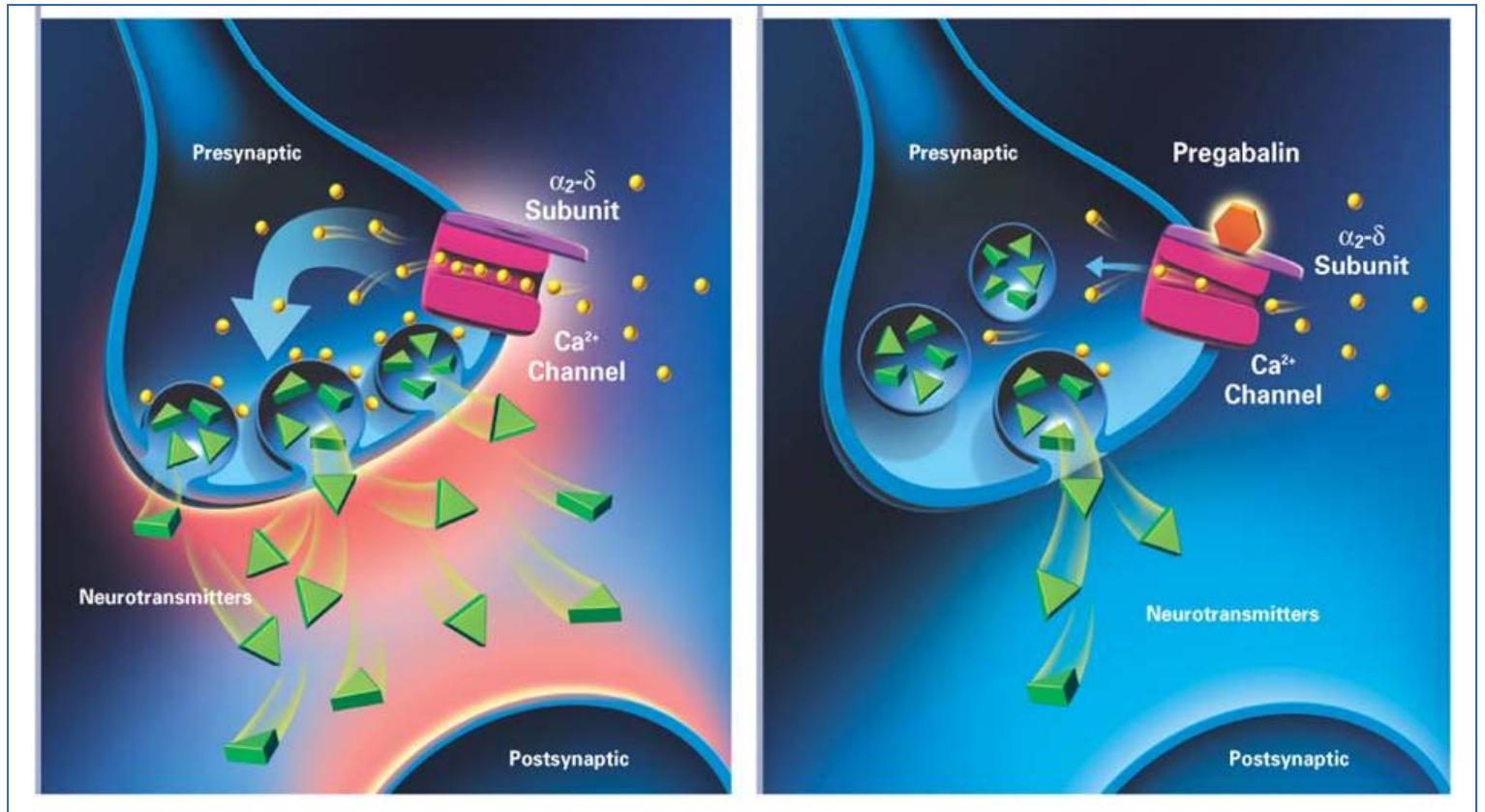
- Gabapentin
- Pregabalin
- 24 RCTs acute pain
- 600 mg optimal initial dose for gabapentin

