



ERAS in 2018

How to reduce perioperative opioids in an ERAS population

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- Post-op nausea, vomiting
- Respiratory depression
- Induced opioid hyperalgesia
- Constipation
- Delirium

Delayed postop recovery Reduced patient satisfaction

- Post operative pain
- Hypertension, tachycardia
- Heart ischemia
- Increased morbidity
- Delayed postop recovery and Reduced patient satisfaction

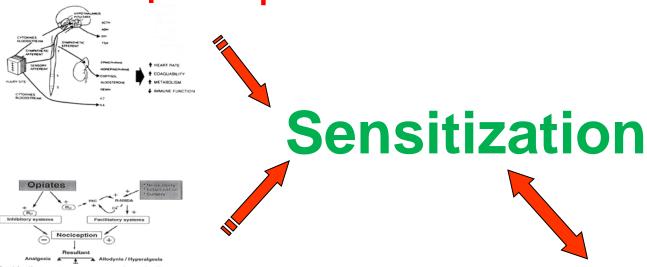






- Lack of good analgesia (block or multimodal)

A- Nociceptive inputs



B- Opioids

« Persistent Post-Surgical Pain »

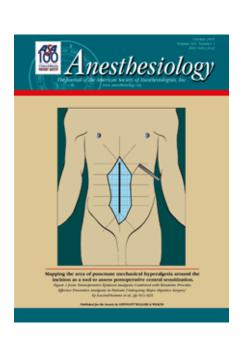


- High intraoperative opioid doses
- Lack of anti-O.I.H. strategy

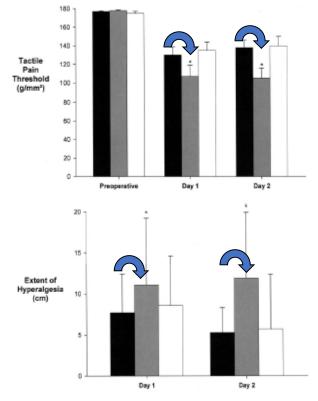
Remifentanil-induced Postoperative Hyperalgesia and Its Prevention with Small-dose Ketamine

Vincent Joly, M.D.,* Philippe Richebe, M.D.,† Bruno Guignard, M.D.,* Dominique Fletcher, M.D.,‡ Pierre Maurette, M.D.,§ Daniel I. Sessler, M.D., Marcel Chauvin, M.D.#

| | Small-dose Remifentanil (n = 25) | Large-dose Remifentanil (n – 25) | Large-dose Remifentanil- Ketamine (n – 24) |
|--|--|--|---|
| Remifentanil dose, mg | 0.9 ± 0.3* | 6.7 ± 3.1 | 6.5 ± 3.4 |
| Desflurane, MAC/h Ephedrine, No. of doses/No. of patients | 0.8 ± 0.2* 9/17 | 0.5 ± 0.2 10/13 | 0.6 ± 0.2 50/15† |
| Final intraoperative temperature, °C | 36.6 ± 0.7 | 36.3 ± 0.8 | 36.3 ± 0.9 |
| Awakening time, min | 14 ± 6 | 13 ± 5 | 14 ± 6 |
| Extubation time, | 16 ± 6 | 14 ± 6 | 15 ± 4 |
| Time to first postoperative morphine, min | 35 (28-46) | 24 (20–33) | 41 (32–52) |
| Morphine given in PACU, mg | 16 (10–24) | 20 (17–27) | 20 (14–23) |
| 0–48 h cumulative postoperative morphine consumption, mg | 68 (50–91) | 86 (59–109) ‡ | 62 (48–87) |
| Postoperative nausea and vomiting, No. of patients | 7 | 8 | 8 |
| Droperidol, No. of doses/No. of patients | 8/7 | 8/8 | 8/8 |



Nociceptive threshold is lowered



Area of Hyperalgesia is bigger

Opioid-induced hyperalgesia in patients after surgery: a systematic review and a meta-analysis

