Practice Guidelines for Acute Pain Management in the Perioperative Setting

An Updated Report by the American Society of Anesthesiologists Task Force on Acute Pain Management

PRACTICE Guidelines are systematically developed recommendations that assist the practitioner and patient in making decisions about health care. These recommendations may be adopted, modified, or rejected according to clinical needs and constraints and are not intended to replace local institutional policies. In addition, Practice Guidelines developed by the American Society of Anesthesiologists (ASA) are not intended as standards or absolute requirements, and their use cannot guarantee any specific outcome. Practice Guidelines are subject to revision as warranted by the evolution of medical knowledge, technology, and practice. They provide basic recommendations that are supported by a synthesis and analysis of

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- What other guideline statements are available on this topic?
 - These Practice Guidelines update the "Practice Guidelines for Acute Pain Management in the Perioperative Setting," adopted by the ASA in 2003 and published in 2004.*
- Why was this guideline developed?
 - In October 2010, the Committee on Standards and Practice Parameters elected to collect new evidence to determine whether recommendations in the existing Practice Guideline were supported by current evidence.
- How does this statement differ from existing guidelines?
 - New evidence presented includes an updated evaluation of scientific literature and findings from surveys of experts and randomly selected ASA members. The new findings did not necessitate a change in recommendations.
- Why does this statement differ from existing guidelines?
- The ASA guidelines differ from the existing guidelines because they provide new evidence obtained from recent scientific literature as well as findings from new surveys of expert consultants and randomly selected ASA members.

the current literature, expert and practitioner opinion, open forum commentary, and clinical feasibility data.

This document updates the "Practice Guidelines for Acute Pain Management in the Perioperative Setting: An Updated Report by the American Society of Anesthesiologists Task Force on Acute Pain Management," adopted by the ASA in 2003 and published in 2004.*

Methodology

A. Definition of Acute Pain Management in the Perioperative Setting

For these Guidelines, acute pain is defined as pain that is present in a surgical patient after a procedure. Such pain may be the result of trauma from the procedure or procedurerelated complications. Pain management in the perioperative setting refers to actions before, during, and after a procedure

Supplemental digital content is available for this article. Direct URL citations appear in the printed text and are available in both the HTML and PDF versions of this article. Links to the digital files are provided in the HTML text of this article on the Journal's Web site (www.anesthesiology.org). that are intended to reduce or eliminate postoperative pain before discharge.

B. Purpose of the Guidelines

The purpose of these Guidelines is to (1) facilitate the safety and effectiveness of acute pain management in the perioperative setting; (2) reduce the risk of adverse outcomes; (3) maintain the patient's functional abilities, as well as physical and psychologic well-being; and (4) enhance the quality of life for patients with acute pain during the perioperative period. Adverse outcomes that may result from the undertreatment of perioperative pain include (but are not limited to) thromboembolic and pulmonary complications, additional time spent in an intensive care unit or hospital, hospital readmission for further pain management, needless suffering, impairment of health-related quality of life, and development of chronic pain. Adverse outcomes associated with the management of perioperative pain include (but are not limited to) respiratory depression, brain or other neurologic injury, sedation, circulatory depression, nausea, vomiting, pruritus, urinary retention, impairment of bowel function, and sleep disruption. Health-related quality of life includes (but is not limited to) physical, emotional, social, and spiritual well-being.

C. Focus

These Guidelines focus on acute pain management in the perioperative setting for adult (including geriatric) and pediatric patients undergoing either inpatient or outpatient surgery. Modalities for perioperative pain management addressed in these Guidelines require a higher level of professional expertise and organizational structure than "as needed" intramuscular or intravenous injections of opioid analgesics. These Guidelines are not intended as an exhaustive compendium of specific techniques.

Patients with severe or concurrent medical illness such as sickle cell crisis, pancreatitis, or acute pain related to cancer or cancer treatment may also benefit from aggressive pain control. Labor pain is another condition of interest to anesthesiologists. However, the complex interactions of concurrent medical therapies and physiologic alterations make it impractical to address pain management for these populations within the context of this document.

Although patients undergoing painful procedures may benefit from the appropriate use of anxiolytics and sedatives in combination with analgesics and local anesthetics when indicated, these Guidelines do not specifically address the use of anxiolysis or sedation during such procedures.

D. Application

These Guidelines are intended for use by anesthesiologists and individuals who deliver care under the supervision of anesthesiologists. The Guidelines may also serve as a resource

† International Anesthesia Research Society, 68th Clinical and Scientific Congress, Orlando, Florida, March 6, 1994.

for other physicians and healthcare professionals who manage perioperative pain. In addition, these Guidelines may be used by policymakers to promote effective and patient-centered care.

Anesthesiologists bring an exceptional level of interest and expertise to the area of perioperative pain management. Anesthesiologists are uniquely qualified and positioned to provide leadership in integrating pain management within perioperative care. In this leadership role, anesthesiologists improve quality of care by developing and directing institution-wide, interdisciplinary perioperative analgesia programs.

E. Task Force Members and Consultants

The original Guidelines were developed by an ASA appointed task force of 11 members, consisting of anesthesiologists in private and academic practices from various geographic areas of the United States, and two consulting methodologists from the ASA Committee on Standards and Practice Parameters.

The Task Force updated the Guidelines by means of a seven-step process. First, they reached consensus on the criteria for evidence. Second, original published research studies from peer-reviewed journals relevant to acute pain management were reviewed and evaluated. Third, expert consultants were asked to: (1) participate in opinion surveys on the effectiveness of various acute pain management recommendations and (2) review and comment on a draft of the updated Guidelines. Fourth, opinions about the updated Guideline recommendations were solicited from a sample of active members of the ASA. Fifth, opinion-based information obtained during an open forum for the original Guidelines, held at a major national meeting,† was reexamined. Sixth, the consultants were surveyed to assess their opinions on the feasibility of implementing the updated Guidelines. Seventh, all available information was used to build consensus to finalize the updated Guidelines. A summary of recommendations may be found in appendix 1.

F. Availability and Strength of Evidence

Preparation of these Guidelines followed a rigorous methodological process. Evidence was obtained from two principal sources: scientific evidence and opinion-based evidence.

Scientific Evidence

Study findings from published scientific literature were aggregated and are reported in summary form by evidence category, as described below. All literature (e.g., randomized controlled trials [RCTs], observational studies, case reports) relevant to each topic was considered when evaluating the findings. However, for reporting purposes in this document, only the highest level of evidence (i.e., level 1, 2, or 3 within

category A, B, or C, as identified below) is included in the summary.

Category A: Supportive Literature

Randomized controlled trials report statistically significant (P < 0.01) differences between clinical interventions for a specified clinical outcome.

- Level 1: The literature contains multiple RCTs, and aggregated findings are supported by meta-analysis.‡
- Level 2: The literature contains multiple RCTs, but the number of studies is insufficient to conduct a viable meta-analysis for the purpose of these Guidelines.
- Level 3: The literature contains a single randomized con-

Category B: Suggestive Literature

Information from observational studies permits inference of beneficial or harmful relationships among clinical interventions and clinical outcomes.

- Level 1: The literature contains observational comparisons (e.g., cohort, case-control research designs) of clinical interventions or conditions and indicates statistically significant differences between clinical interventions for a specified clinical outcome.
- Level 2: The literature contains noncomparative observational studies with associative (*e.g.*, relative risk, correlation) or descriptive statistics.
- Level 3: The literature contains case reports.

Category C: Equivocal Literature

The literature cannot determine whether there are beneficial or harmful relationships among clinical interventions and clinical outcomes.

- Level 1: Meta-analysis did not find significant differences (P > 0.01) among groups or conditions.
- Level 2: The number of studies is insufficient to conduct meta-analysis, and (1) RCTs have not found significant differences among groups or conditions or (2) RCTs report inconsistent findings.
- Level 3: Observational studies report inconsistent findings or do *not* permit inference of beneficial or harmful relationships.