The Calcium Channel



Adapted from Schwartz.<sup>1</sup>

# Gabapentinoids



PANDEY J NEUROSURG ANESTHESIOL 2005 TIIPPANA ANESTH ANALG 2007

# Gabapentin

## **Do Surgical Patients Benefit from Perioperative Gabapentin/Pregabalin? A Systematic Review of Efficacy and Safety**

Elina M. Tiippana, MD\*

Katri Hamunen, MD, PhD\*

Vesa K. Kontinen, MD, PhD\*#

Eija Kalso, MD, PhD\*#

**BACKGROUND:** Gabapentin and pregabalin have antialloodynic and antihyperalgesic properties useful for treating neuropathic pain. These properties may also be beneficial in acute postoperative pain. In this study we evaluated randomized, controlled trials examining the analgesic efficacy, adverse effects, and clinical value of gabapentinoids in postoperative pain.

**METHODS:** A systematic search of Medline, PubMed, and Cochrane Central Register of Controlled Trials (CENTRAL) databases yielded 22 randomized, controlled trials on perioperative administration of gabapentinoids for postoperative pain relief.

# Gabapentin

- Reduce opioid consumption 16-67%
- Reduce opioid related side effects
- Additive effect with NSAIDs
- Anxiolysis
- Improve functional recovery
- No increase adverse effects
- Pregabalin 6x more potent

# Surgical Outcomes

## **Preoperative Gabapentin Decreases Anxiety and Improves Early Functional Recovery from Knee Surgery**

Christophe Ménigaux, MD\*, Frédéric Adam, MD\*, Bruno Guignard, MD\*, Daniel I. Sessler, MD†, and Marcel Chauvin, MD\*

**Table 2.** Knee Flexion During Physiotherapy

	Postoperative Day 1			Postoperative Day 2		
	Control	Gabapentin	P value	Control	Gabapentin	P value
First passive flexion (°)	50 ± 17	70 ± 14	0.001	65 ± 15	75 ± 14	0.015
First active flexion (°)	44 ± 19	65 ± 14	0.001	62 ± 16	71 ± 17	0.034
Maximal passive flexion (°)	68 ± 15	78 ± 10	0.022	81 ± 7	85 ± 11	0.030
Maximal active flexion (°)	64 ± 18	76 ± 9	0.020	77 ± 11	84 ± 12	0.007

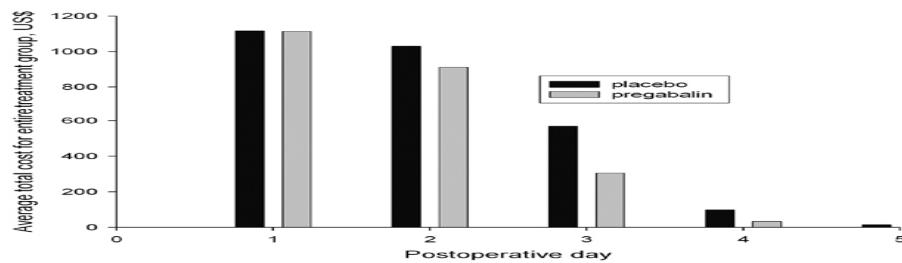
- ↓ anxiety >50%
- ↓ morphine use 58%
- ↑ active & passive flexion POD 1&2

# Surgical Outcomes

## Perioperative Pregabalin Improves Postoperative Outcomes after Total Knee Arthroplasty

Reduced Costs and Time to Discharge Following Total Knee Arthroplasty with Perioperative Pregabalin