D. Fletcher^{1,2,3*} and V. Martinez^{1,2,3}

British Journal of Anaesthesia 112 (6): 991–1004 (2014)

| Study (first author, year) | Number of patients in control or low opioid dose group | Number of patients in high opioid dose group | Patients/surgery | Intervention | Outcomes |
|----------------------------------|---|---|------------------------------------|--|--|
| Agata ⁵³ (2010) | 15 low dose | 15 | Elective orthognatic surgery | I.V. remifentanil (0.15 μ g kg min ⁻¹) vs (> 0.3 μ g kg min ⁻¹) | Pain VAS at rest at 1, 3, 6, 12 and 24 h. PCA i.v. fentanyl 24 h. Haemodynamic variables 12 h. PONV and shivering 24 h |
| Carvalho ⁴⁴ (2012) | 9 control 9 low dose | 9 | Caesarean section | Intrathecal single shot fentanyl (5 μg) vs (25 μg) | Pain VAS at rest, oxygen saturation and respiratory rate 30 min, 1, 4, 8, 12 and 24 h. Intraoperative pain, nausea, hypotension, and vasopressor use. PCA i.v. morphine 24 h |
| Chia ⁷ (1999) | 30 low dose | 30 | Hysterectomy | $1 \mu g kg^{-1}$ fentanyl bolus vs 15 $\mu g kg^{-1}$ bolus plus 100 $\mu g h^{-1}$ infusion | Pain VAS at rest 4, 8, 12, and 16 h. Haemodynamic, arterial blood gas, and sedation scores. PCA i.v. morphine 24 h |
| Cho ⁴⁰ (2008) | 30 control 30 low dose | 30 | Gynaecology | I.V. remifentanil (target 1 ng ml ⁻¹) vs high-dose remifentanil (target 3 ng ml ⁻¹) | Pain VAS at rest 15,30,45,60 min and 6,12,24, and 48 h. Sedation, agitation. PCA i.v. morphine 48 h. PONV requiring antiemetic |
| Cooper ⁶ (1997) | 30 control | 30 | Caesarean section | Intrathecal single shot fentanyl (25 µg) vs placebo | Intraoperative most severe pain; intraoperative nausea, vomiting, drowsiness. Pain VAS at rest and during coughing at 15 min, 3, 6, 10, and 23 h. PON, POV, pruritus, drowsiness. PCA i.v. morphine 24 h |
| Cooper ⁴⁵ (2002) | 18 control | 18 | Caesarean section | Intrathecal single shot fentanyl (25 μg) vs placebo | Pain VAS at rest and during coughing in PACU and then at 2, 4, 10, and 20 h. Intraoperative pain; PON, POV, pruritus, drowsiness. PCA epidural fentanyl |
| Cortinez ¹⁰ (2001) | 30 control | 30 | Gynaecology | I.V. remifentanil (0.23 μ g kg min $^{-1}$) vs placebo | Pain VAS during coughing at 15, 30,45, 90 min, 2, and 24 h. PCA i.v. morphine 24 h, PONV, sedation, hypoxemia (pulse oximeter), respiratory depression; patient satisfaction |
| Fechner ²¹ (2013) | 18 low dose | 16 | Coronary artery bypass graft | I.V. sufentanil (target 0.4 ng ml ⁻¹) vs remifentanil (target 0.8 ng ml ⁻¹) | Pain NRS at rest and during deep inspiration, PCA i.v. morphine 48 h. Cognitive function, sedation, constipation, PONV. Primary and secondary hyperalgesia |
| Guignard ⁹ (2000) | 25 low dose | 24 | Colorectal surgery | I.V. remifentanil (0.1 μ g kg min ⁻¹) vs (0.3 μ g kg min ⁻¹) | Pain VAS at rest at 24 h. PCA i.v. morphine 48 h. PON, POV, pruritus, dysphoria, diplopia, hallu anations |
| Hansen ⁴³ (2005) | 18 control | 21 | Major abdominal surgery | I.V. remifentanil (0.4 µg kg min ⁻¹) vs placebo | Summed pain VAS at rest and during coughing at 4, 6, and 24 h. PCA i.v. morphine 24 h. PON, POV, sedation |
| Joly ¹¹ (2005) | 25 low dose | 25 | Major abdominal surgery | I.V. remifentanil (0.05 μ g kg min ⁻¹) vs (0.4 μ g kg min ⁻¹) | Pain verbal scale for 3 h then pain VAS at rest every 4 h for 44 h. Pain VAS when peak flow measurement at 24 and 48 h. PCA i.v. |
| Continue | d | | | | morphine 48 h. PONV, laryngospasm, bronchospasm, respiratory depression, muscular rigidity, agitation, and shivering Primary and secondary hyperalgesia |