Category D: Insufficient Evidence from Literature

The *lack* of scientific evidence in the literature is described by the following terms.

Inadequate: The available literature cannot be used to assess relationships among clinical interventions and clinical outcomes. The literature either does not meet the criteria for content as defined in the "Focus" of the Guidelines or does not permit a clear interpretation of findings due to methodological concerns (e.g., confounding in study design or implementation).

Silent: No identified studies address the specified relationships among interventions and outcomes.

Opinion-based Evidence

All opinion-based evidence (*e.g.*, survey data, open-forum testimony, Internet-based comments, letters, editorials) relevant to each topic was considered in the development of these updated Guidelines. However, only the findings obtained from formal surveys are reported.

Opinion surveys were developed for this update by the Task Force to address each clinical intervention identified in the document. Identical surveys were distributed to expert consultants and ASA members.

Category A: Expert Opinion

Survey responses from Task Force-appointed expert consultants are reported in summary form in the text, with a complete listing of consultant survey responses reported in appendix 2.

Category B: Membership Opinion

Survey responses from active ASA members are reported in summary form in the text, with a complete listing of ASA member survey responses reported in appendix 2.

Opinion survey responses are recorded using a 5-point scale and summarized based on median values.§

Strongly Agree: Median score of 5 (At least 50% of the responses are 5)

Agree: Median score of 4 (At least 50% of the responses are 4 or 4 and 5)

Equivocal: Median score of 3 (At least 50% of the responses are 3, or no other response category or combination of similar categories contain at least 50% of the responses)

Disagree: Median score of 2 (At least 50% of responses are 2 or 1 and 2)

Strongly Disagree: Median score of 1 (At least 50% of responses are 1)

Category C: Informal Opinion

Open-forum testimony from the previous update, Internetbased comments, letters, and editorials are all informally evaluated and discussed during the development of Guideline recommendations. When warranted, the Task Force may add educational information or cautionary notes based on this information.

[‡] All meta-analyses are conducted by the American Society of Anesthesiologists methodology group. Meta-analyses from other sources are reviewed but not included as evidence in this document.

[§] When an equal number of categorically distinct responses are obtained, the median value is determined by calculating the arithmetic mean of the two middle values. Ties are calculated by a predetermined formula.

Guidelines

I. Institutional Policies and Procedures for Providing Perioperative Pain Management

Institutional policies and procedures include (but are not limited to) (1) education and training for healthcare providers, (2) monitoring of patient outcomes, (3) documentation of monitoring activities, (4) monitoring of outcomes at an institutional level, (5) 24-h availability of anesthesiologists providing perioperative pain management, and (6) use of a dedicated acute pain service.

Observational studies report that education and training programs for healthcare providers are associated with decreased pain levels, ¹⁻⁴ decreased nausea and vomiting, ² and improved patient satisfaction ¹ (*Category B2 evidence*), although the type of education and training provided varied across the studies. Published evidence is insufficient to evaluate the impact of monitoring patient outcomes at either the individual patient or institutional level, and the 24-h availability of anesthesiologists (*Category D evidence*). Observational studies assessing documentation activities suggest that pain outcomes are not fully documented in patient records (*Category B2 evidence*). ⁵⁻¹¹ Observational studies indicate that acute pain services are associated with reductions in perioperative pain (*Category B2 evidence*), ^{12–20} although treatment components of the acute pain services varied across the studies.

The consultants and ASA members strongly agree that anesthesiologists offering perioperative analgesia services should provide, in collaboration with other healthcare professionals as appropriate, ongoing education and training of hospital personnel regarding the effective and safe use of the available treatment options within the institution. The consultants and ASA members also strongly agree that anesthesiologists and other healthcare providers should use standardized, validated instruments to facilitate the regular evaluation and documentation of pain intensity, the effects of pain therapy, and side effects caused by the therapy. The ASA members agree and the consultants strongly agree that: (1) anesthesiologists responsible for perioperative analgesia should be available at all times to consult with ward nurses, surgeons, or other involved physicians, and should assist in evaluating patients who are experiencing problems with any aspect of perioperative pain relief; (2) anesthesiologists should provide analgesia services within the framework of an Acute Pain Service and participate in developing standardized institutional policies and procedures; and (3) an integrated approach to perioperative pain management (e.g., oradministering, and transitioning therapies, transferring responsibility for pain therapy, outcomes assessment, continuous quality improvement) should be used to minimize analgesic gaps.

Recommendations for Institutional Policies and Procedures. Anesthesiologists offering perioperative analgesia services should provide, in collaboration with other healthcare professionals as appropriate, ongoing education and training

to ensure that hospital personnel are knowledgeable and skilled with regard to the effective and safe use of the available treatment options within the institution. Educational content should range from basic bedside pain assessment to sophisticated pain management techniques (e.g., epidural analgesia, patient controlled analgesia, and various regional anesthesia techniques) and nonpharmacologic techniques (e.g., relaxation, imagery, hypnotic methods). For optimal pain management, ongoing education and training are essential for new personnel, to maintain skills, and whenever therapeutic approaches are modified.

Anesthesiologists and other healthcare providers should use standardized, validated instruments to facilitate the regular evaluation and documentation of pain intensity, the effects of pain therapy, and side effects caused by the therapy.

Analgesic techniques involve risk for adverse effects that may require prompt medical evaluation. Anesthesiologists responsible for perioperative analgesia should be available *at all times* to consult with ward nurses, surgeons, or other involved physicians, and should assist in evaluating patients who are experiencing problems with any aspect of perioperative pain relief.

Anesthesiologists providing perioperative analgesia services should do so within the framework of an Acute Pain Service and participate in developing standardized institutional policies and procedures. An integrated approach to perioperative pain management that minimizes analgesic gaps includes ordering, administering, and transitioning therapies, and transferring responsibility for perioperative pain therapy, as well as outcomes assessment and continuous quality improvement.

II. Preoperative Evaluation of the Patient

Preoperative patient evaluation and planning is integral to perioperative pain management. Proactive individualized planning is an anticipatory strategy for postoperative analgesia that integrates pain management into the perioperative care of patients. Patient factors to consider in formulating a plan include type of surgery, expected severity of postoperative pain, underlying medical conditions (e.g., presence of respiratory or cardiac disease, allergies), the risk—benefit ratio for the available techniques, and a patient's preferences or previous experience with pain.

Although the literature is insufficient regarding the efficacy of a preoperative directed pain history, a directed physical examination, or consultations with other healthcare providers (*Category D evidence*), the Task Force points out the obvious value of these activities. One observational study in a neonatal intensive care unit suggests that the implementation of a pain management protocol may be associated with reduced analgesic use, shorter time to extubation, and shorter times to discharge (*Category B2 evidence*).²¹

The ASA members agree and the consultants strongly agree that a directed history, a directed physical examination,

and a pain control plan should be included in the anesthetic preoperative evaluation.

Recommendations for Preoperative Evaluation of the Patient. A directed pain history, a directed physical examination, and a pain control plan should be included in the anesthetic preoperative evaluation.

III. Preoperative Preparation of the Patient

Preoperative patient preparation includes (1) adjustment or continuation of medications whose sudden cessation may provoke a withdrawal syndrome, (2) treatments to reduce preexisting pain and anxiety, (3) premedications before surgery as part of a multimodal analgesic pain management program, and (4) patient and family education, including behavioral pain control techniques.

There is insufficient literature to evaluate the impact of preoperative adjustment or continuation of medications whose sudden cessation may provoke an abstinence syndrome (*Category D evidence*). Similarly, there is insufficient literature to evaluate the efficacy of the preoperative initiation of treatment either to reduce preexisting pain or as part of a multimodal analgesic pain management program (*Category D evidence*). RCTs are equivocal regarding the impact of patient and family education on patient pain, analgesic use, anxiety, and time to discharge, although features of patient and family education varied across the studies (*Category C2 evidence*). ^{22–35}

The consultants and ASA members strongly agree that patient preparation for perioperative pain management should include appropriate adjustments or continuation of medications to avert an abstinence syndrome, treatment of preexistent pain, or preoperative initiation of therapy for postoperative pain management. The ASA members agree and the consultants strongly agree that anesthesiologists offering perioperative analgesia services should provide, in collaboration with others as appropriate, patient and family education. The consultants and ASA members agree that perioperative patient education should include instruction in behavioral modalities for control of pain and anxiety.