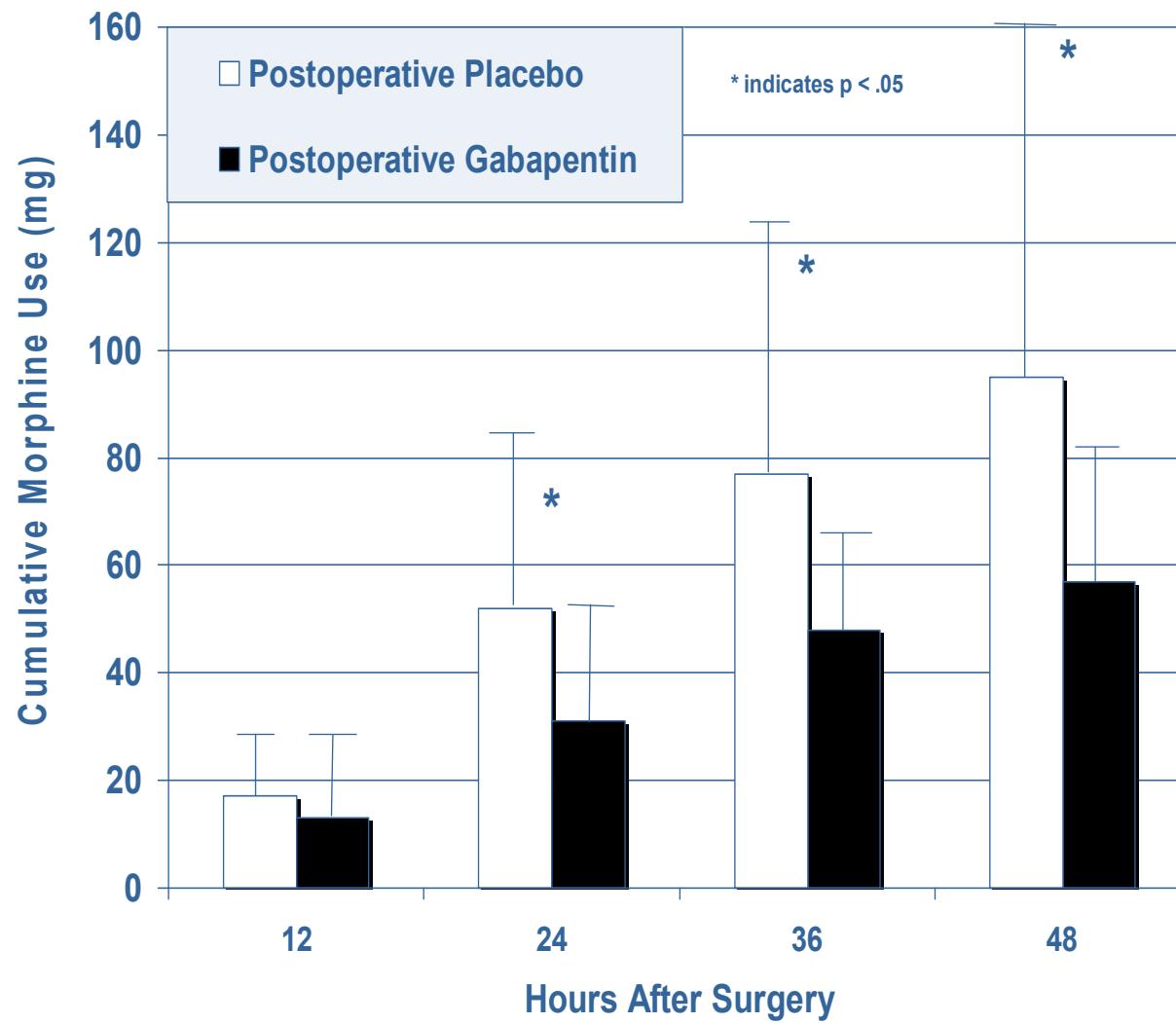
Asokumar Buvanendran, M.D., Scott S. Reuben, M.D., Maruti Kari, M.D., Jeffrey S. Kroin, Ph.D., Mario Moric, Ph.D.  
Anesthesiology, Rush Medical College, Chicago, Illinois