27 studies over 15 years

About 1500 patients

All types of opioids

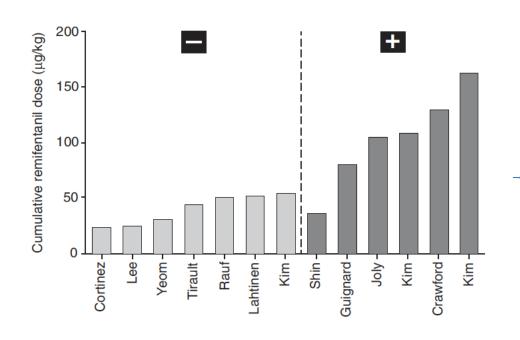
Mostly remifentanil, though

All surgeries

Intraoperative Use of Remifentanil for TIVA: Postoperative Pain, Acute Tolerance, and Opioid-Induced Hyperalgesia

Martin S. Angst, MD

Journal of Cardiothoracic and Vascular Anesthesia, Vol 29, No S1 (June), 2015: pp S16-S22



Remifentanil studies:

- light gray bars = « small doses » of remifentanil = negative study regarding OIH (23-54 mcg/kg)
- dark gray bars = « high doses of remifentanil » = positive studies regarding OIH (35-162 mcg/kg)

Chronic Pain as an Outcome of Surgery

Anesthesiology 2000; 93:1123-33

A Review of Predictive Factors

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

Persistent postsurgical pain: risk factors and prevention

Henrik Kehlet, Troels S Jensen, Clifford J Woolf

Lancet 2006; 367: 1618-25

| | Estimated incidence of chronic pain | Estimated chronic severe (disabling) pain (>5 out of score of 10) | US surgical volumes (1000s)† |
|---|-------------------------------------|---|---------------------------------|
| Amputation ² | 30–50% | 5–10% | 159 (lower limb only) |
| Breast surgery (lumpectomy and mastectomy) ³ | 20-30% | 5-10% | 479 |
| Thoracotomy ⁴⁻⁷ | 30-40% | 10% | Unknown |
| Inguinal hernia repair ⁸⁻¹⁰ | 10% | 2–4% | 609 |
| Coronary artery bypass surgery ¹¹⁻¹³ | 30-50% | 5–10% | 598 |
| Caesarean section ¹⁴ | 10% | 4% | 220 |

Narcotic Sparing Analgesia in ERAS *Before surgery*

- Proper patient evaluation: chronic pain? Chronic opioid exposure...
- Patient Education prior surgery
- Set proper expectations for your patient
- Start multimodal analgesia before entering the OR
- Place preoperative epidural WHEN necessary prior to OR entrance if possible and test it

Narcotic Sparing Analgesia in ERAS During surgery



- Epidural versus Regional: Epidural is no more recommended for colorectal laparoscopic surgery!
- Prefer TAP block or RS blocks if laparoscopic surgery or preperitoneal infiltration
- IV lidocaine is also an option when epidural is not needed
- NSAIDs have to be discussed with the surgical team and based on patient's evaluation
- Adjuvant analgesics: NMDA modulators (Ketamine, N2O, dextromethorphan, magnesium might be added),
 Alpha-2 agonists might be used
- Gabapentinoids are not recommended in this type of surgery



• Intraoperative nociception MONITORS

Narcotic Sparing Analgesia in ERAS After surgery

- Proper orders must be implemented for each chosen strategy: PCEA, Regional Blocks (TAP, RS, CWI), PCA, multimodal analgesia permitted...
- Multimodal must be continued
- If epidural placement, a STOP-test must be proposed at POD2 to avoid delaying the patient's discharge
- Ketamine, IV lidocaine might be continued postoperatively when needed

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