Recommendations for Preoperative Preparation of the Patient. Patient preparation for perioperative pain management should include appropriate adjustments or continuation of medications to avert an abstinence syndrome, treatment of preexistent pain, or preoperative initiation of therapy for postoperative pain management.

Anesthesiologists offering perioperative analgesia services should provide, in collaboration with others as appropriate, patient and family education regarding their important roles in achieving comfort, reporting pain, and in proper use of the recommended analgesic methods. Common misconceptions that overestimate the risk of adverse effects and addiction should be dispelled. Patient education for optimal use of patient-controlled analgesia (PCA) and other sophisticated methods, such as patient-controlled epidural analgesia,

might include discussion of these analgesic methods at the time of the preanesthetic evaluation, brochures and videotapes to educate patients about therapeutic options, and discussion at the bedside during postoperative visits. Such education may also include instruction in behavioral modalities for control of pain and anxiety.

IV. Perioperative Techniques for Pain Management

Perioperative techniques for postoperative pain management include but are not limited to the following single modalities: (1) central regional (*i.e.*, neuraxial) opioid analgesia; (2) PCA with systemic opioids; and (3) peripheral regional analgesic techniques, including but not limited to intercostal blocks, plexus blocks, and local anesthetic infiltration of incisions.

Central regional opioid analgesia: Randomized controlled trials report improved pain relief when use of preincisional epidural or intrathecal morphine is compared with preincisional oral, intravenous, or intramuscular morphine (*Category A2 evidence*). ^{36–39} RCTs comparing preoperative or preincisional intrathecal morphine or epidural sufentanil with saline placebo report inconsistent findings regarding pain relief (*Category C2 evidence*). ^{40–43} RCTs comparing preoperative or preincisional epidural morphine or fentanyl with postoperative epidural morphine or fentanyl are equivocal regarding postoperative pain scores (*Category C2 evidence*). ^{44,45}

Meta-analyses of RCTs^{46–54} report improved pain relief and increased frequency of pruritus in comparisons of postincisional epidural morphine and saline placebo (*Category A1 evidence*); findings for the frequency of nausea or vomiting were equivocal (*Category C1 evidence*). Meta-analyses of RCTs comparing postincisional epidural morphine with intramuscular morphine report improved pain relief and an increased frequency of pruritus (*Category A1 evidence*). One RCT reports improved pain scores and less analgesic use when postincisional intrathecal fentanyl is compared with no postincisional spinal treatment (*Category A3 evidence*).

One RCT reports improved pain scores when postoperative epidural morphine is compared with postoperative epidural saline (*Category A3 evidence*). Meta-analyses of RCTs^{62–70} report improved pain scores and a higher frequency of pruritus and urinary retention when postoperative epidural morphine is compared with intramuscular morphine (*Category A3 evidence*); findings for nausea and vomiting are equivocal (*Category C2 evidence*). Findings from RCTs are equivocal regarding the analgesic efficacy of postoperative epidural fentanyl compared with postoperative IV fentanyl (*Category C2 evidence*)^{71–74}; meta-analytic findings are equivocal for nausea and vomiting and pruritus (*Category C1 evidence*). T2–76

PCA with systemic opioids: Randomized controlled trials report equivocal findings regarding the analgesic efficacy of IV PCA techniques compared with nurse or staff-administered intravenous analgesia (*Category C2 evidence*).^{77–80}

Meta-analysis of RCTs reports improved pain scores when IV PCA morphine is compared with intramuscular morphine (*Category A1 evidence*). Findings from meta-analysis of RCTs comparing epidural PCA and IV PCA opioids are equivocal regarding analgesic efficacy (*Category C1 evidence*). Findings from meta-analyses of RCTs and indicate more analgesic use when IV PCA with a background infusion of morphine is compared with IV PCA without a background infusion (*Category A1 evidence*); findings were equivocal regarding pain relief, nausea and vomiting, pruritus, and sedation (*Category C1 evidence*).

Peripheral regional techniques: For these Guidelines, peripheral regional techniques include peripheral nerve blocks (e.g., intercostal, ilioinguinal, interpleural, or plexus blocks), intraarticular blocks, and infiltration of incisions. RCTs indicate that preincisional intercostal or interpleural bupivacaine compared with saline is associated with improved pain relief (Category A2 evidence). RCTs report improved pain relief and reduced analgesic consumption when postincisional intercostal or interpleural bupivacaine is compared with saline (Category A2 evidence). Meta-analyses of RCTs report equivocal findings for pain relief and analgesic used when postoperative intercostal or interpleural blocks are compared with saline (Category C1 evidence). 110–117

Randomized controlled trials report equivocal pain relief findings when preincisional plexus blocks with bupivacaine are compared with saline (Category C2 evidence). 118-121 Meta-analyses of RCTs^{118–122} report less analgesic use when preincisional plexus blocks with bupivacaine are compared with saline (Category A1 evidence); findings are equivocal for nausea and vomiting (Category C1 evidence). Meta-analysis of RCTs reports lower pain scores when preincisional plexus and other blocks are compared with no block (Category A1 evidence). 123-127 RCTs report equivocal findings for pain scores and analgesic use when postincisional plexus and other blocks are compared with saline or no block (Category C2 evidence). 124,128-132 RCTs report equivocal findings for pain scores and analgesic use when postincisional intraarticular opioids or local anesthetics are compared with saline (Category C2 evidence). 133-139

Meta-analysis of RCTs reports improved pain scores when preincisional infiltration of bupivacaine is compared with saline (Category A1 evidence)^{140–148}; findings for analgesic use are equivocal (Category C1 evidence).^{140,145,147,148–150} Meta-analyses of RCTs are equivocal for pain scores and analgesic use when postincisional infiltration of bupivacaine is compared with saline (Category C1 evidence).^{140,151–160} Meta-analysis of RCTs reports equivocal pain score findings when preincisional infiltration of bupivacaine is compared with postincisional infiltration of bupivacaine (Category C1 evidence).^{140,145,161–164} Meta-analysis of RCTs reports improved pain scores and reduced analgesic use when preincisional infiltration of ropivacaine is compared with saline (Category A1 evidence).^{164–171}

The consultants and ASA members strongly agree that anesthesiologists who manage perioperative pain should use therapeutic options such as epidural or intrathecal opioids, systemic opioid PCA, and regional techniques after thoughtfully considering the risks and benefits for the individual patient; they also strongly agree that these modalities should be used in preference to intramuscular opioids ordered "as needed." The consultants and ASA members also strongly agree that the therapy selected should reflect the individual anesthesiologist's expertise, as well as the capacity for safe application of the modality in each practice setting. Moreover, the consultants and ASA members strongly agree that special caution should be taken when continuous infusion modalities are used, as drug accumulation may contribute to adverse events.

Recommendations for Perioperative Techniques for Pain Management. Anesthesiologists who manage perioperative pain should use therapeutic options such as central regional (i.e., neuraxial) opioids, systemic opioid PCA, and peripheral regional techniques after thoughtfully considering the risks and benefits for the individual patient. These modalities should be used in preference to intramuscular opioids ordered "as needed." The therapy selected should reflect the individual anesthesiologist's expertise, as well as the capacity for safe application of the modality in each practice setting. This capacity includes the ability to recognize and treat adverse effects that emerge after initiation of therapy. Special caution should be taken when continuous infusion modalities are used, as drug accumulation may contribute to adverse events.

V. Multimodal Techniques for Pain Management

Multimodal techniques for pain management include the administration of two or more drugs that act by different mechanisms for providing analgesia. These drugs may be administered *via* the same route or by different routes.