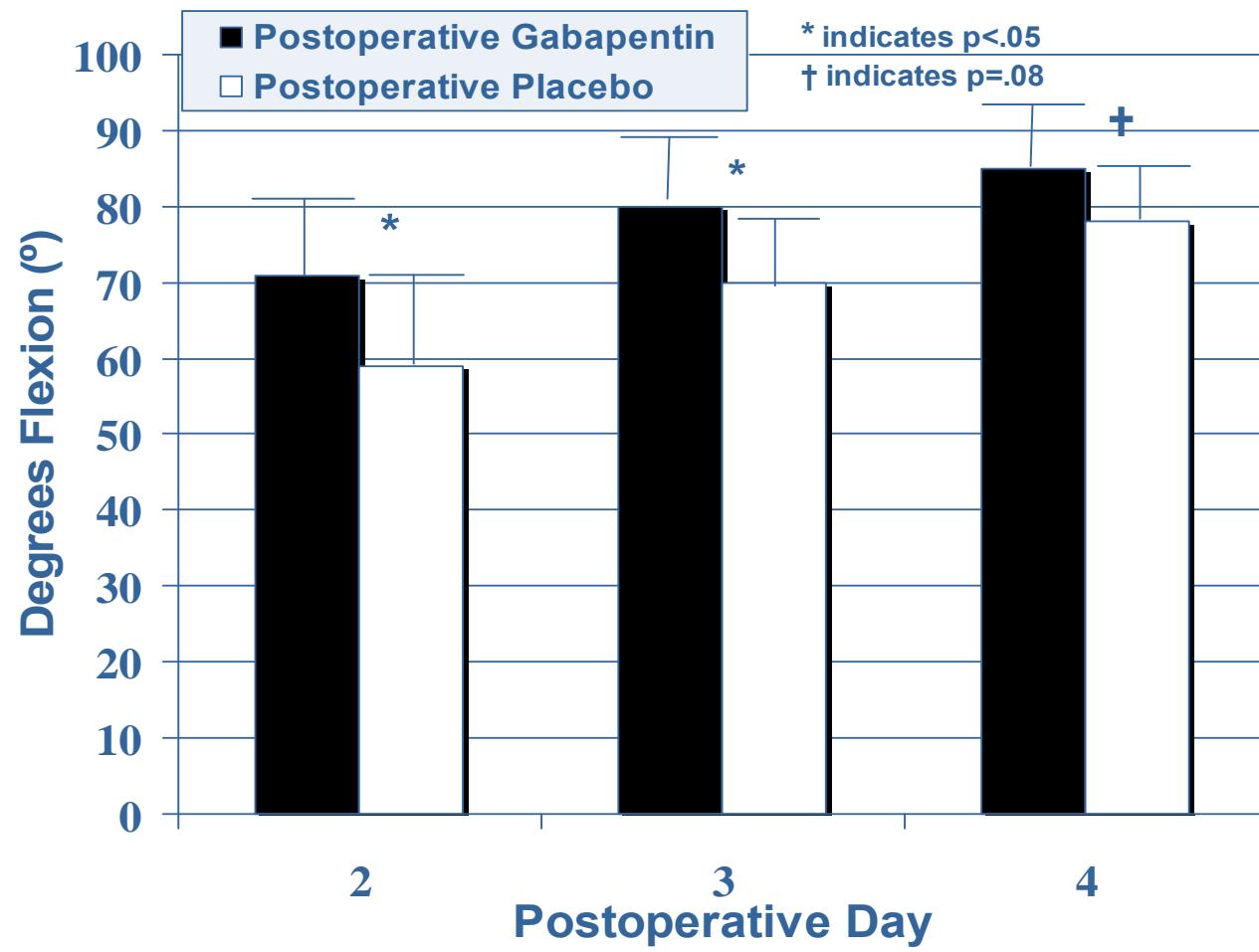
- flexion, sleep, ↓ PCEA, ↓nausea
- ↓ time to discharge, ↓ cost