Multimodal techniques with central regional analgesics: Meta-analyses of RCTs 46,49,172–176 report improved pain scores (Category A1 evidence) and equivocal findings for nausea and vomiting and pruritus (Category C1 evidence) when epidural morphine combined with local anesthetics is compared with epidural morphine alone. Meta-analyses of RCTs¹⁷⁷⁻¹⁸⁸ report improved pain scores and more motor weakness when epidural fentanyl combined with local anesthetics is compared with epidural fentanyl alone (Category A1 evidence); equivocal findings are reported for nausea and vomiting and pruritus (Category C1 evidence). Meta-analyses of RCTs^{49,172,176,189–194} report improved pain scores, greater pain relief, and a higher frequency of pruritus (Category A1 evidence) when epidural morphine combined with bupivacaine is compared with epidural bupivacaine alone; equivocal findings are reported for nausea and vomiting (Category C1 evidence). RCTs report equivocal findings when epidural fentanyl combined with bupivacaine is compared with epidural bupivacaine alone (Category C2 evidence). 179-181,188 Meta-analysis of RCTs for the above comparison reports higher frequency of pruritus (Category A1 evidence) 180,181,188,195,196 with equivocal findings for nausea and vomiting (Category C1 evidence). 179–181,188,195–197 RCTs report equivocal findings for pain scores, nausea and vomiting, pruritus, and motor weakness when epidural fentanyl with ropivacaine is compared with epidural ropivacaine (Category C2 evidence). 198–201 Meta-analyses of RCTs 200,202–206 are equivocal for pain scores (Category C2 evidence) and a higher frequency of pruritus when epidural sufentanil combined with ropivacaine is compared with epidural ropivacaine (Category A1 evidence). Meta-analysis of RCTs is equivocal for pain scores when epidural opioids combined with clonidine is compared with epidural opioids (Category C1 evidence). 207–212

Multimodal techniques with systemic analgesics: Metaanalyses of RCTs²¹³⁻²²⁰ report improved pain scores and reduced analgesic use (Category A1 evidence) when intravenous morphine combined with ketorolac is compared with intravenous morphine; equivocal findings are reported for nausea and vomiting (Category C1 evidence). Meta-analyses of RCTs²²¹⁻²²⁶ report equivocal findings for pain scores, analgesic use, or nausea scores when intravenous morphine combined with ketamine is compared with intravenous morphine (Category C1 evidence). RCTs report inconsistent findings for pain scores and morphine use when intravenous patient-controlled opioid analgesia (IV PCA) combined with oral cyclooxygenase-2 (COX-2) selective nonsteroidal antiinflammatory drugs (NSAIDs)²²⁷ or nonselective NSAIDs^{228,229} are compared with IV PCA opioids alone; findings for acetaminophen are equivocal (Category C2 evidence).230 Meta-analyses of RCTs report lower pain scores and reduced opioid use when IV opioids combined with calcium channel blockers (i.e., gabapentin, pregabalin) is compared with IV opioids alone (Category A1 evidence) $^{231-240}$; no differences in nausea or vomiting are reported (Category C1 evidence). 233-236,238,241

The consultants and ASA members strongly agree that whenever possible, anesthesiologists should use multimodal pain management therapy. The ASA members agree and the consultants strongly agree that acetaminophen should be considered as part of a postoperative multimodal pain management regimen; both the consultants and ASA members agree that COX-2 selective NSAIDs (COXIBs), nonselective NSAIDs, and calcium channel α -2- δ antagonists (gabapentin and pregabalin) should be considered as part of a postoperative multimodal pain management regimen. Moreover, the ASA members agree and the consultants strongly agree that, unless contraindicated, patients should receive an around-the-clock regimen of NSAIDs, COXIBs, or acetaminophen. Both the consultants and ASA members strongly agree that (1) regional blockade with local anesthetics should be considered as part of a multimodal approach for pain management; (2) dosing regimens should be administered to optimize efficacy while minimizing the risk of adverse events; and (3) the choice of medication, dose, route, and duration of therapy should be individualized.

Recommendations for Multimodal Techniques. Whenever possible, anesthesiologists should use multimodal pain management therapy. Central regional blockade with local anesthetics should be considered. Unless contraindicated, patients should receive an around-the-clock regimen of COXIBs, NSAIDs, or acetaminophen. Dosing regimens should be administered to optimize efficacy while minimizing the risk of adverse events. The choice of medication, dose, route, and duration of therapy should be individualized.

VI. Patient Subpopulations

Some patient groups are at special risk for inadequate pain control and require additional analgesic considerations. Patient populations at risk include (1) pediatric patients, (2) geriatric patients, and (3) critically ill or cognitively impaired patients, or other patients who may have difficulty communicating. The Task Force believes that genetics and gender modify the pain experience and response to analgesic therapies. In addition, the Task Force believes that patient race, ethnicity, culture, gender, and socioeconomic status influence access to treatment as well as pain assessment by healthcare providers.

Pediatric Patients. The Task Force believes that optimal care for infants and children (including adolescents) requires special attention to the biopsychosocial nature of pain. This specific patient population presents developmental differences in their experience and expression of pain and suffering, and their response to analgesic pharmacotherapy. Caregivers in both the home and hospital may have misperceptions regarding the importance of analgesia as well as its risks and benefits. In the absence of a clear source of pain or obvious pain behavior, caregivers may assume that pain is not present and defer treatment. Safe methods for providing analgesia are underused in pediatric patients for fear of opioid-induced respiratory depression.

The emotional component of pain is particularly strong in infants and children. Absence of parents, security objects, and familiar surroundings may cause as much suffering as the surgical incision. Children's fear of injections makes intramuscular or other invasive routes of drug delivery aversive. Even the valuable technique of topical analgesia before injections may not lessen this fear.

A variety of techniques may be effective in providing analgesia in pediatric patients. Many are the same as for adults, although some (e.g., caudal analgesia) are more commonly used in children. The Task Force believes that it is important for caregivers to recognize that pediatric patients require special consideration to ensure optimal perioperative analgesia.

The ASA members and consultants strongly agree that (1) perioperative care for children undergoing painful procedures or surgery requires developmentally appropriate pain assessment and therapy; (2) analgesic therapy should depend upon age, weight, and comorbidity, and unless contraindi-

cated should involve a multimodal approach; and (3) because many analgesic medications are synergistic with sedating agents, it is imperative that appropriate monitoring be used during the procedure and recovery. The ASA members agree and the consultants strongly agree that behavioral techniques, especially important in addressing the emotional component of pain, should be applied whenever feasible.

Recommendations for Pediatric Patients. Aggressive and proactive pain management is necessary to overcome the historic undertreatment of pain in children. Perioperative care for children undergoing painful procedures or surgery requires developmentally appropriate pain assessment and therapy. Analgesic therapy should depend upon age, weight, and comorbidity, and unless contraindicated should involve a multimodal approach. Behavioral techniques, especially important in addressing the emotional component of pain, should be applied whenever feasible.

Sedative, analgesic, and local anesthetics are all important components of appropriate analgesic regimens for painful procedures. Because many analgesic medications are synergistic with sedating agents, it is imperative that appropriate monitoring be used during the procedure and recovery.

Geriatric Patients. Elderly patients suffer from conditions such as arthritis or cancer that render them more likely to undergo surgery. The Task Force believes that pain is often undertreated, and elderly individuals may be more vulnerable to the detrimental effects of such undertreatment. The physical, social, emotional, and cognitive changes associated with aging have an impact on perioperative pain management. These patients may have different attitudes than younger adult patients in expressing pain and seeking appropriate therapy. Altered physiology changes the way analgesic drugs and local anesthetics are distributed and metabolized and frequently requires dose alterations. Techniques effective in younger adults may also benefit geriatric patients without an age-related increase in adverse effects. One observational study suggests that perioperative analgesics are provided in lower dosages to older adults than to younger adults (Category B2 evidence). 242 The Task Force believes that, although the reasons for lower perioperative analgesic doses in the elderly are unclear, undertreatment of pain in elderly persons is widespread.

The ASA members and consultants strongly agree that (1) pain assessment and therapy should be integrated into the perioperative care of geriatric patients; (2) pain assessment tools appropriate to a patient's cognitive abilities should be used; and (3) dose titration should be done to ensure adequate treatment while avoiding adverse effects such as somnolence in this vulnerable group, who may be taking other medications. The ASA members agree and the consultants strongly agree that extensive and proactive evaluation and questioning should be conducted to overcome barriers that hinder communication regarding unrelieved pain.

Recommendations for Geriatric Patients. Pain assessment and therapy should be integrated into the perioperative care

of geriatric patients. Pain assessment tools appropriate to a patient's cognitive abilities should be used. Extensive and proactive evaluation and questioning may be necessary to overcome barriers that hinder communication regarding unrelieved pain. Anesthesiologists should recognize that geriatric patients may respond differently than younger patients to pain and analgesic medications, often because of comorbidity. Vigilant dose titration is necessary to ensure adequate treatment while avoiding adverse effects such as somnolence in this vulnerable group, who are often taking other medications (including alternative and complementary agents).

Other Subpopulations. Patients who are critically ill, cognitively impaired (e.g., Alzheimer's disease), or who otherwise have difficulty communicating (e.g., cultural or language barriers) present unique challenges to perioperative pain management. The Task Force believes that techniques that reduce drug dosages required to provide effective analgesia (e.g., regional analgesia and multimodal analgesia) may be suitable for such patients. Behavioral modalities and techniques such as PCA that depend upon self-administration of analgesics are generally less suitable for the cognitively impaired. The literature is insufficient to evaluate the application of pain assessment methods or pain management techniques specific to these populations (Category D evidence).

The consultants and ASA members strongly agree that anesthesiologists should recognize that patients who are critically ill, cognitively impaired, or have communication difficulties may require additional interventions to ensure optimal perioperative pain management. Moreover, the ASA members agree and the consultants strongly agree that anesthesiologists should consider a therapeutic trial of an analgesic in patients with increased blood pressure and heart rate or agitated behavior, when causes other than pain have been excluded.

Recommendations for Other Subpopulations. Anesthesiologists should recognize that patients who are critically ill, cognitively impaired, or have communication difficulties may require additional interventions to ensure optimal perioperative pain management. Anesthesiologists should consider a therapeutic trial of an analgesic in patients with increased blood pressure and heart rate or agitated behavior when causes other than pain have been excluded.

Appendix 1: Summary of Recommendations

I. Institutional Policies and Procedures for Providing Perioperative Pain Management

- Anesthesiologists offering perioperative analgesia services should provide, in collaboration with other healthcare professionals as appropriate, ongoing education and training to ensure that hospital personnel are knowledgeable and skilled with regard to the effective and safe use of the available treatment options within the institution.
 - Educational content should range from basic bedside pain assessment to sophisticated pain management techniques (e.g.,

- epidural analgesia, PCA, and various regional anesthesia techniques) and nonpharmacologic techniques (*e.g.*, relaxation, imagery, hypnotic methods).
- For optimal pain management, ongoing education and training are essential for new personnel, to maintain skills, and whenever therapeutic approaches are modified.
- Anesthesiologists and other healthcare providers should use standardized, validated instruments to facilitate the regular evaluation and documentation of pain intensity, the effects of pain therapy, and side effects caused by the therapy.
- Anesthesiologists responsible for perioperative analgesia should be available at all times to consult with ward nurses, surgeons, or other involved physicians.
 - They should assist in evaluating patients who are experiencing problems with any aspect of perioperative pain relief.
- Anesthesiologists providing perioperative analgesia services should do so within the framework of an Acute Pain Service.
 - They should participate in developing standardized institutional policies and procedures.

II. Preoperative Evaluation of the Patient

 A directed pain history, a directed physical examination, and a pain control plan should be included in the anesthetic preoperative evaluation.

III. Preoperative Preparation of the Patient

- Patient preparation for perioperative pain management should include appropriate adjustments or continuation of medications to avert an abstinence syndrome, treatment of preexistent pain, or preoperative initiation of therapy for postoperative pain management.
- Anesthesiologists offering perioperative analgesia services should provide, in collaboration with others as appropriate, patient and family education regarding their important roles in achieving comfort, reporting pain, and in proper use of the recommended analgesic methods.
 - Common misconceptions that overestimate the risk of adverse effects and addiction should be dispelled.
 - Patient education for optimal use of PCA and other sophisticated methods, such as patient-controlled epidural analgesia, might include discussion of these analgesic methods at the time of the preanesthetic evaluation, brochures and videotapes to educate patients about therapeutic options, and discussion at the bedside during postoperative visits.
 - Such education may also include instruction in behavioral modalities for control of pain and anxiety.

IV. Perioperative Techniques for Pain Management

- Anesthesiologists who manage perioperative pain should use therapeutic options such as epidural or intrathecal opioids, systemic opioid PCA, and regional techniques after thoughtfully considering the risks and benefits for the individual patient.
 - These modalities should be used in preference to intramuscular opioids ordered "as needed."
- The therapy selected should reflect the individual anesthesiologist's expertise, as well as the capacity for safe application of the modality in each practice setting.
 - This capacity includes the ability to recognize and treat adverse effects that emerge after initiation of therapy.
- Special caution should be taken when continuous infusion modalities are used because drug accumulation may contribute to adverse events.

V. Multimodal Techniques for Pain Management

- Whenever possible, anesthesiologists should use multimodal pain management therapy.
 - Unless contraindicated, patients should receive an aroundthe-clock regimen of NSAIDs, COXIBs, or acetaminophen.
 - O Regional blockade with local anesthetics should be considered.
- Dosing regimens should be administered to optimize efficacy while minimizing the risk of adverse events.
- The choice of medication, dose, route, and duration of therapy should be individualized.

VI. Patient Subpopulations

- · Pediatric patients
 - Aggressive and proactive pain management is necessary to overcome the historic undertreatment of pain in children.
 - Perioperative care for children undergoing painful procedures or surgery requires developmentally appropriate pain assessment and therapy.
 - Analgesic therapy should depend upon age, weight, and comorbidity, and unless contraindicated should involve a multimodal approach.
 - Behavioral techniques, especially important in addressing the emotional component of pain, should be applied whenever feasible.
 - Sedative, analgesic, and local anesthetics are all important components of appropriate analgesic regimens for painful procedures.
 - Because many analgesic medications are synergistic with sedating agents, it is imperative that appropriate monitoring be used during the procedure and recovery.
- · Geriatric patients
 - Pain assessment and therapy should be integrated into the perioperative care of geriatric patients.
 - Pain assessment tools appropriate to a patient's cognitive abilities should be used. Extensive and proactive evaluation and questioning may be necessary to overcome barriers that hinder communication regarding unrelieved pain.
 - Anesthesiologists should recognize that geriatric patients may respond differently than younger patients to pain and analgesic medications, often because of comorbidity.
 - Vigilant dose titration is necessary to ensure adequate treatment while avoiding adverse effects such as somnolence in this vulnerable group, who are often taking other medications (including alternative and complementary agents).
- Other subpopulations
 - Anesthesiologists should recognize that patients who are critically ill, cognitively impaired, or have communication difficulties may require additional interventions to ensure optimal perioperative pain management.
 - Anesthesiologists should consider a therapeutic trial of an analgesic in patients with increased blood pressure and heart rate or agitated behavior when causes other than pain have been excluded.

Appendix 2: Methods and Analyses

A. State of the Literature

For these updated Guidelines, a review of studies used in the development of the original Guidelines was combined with studies published subsequent to approval of the original Guidelines in 2003.* The scientific assessment of these Guidelines was based on evidence linkages or statements regarding potential

relationships between clinical interventions and outcomes. The interventions listed below were examined to assess their relationship to a variety of outcomes related to the management of acute pain in the perioperative setting.