ASA abstracts A392,393 2008

# Gabapentin TKA



# Gabapentin TKA



# Dexamethasone

- Reduce opioid consumption
- Reduce pain scores
- Reduce nausea & vomiting

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## CURRENT CONCEPTS REVIEW EFFICACY AND SAFETY OF STEROID USE FOR POSTOPERATIVE PAIN RELIEF

UPDATE AND REVIEW OF THE MEDICAL LITERATURE

BY ANGELO SALERNO, DIP APP Sc, GRAD DIP, MPOD (POD SURG), AND ROBERT HERMANN, DPS

BISGAARD ANN SURG 2003 HOLTE J AM COLL SURG 2002 KARDASH ANESTH 33  
ANALG 2008

# Local Anesthetic

- Improves dynamic pain
  - Improves rehabilitation
- 
- Infiltration
  - Nerve block
  - Neuraxial block

KLEIN ANESTH ANALG 2005 GERGES J CLIN ANESTH 2006  
NODA MASUI 1990 WANG REG ANESTH PAIN MED 2002 ILFIELD ANESTH ANALG 2006  
CAPDEVILA ANESTH 1999 CHELLY J ARTHOPLSTY 2001 SINGELYN ANESTH ANALG  
1998

# Thoracic Epidural

Anesthesiology 2002; 97:540-9

© 2002 American Society of Anesthesiologists, Inc. Lippincott Williams & Wilkins, Inc.

## ***Epidural Analgesia Enhances Functional Exercise Capacity and Health-related Quality of Life after Colonic Surgery***

### *Results of a Randomized Trial*

Franco Carli, M.D., M.Phil.,\* Nancy Mayo, Ph.D.,† Kristine Klubien, M.D.,‡ Thomas Schriker, M.D., Ph.D.,‡  
Judith Trudel, M.D., M.Sc.,§ Paul Belliveau, M.D.§

- Hastens return of bowel function for colon surgery
- Improves rehabilitation &quality of life

CARLI ANESTHESIOL 2002 97:540

BASSE ANN SURG 2000 232:51

# Controlled-release Opioid

- Oxycontin, Hydromorphone, MS contin
- Oxycontin : Biphasic release
- Constant low blood levels
- 12 h duration
- Improved rehabilitation

# Ketamine / Clonidine / Lidocaine/ Tramadol

- Decrease pain, opioid consumption
- High side effect profile-clonidine
- Difficult administration (ketamine, lidocaine)-must do iv /infusion
- Tramadol well tolerated by elderly

BEAULIEU CAN J ANESTH 2007 McCARTNEY ANESTH ANALG 2004 LAUWICK CJA 2008  
LIKAR CLIN THER 2006 ROSENTHAL J AM GERIATR SOC 2004

# Multimodal Analgesia

## EFFECTS ARE ADDITIVE

- Acetaminophen      FLETCHER      CAN J ANESTH 1997
- NSAIDs /coxib      GILRON PAIN 2005      TURAN ANESTH ANALG 2006
- Gabapentin / pregabalin      REUBEN ANESTH ANALG 2006      KOC ANESTH ANALG 2007
- Dexamethasone      KJETIL ANESTH ANALG 2007
- Local Anesthetic      FASSOUAKI ANESTH ANALG 2005

# Multimodal Analgesia

## REDUCES PCEA USE

- Celecoxib - TKA

BUVANENDRAN JAMA 2003

↓ PCEA use 60%

- Gabapentin - LES

TURAN BR J ANAESTH 2006

↓ PCEA use 60%

# Clinical Pathways

## PREOPERATIVE

- celecoxib, acetaminophen, gabapentin, dexamethasone,  
± oxycontin

## INTRAOPERATIVE

- local infiltration, nerve block, neuraxial block,  
+/- ketamine

# Clinical Pathways

## POSTOPERATIVE

- celecoxib, acetaminophen, gabapentin, oxycontin, continuous nerve block, wound infiltration, epidural