Institutional Policies and Procedures for Providing Perioperative Pain Management

Education and training of healthcare providers

Monitoring of patient outcomes

Documentation of monitoring activities

Monitoring of outcomes at an institutional level

24-h availability of anesthesiologists providing perioperative pain management

Acute pain service

Preoperative Evaluation of the Patient

A directed pain history (e.g., medical record review and patient interview to include current medications, adverse effects, preexisting pain conditions, medical conditions that would influence a pain therapy, nonpharmacologic pain therapies, alternative and complementary therapies)

A directed physical examination

Consultations with other healthcare providers (e.g., nurses, surgeons, pharmacists)

Preoperative Preparation of the Patient

Preoperative adjustment or continuation of medications whose sudden cessation may provoke an abstinence syndrome

Preoperative treatment(s) to reduce preexisting pain and anxiety Premedication(s) before surgery as part of a multimodal analgesic pain management program

Patient and family education

Perioperative Techniques for Pain Management

Epidural or intrathecal analgesia with opioids (vs. epidural placebo, epidural local anesthetics, or IV, intramuscular, or oral opioids) Patient-controlled analgesia with opioids:

IV PCA versus nurse-controlled or continuous IV

IV PCA versus intramuscular

Epidural PCA versus epidural bolus or infusion

Epidural PCA versus IV PCA

IV PCA with background infusion of opioids *versus* no background infusion

Regional analgesia with local anesthetics or opioids

Intercostal or interpleural blocks

Plexus and other blocks

Intraarticular opioids, local anesthetics or combinations Infiltration of incisions

Multimodal Techniques (Epidural, IV, or Regional Techniques)

Two or more analgesic agents, one route *versus* a single agent, one route Epidural or intrathecal analgesia with opioids combined with:

Local anesthetics versus epidural opioids

Local anesthetics versus epidural local anesthetics

Clonidine versus epidural opioids

IV opioids combined with:

Clonidine versus IV opioids

Ketorolac versus IV opioids

Ketamine versus IV opioids

Oral opioids combined with NSAIDs, COXIBs, or acetaminophen *versus* oral opioids

Two or more drug delivery routes versus a single route

Epidural or intrathecal analgesia with opioids combined with IV, intramuscular, oral, transdermal, or subcutaneous analgesics *versus* epidural opioids

IV opioids combined with oral NSAIDs, COXIBs, or acetaminophen *versus* IV opioids

Nonpharmacologic, alternative, or complementary pain management combined with pharmacologic pain management *versus* pharmacologic pain management

Special Patient Populations

Pain management techniques for pediatric patients

Pain assessment techniques

Dose level adjustments

Avoidance of repetitive diagnostic evaluation (heel sticks) for neonates Pain management techniques for geriatric patients

Pain assessment techniques

Dose level adjustments

Pain management techniques for other special populations (e.g., cognitively impaired, critically ill, patients with difficulty communicating)

Pain assessment methods specific to special populations

Pain management techniques specific to special populations

For the literature review, potentially relevant clinical studies were identified *via* electronic and manual searches of the literature. The electronic and manual searches covered a 49-yr period from 1963 through 2011. More than 2,000 citations were identified initially, yielding a total of 1,784 nonoverlapping articles that addressed topics related to the evidence linkages. After the articles were reviewed, 1,153 studies did not provide direct evidence and were eliminated subsequently. A total of 631 articles contained direct linkage-related evidence. A complete bibliography used to develop these Guidelines, organized by section, is available as Supplemental Digital Content 2, http://links.lww.com/ALN/A781.

Initially, each pertinent outcome reported in a study was classified as supporting an evidence linkage, refuting a linkage, or equivocal. The results were then summarized to obtain a directional assessment for each evidence linkage before conducting formal meta-analyses. Literature pertaining to four evidence linkage categories contained enough studies with well-defined experimental designs and statistical information sufficient for meta-analyses (table 1). These linkages were: (1) epidural or intrathecal opioids, (2) patient-controlled analgesia, (3) regional analgesia, and (4) two or more anesthetic drugs *versus* a single drug.

General variance-based, effect-size estimates or combined probability tests were obtained for continuous outcome measures, and Mantel-Haenszel odds ratios were obtained for dichotomous outcome measures. Two combined probability tests were used as follows: (1) the Fisher combined test, producing chi-square values based on logarithmic transformations of the reported P values from the independent studies, and (2) the Stouffer combined test, providing weighted representation of the studies by weighting each of the standard normal deviates by the size of the sample. An odds ratio procedure based on the Mantel-Haenszel method for combining study results using 2 × 2 tables was used with outcome frequency information. An acceptable significance level was set at P < 0.01 (one-tailed). Tests for heterogeneity of the independent studies were conducted to assure consistency among the study results. DerSimonian-Laird random-effects odds ratios were obtained when significant heterogeneity was found (P < 0.01). To control for potential publishing bias, a "fail-safe n" value was calculated. No search for unpublished studies was conducted, and no

reliability tests for locating research results were done. To be accepted as significant findings, Mantel-Haenszel odds ratios must agree with combined test results whenever both types of data are assessed. In the absence of Mantel-Haenszel odds ratios, findings from both the Fisher and weighted Stouffer combined tests must agree with each other to be acceptable as significant.

For the previous update of the Guidelines, interobserver agreement among Task Force members and two methodologists was established by interrater reliability testing. Agreement levels using a kappa (k) statistic for two-rater agreement pairs were as follows: (1) type of study design, k=0.63-0.94; (2) type of analysis, k=0.39-0.89; (3) evidence linkage assignment, k=0.74-0.96; and (4) literature inclusion for database, k=0.75-0.88. Three-rater chance-corrected agreement values were: (1) study design, Sav = 0.80, Var (Sav) = 0.007; (2) type of analysis, Sav = 0.59, Var (Sav) = 0.032; (3) linkage assignment, Sav = 0.73 Var (Sav) = 0.010; (4) literature database inclusion, Sav = 0.83 Var (Sav) = 0.015. These values represent moderate levels of agreement. For the updated Guidelines, the same two methodologists involved in the original Guidelines conducted the literature review.

The findings of the literature analyses were supplemented by the opinions of Task Force members after considering opinions derived from a variety of sources, including informal commentary and comments from postings of the draft document on the ASA web site. In addition, opinions obtained from consultant surveys, open forum commentary, and other sources used in the original Guidelines were reviewed and considered.

B. Consensus-based Evidence

Consensus was obtained from multiple sources, including (1) survey opinion from consultants who were selected based on their knowledge or expertise in acute pain management, (2) survey opinions solicited from active members of the ASA, (3) testimony from attendees of a publicly held open forum at a national anesthesia meeting (original Guidelines only), (4) Internet commentary, and (5) Task Force opinion and interpretation. The survey rate of return was 62% (n = 53 of 85) for the consultants (table 2), and 268 surveys were received from active ASA members (table 3).

For the previous update of the Guidelines, an additional survey was sent to the expert consultants asking them to indicate which, if any, of the evidence linkages would change their clinical practices if the Guidelines were instituted. The rate of return was 70.1% (n = 61 of 87). The percentages of responding consultants expecting *no change* associated with each linkage were as follows: (1) proactive planning 82.0%, (2) education and training 88.5%, (3) education or participation of patient and family 80.3%, (4) monitoring or documentation 77.0%, (5) availability of anesthesiologists 90.2%, (6) institutional protocols 86.9%, (7) use of PCA, epidural, or regional techniques 90.2%, (8) use of multimodality techniques 88.5%, (9) organizational characteristics 90.2%, (10) pediatric techniques 95.1%, (11) geriatric techniques 91.8%, and (12) ambulatory surgery techniques 85.2%.

Sixty-five percent of the respondents indicated that the Guide-lines would have *no effect* on the amount of time spent on a typical case, and 24% indicated that there would be an increase of the amount of time spent on a typical case with the implementation of these Guidelines (mean time increase = 3.4 min). Eighty-nine percent indicated that new equipment, supplies, or training would *not* be needed to implement the Guidelines, and 92% indicated that implementation of the Guidelines would *not* require changes in practice that would affect costs.

Table 1. Meta-analysis Summary

		Fisher	r Weighted					Heterog	eneity	
		Chi-	P	Stouffer	P	Effect	Odds	Confidence		Effect
Evidence Linkages	Ν	square	Value	Zc	Value	Size	Ratio	Interval	Values	Size
Perioperative techniques Epidural/intrathecal opioids Postincisional Morphine vs. saline										
Pain scores or relief Nausea or vomiting Pruritus Morphine vs. IM morphine	6 9 8	51.50	0.001	-3.50	0.001	-0.35	1.17* 7.35	0.32–5.56 3.84–14.08	0.342	0.694 0.001 0.186
Pain scores Pruritus Postoperative	6	52.16	0.001	-3.79	0.001	-0.44	6.24	2.28–17.08	0.995	0.788 0.779
Morphine vs. IM morphine Pain scores or relief Nausea or vomiting Pruritus Urinary retention Fentanyl vs. IV fentanyl	7 9 5 7	81.29	0.001	-7.52	0.001	-0.57	0.76 5.45 3.10	0.35–1.66 1.62–18.36 1.31–7.32	0.097	0.001 0.442 0.980 0.865
Nausea or vomiting Pruritus	5 5						0.73* 1.17	0.08–4.92 0.30–4.54		0.001 0.731
PCA IV PCA vs. IM morphine									(cor	ntinued)

Table 1. Continued

	Fishe Chi	- P	Weighted Stouffer	P				Heterog P	Effect
Evidence Linkages	V squa	re Value	Zc	Value	Size	Ratio	Interval	Values	Size
Pain scores Epidural PCA vs. IV PCA opioids	8 52.2	6 0.001	-4.01	0.001	-0.22			0.700	0.550
Pain scores	5 37.9	0.001	-2.17	0.015	-0.33			0.999	0.951
PCA with background morphine Pain scores or relief	6 25.9	0.011	-2.25	0.012	0.07			0.315	0.138
	0 20.3		6.12	0.001	0.35			0.001	0.001
9	9					1.01	0.57-1.78		0.666
	7 6 16.4	4 0.172	-1.62	0.053	-0.03	0.99	0.43–2.29	0.675	0.522 0.628
Regional analgesia	0 10.4	·4 U.172	1.02	0.055	0.03			0.073	0.020
Intercostal or interpleural block	(S								
Postoperative vs. saline			4.70					0.000	0.470
	7 54.1		-1.79		-0.38			0.663	0.479
Analgesic use Plexus and other blocks	5 36.3	0.001	-1.51	0.066	-0.34			0.263	0.381
Preincisional vs. saline									
	5 52.1	3 0.001	-5.62	0.001	-0.37			0.146	0.057
	5					0.51	0.15-1.73		0.769
Preincisional vs. no block									
	5 45.1	5 0.001	-4.41	0.001	-0.32			0.061	0.174
Infiltration of incisions Preincisional bupivacaine vs	calina								
•	. <i>saii le</i> 9 84.8	3 0.001	-3.51	0.001	-0.32			0.002	0.001
	6 21.2		-2.01		-0.11			0.662	0.605
Postincisional bupivacaine v	s. saline	!							
	8 42.5		-2.10		-0.17			0.044	0.051
	9 53.7	1 0.001	-2.12	0.017	-0.20			0.039	0.024
Pre- vs. postincisional bupiv Pain scores	acaine 6 39.2	8 0.001	1.02	0.154	0.02			0.001	0.001
Preincisional ropivacaine vs.		.0.001	1.02	0.154	0.02			0.001	0.001
	5 44.1	4 0.001	-3.96	0.001	-0.31			0.964	0.556
	7 45.5	0.001	-3.90	0.001	-0.43			0.001	0.001
Multimodality techniques									
Two or more vs. single drug, sar			la i a						
Epidural morphine + local a Pain scores	7 42.9		–2.32	0.010	-0.22			0.466	0.167
	, 42.5 6	0.001	2.02	0.010	0.22	0.80	0.40-1.57	0.400	0.829
	6						0.93-4.36		0.176
Epidural fentanyl + local and									
Pain scores 1		1 0.001	-3.11	0.001	-0.29	0.77	0.40.4.07	0.006	0.001
3	1 2						0.46–1.27 0.55–1.56		0.304 0.266
	9						1.57–6.65		0.200
Epidural morphine + bupiva		. bupivaca	aine			0.20	1.07 0.00		0.011
		0.001	-3.03	0.001	-0.25			0.470	0.245
	5						1.31–8.92		0.352
9	8						0.62-2.48		0.858
Pruritus Epidural fentanyl + bupivaca	6 aine vs	hunivacair	26			7.35	2.82–19.15		0.584
	ii ie vs. i 7	Zapivacali				1.27	0.58–2.80		0.329
	5						1.02-8.23		0.840
Epidural sufentanil + ropivad					_				
		4 0.001	-2.09	0.018	-0.17	4.00	0.01.0.07	0.730	0.425
Pruritus	6					4.32	2.31–8.07	loor	0.705 ntinued)
								(001	iai iaca)

Table 1. Continued

		Fisher	Weighted				Heterogeneity			
		Chi-	Р	Stouffer	P	Effect	Odds	Confidence	P	Effect
Evidence Linkages	Ν	square	Value	Zc	Value	Size	Ratio	Interval	Values	Size
Epidural opioids + clonid	ine v	s. opioid	S							
Pain scores	6	45.77	0.001	-1.27	0.102	-0.12			0.001	0.001
IV morphine + ketorolac	vs. IV	/ morphir	пе							
Pain scores	6	44.18	0.001	-3.95	0.001	-0.30			0.987	0.992
Analgesic use	6	72.42	0.001	-7.17	0.001	-0.59			0.001	0.001
Nausea or vomiting	6						1.04	0.54 - 2.00		0.937
IV morphine + ketamine v	/s. IV	' morphir	ne							
Pain scores or relief	6	39.95	0.001	-0.81	0.209	-0.11			0.056	0.001
Analgesic use	6	37.12	0.001	-1.00	0.159	-0.08			0.027	0.001
Nausea	6	26.45	0.009	0.48	0.316	-0.04			0.165	0.037
Two or more routes vs. single	rout	е								
IV opioids combined with	calc	ium char	nel blo	ckers (gaba _l	pentin, _l	pregaba	lin) vs.	IV opioids		
Pain scores	7	54.03	0.001	-3.82	0.001	-0.29			0.700	0.850
Opioid use	10	111.66	0.001	-12.07	0.001	-0.48			0.001	0.001
Nausea	6						1.04	0.55-1.98		0.800
Vomiting	5						0.86	0.41–1.83		0.970

^{*} Random effects odds ratio.