## HOME

- Continue oral multimodal analgesia at home days to 2 weeks

# Clinical example I

## L4/5 Laminectomy

### PREOP (2 hr)

- celecoxib 400 mg
- acetaminophen 1000mg
- gabapentin 600 mg
- oxycontin 20 mg

# Clinical example I

## INTRAOP

- ketamine 20 mg iv
- dexamethasone 8 mg iv
- Local infiltration 20 ml 0.5% bupivacaine by surgeon

# Clinical example I

## POSTOP

- celecoxib 200 mg q12h
- acetaminophen 1000mg q6h
- gabapentin 100-200 mg q8h
- oxycontin 10 mg q8h
- oxycodone 5-10 mg prn b/t

# Clinical example II

THA

PREOP (2 hr)

- celecoxib 400 mg
- acetaminophen 1000mg
- gabapentin 600 mg
- Spinal: 15mg 0.5%bupivacaine  
10ug fentanyl + 100ug epimorph

# Clinical example II

## INTRAOP

- ketamine 20 mg iv
- dexamethasone 8 mg iv
- extensive local infiltration 0.2% ropivacaine by surgeon

# Clinical example II

## POSTOP

- celecoxib 200 mg q12h
- acetaminophen 1000mg q6h
- gabapentin 100-200 mg q8h
- oxycontin 10-20 mg in PACU & 5-10 mg q8h
- oxycodone 5-10 mg prn b/t

# Clinical example III

**TKA**

**PREOP (2 hr)**

- celecoxib 400 mg
- acetaminophen 1000mg
- gabapentin 600 mg

CFNB/FNB 20 ml 0.5% ropivacaine  
SNB            20 ml 0.5% ropivacaine

# Clinical example III

## PREOP

- spinal 10 mg 0.5% bupivacaine  
10 µg fentanyl

## POSTOP

- celecoxib 200 mg q12h
- acetaminophen 1000mg q6h
- gabapentin 100-200 mg q8h
- oxycontin 10-20 mg in PACU & 5-10 mg q8h
- PCA or oxycodone 5-10 mg prn b/t

# Clinical example III

## PREOP

- spinal 10 mg 0.5% bupivacaine  
10 µg fentanyl

## INTRAOP

- ketamine 20 mg iv
- dexamethasone 8 mg iv

# Clinical example IV

## Lap-Chole

### PREOP (2 hr)

- celecoxib 400 mg
- acetaminophen 1000 mg
- gabapentin 600 mg
- oxycontin 10 mg

# Clinical example IV

## INTRAOP

- ketamine 20mg iv
- dexamethasone 8 mg iv
- Infiltration 0.5% bupivacaine  
umbilical port: skin to  
preperitoneal
- Infiltration port sites 0.5%  
bupivacaine

# Clinical example IV

## INTRAOP

- 2I RL

# SB order sheet



## PHYSICIAN'S ORDERS

### PHYSICIAN'S ORDERS

All orders shall be DATED, TIMED, and SIGNED  
All medication orders shall be written in the GENERIC or non-proprietary name.  
All orders shall be written legibly using ball point pen.

### PATIENT IDENTIFICATION

TIME & DATE				SIGNATURE OF NURSE WHEN DRUG ADMINISTERED
YYYY/MM/DD				
<b>HOLLAND ORTHOPAEDIC &amp; ARTHRITIC CENTRE ACUTE PAIN SERVICE (APS) PRE-OPERATIVE MEDICATION ORDER FORM</b>				
	Yes	No		Doctor must check off appropriate orders.
	<input checked="" type="checkbox"/>			Give the following medication(s) with a sip of water 1 - 2 hours pre-operatively
			1	acetaminophen _____ mg po
			2	celecoxib _____ mg po (hold if creatinine >106umol/L).
			3	gabapentin _____ mg po
			4	
			5	
			6	
Doctor's Signature:			PRINT NAME:	

# Summary

- Use preventative, perioperative, multimodal analgesia in every case
- This will reduce opioid use, improve outcome and reduce chronic pain