Table 2. Consultant Survey Responses*

			Р	ercent Re	sponding to E	ach Item	
		N	Strongly Agree	Agree	Equivocal	Disagree	Strongly Disagree
Pro	Anesthesiologists offering perioperative Anesthesiologists offering perioperative analgesia services should provide, in collaboration with other healthcare professionals as appropriate, ongoing education and training of hospital personnel regarding the effective and safe use of the						
2.	available treatment options within the institution Anesthesiologists and other healthcare providers should use standardized, validated instruments to facilitate the regular evaluation and documentation of pain intensity, the effects of pain therapy, and side effects caused by the therapy	53	86.8* 67.9*	11.3 26.4	1.9 5.7	0.0	0.0
3.	Anesthesiologists responsible for perioperative analgesia should be available at all times to consult with ward nurses, surgeons, or other involved physicians and should assist in evaluating patients who are experiencing problems with any aspect of perioperative pain relief	53	56.6*	26.4	17.0	0.0	0.0
4.	Anesthesiologists should provide analgesia services within the framework of an Acute Pain Service and participate in developing standardized institutional policies and procedures	53	73.6*	26.4	0.0	0.0	0.0 (continued)

 $IM = intramuscular; \ IV = intravenous; \ PCA = patient-controlled \ analgesia.$

Table 2. Continued

		Р	ercent Re	sponding to E	ach Item	
	N	Strongly Agree	Agree	Equivocal	Disagree	Strongly Disagree
5. An integrated approach to perioperative pain management (e.g., ordering, administering, and transitioning therapies, transferring responsibility for pain therapy, outcomes assessment, continuous quality improvement) should be used to minimize		70.04	24.5			
analgesic gaps	53	73.6*	24.5	1.9	0.0	0.0
 II. Preoperative Evaluation of the Patient 6. A directed pain history, a directed physical examination, and a pain control plan should be included in the anesthetic preoperative evaluation 	52	57.7*	36.5	3.8	1.9	0.0
 III. Preoperative Preparation of the Patient 7. Patient preparation for perioperative pain management should include appropriate adjustments or continuation of medications to avert an abstinence syndrome, treatment of preexistent pain, or preoperative initiation of therapy for 						
postoperative pain management 8. Anesthesiologists offering perioperative analgesia services should provide, in collaboration with others as appropriate,	53	77.4*	18.9	3.8	0.0	0.0
patient and family education 9. Perioperative patient education should include instruction in behavioral modalities	53	50.9*	35.8	7.5	5.7	0.0
for control of pain and anxiety	53	37.7	39.6*	13.2	7.5	1.9
IV. Perioperative Techniques for Pain Managem 10. Anesthesiologists who manage perioperative pain should use therapeutic options such as epidural or intrathecal opioids, systemic opioid PCA, and regional techniques after thoughtfully considering the risks and benefits for the individual	ent					
patient 11. These modalities should be used in preference to intramuscular opioids	53	86.8*	13.2	0.0	0.0	0.0
ordered "as needed" 12. The therapy selected should reflect the individual anesthesiologist's expertise, as	53	79.2*	11.3	3.8	1.9	3.8
well as the capacity for safe application of the modality in each practice setting 13. Special caution should be taken when continuous infusion modalities are used because drug accumulation may	53	79.2*	17.0	0.0	3.8	0.0
contribute to adverse events	53	69.8*	26.4	1.9	1.9	0.0
V. Multimodal Techniques for Pain Management 14. Whenever possible, anesthesiologists should use multimodal pain management						_
therapy	53	71.7*	28.3	0.0	0.0	0.0 (continued)

Table 2. Continued

		F	ercent Re	sponding to E	ach Item	
	N	Strongly Agree	Agree	Equivocal	Disagree	Strongly Disagree
15. The following drugs should be consider as part of a postoperative multimodal management regimen:						
COX-2 selective NSAIDs (COXIBs) Nonselective NSAIDs Acetaminophen Calcium channel α -2- δ antagonists	53 52 53	49.1 19.2 62.3*	34.0* 57.7* 32.1	15.1 23.1 5.7	1.9 0.0 0.0	0.0 0.0 0.0
gabapentin, pregabalin) 16. Unless contraindicated, all patients si receive an around-the-clock regimen	53 nould	22.6	50.9*	26.4	0.0	0.0
NSAIDs, COXIBs, or acetaminophen 17. Regional blockade with local anesthe should be considered as part of a	51	54.9*	23.5	7.8	9.8	3.9
multimodal approach for pain managers. 18. Dosing regimens should be administed to optimize efficacy while minimizing	ered	73.1*	25.0	1.9	0.0	0.0
risk of adverse events 19. The choice of medication, dose, route, a duration of therapy should be individualized.	52 nd	86.5* 73.1*	13.5 26.9	0.0	0.0	0.0
 VI. Patient Subpopulations Pediatric patients 20. Perioperative care for children undergouinful procedures or surgery require developmentally appropriate pain assessment and therapy 21. Analgesic therapy should depend upon 	53	73.6*	24.5	1.9	0.0	0.0
age, weight, and comorbidity and unl contraindicated should involve a multimodal approach 22. Behavioral techniques, especially imp	ess 53 portant	67.9*	30.2	1.9	0.0	0.0
in addressing the emotional compone pain, should be applied whenever fea 23. Because many analgesic medications synergistic with sedating agents, it is imperative that appropriate monitorin- used during the procedure and recov	sible 53 are	50.9* 83.0*	30.2 17.0	18.9	0.0	0.0
Geriatric patients 24. Pain assessment and therapy should integrated into the perioperative care	be	00.0	17.0	0.0	0.0	0.0
geriatric patients 25. Pain assessment tools appropriate to	53	73.6*	26.4	0.0	0.0	0.0
patient's cognitive abilities should be 26. Extensive and proactive evaluation ar questioning should be conducted to overcome barriers that hinder	used 53	77.4*	22.6	0.0	0.0	0.0
communication regarding unrelieved	pain 53	58.5*	35.8	5.7	0.0	0.0 (continued)

Table 2. Continued

	Percent Responding to Each Item								
	N	Strongly Agree	Agree	Equivocal	Disagree	Strongly Disagree			
27. Dose titration should be done to ensure adequate treatment while avoiding adverse effects such as somnolence in this vulnerable group, who may be taking other medications	53	77.4*	22.6	0.0	0.0	0.0			
Other Subpopulations 28. Anesthesiologists should recognize that patients who are critically ill, cognitively impaired, or have communication difficulties may require additional interventions to ensure optimal perioperative pain management 29. Anesthesiologists should consider a therapeutic trial of an analgesic in patients	53	73.6*	24.5	1.9	0.0	0.0			
with elevated blood pressure and heart rate or agitated behavior when causes other than pain have been excluded	53	50.9*	37.7	9.4	1.9	0.0			

^{*} Indicates the median.

Table 3. ASA Member Survey Responses*

		Pe	ercent Res	ponding to E	ach Item	
	N	Strongly Agree	Agree	Equivocal	Disagree	Strongly Disagree
I. Institutional Policies and Procedures for Providing Perioperative Pain Management 1. Anesthesiologists offering perioperative analgesia services should provide, in collaboration with other healthcare professionals as appropriate, ongoing education and training of hospital personnel regarding the effective and safe use of the available treatment options within the Institution 2. Anesthesiologists and other healthcare providers should use standardized, validated instruments to facilitate the regular evaluation and documentation of poin intensity, the effects of poin thereby.	268	53.0*	37.7	4.1	3.7	1.5
pain intensity, the effects of pain therapy, and side effects caused by the therapy 3. Anesthesiologists responsible for perioperative analgesia should be available at all times to consult with ward nurses, surgeons, or other involved physicians and should assist in evaluating patients who are experiencing problems with any aspect of perioperative pain relief	268 267	52.2* 38.9	35.5 36.0*	7.5	3.7	1.1 2.6 (continued)

COX-2 = cyclooxygenase-2; N = number of consultants who responded to each item; <math>NSAID = nonsteroidal antiinflammatory drug; PCA = patient-controlled analgesia.

Table 3. Continued

		Pe	rcent Res	ponding to E	ach Item	
	N	Strongly Agree	Agree	Equivocal	Disagree	Strongly Disagree
 4. Anesthesiologists should provide analgesia services within the framework of an Acute Pain Service and participate in developing standardized institutional policies and Procedures 5. An integrated approach to perioperative pain management (e.g., ordering, administering, and transitioning therapies, transferring responsibility for pain therapy, 	268	39.9	39.2*	14.9	3.4	2.6
outcomes assessment, continuous quality improvement) should be used to minimize analgesic gaps	269	46.5	44.6*	7.4	1.5	0.0
 II. Preoperative Evaluation of the Patient 6. A directed pain history, a directed physical examination, and a pain control plan should be included in the anesthetic preoperative evaluation 	267	30.3	39.7*	18.4	9.4	2.2
 III. Preoperative Preparation of the Patient 7. Patient preparation for perioperative pain management should include appropriate adjustments or continuation of medications to avert an abstinence syndrome, treatment of preexistent pain, or preoperative initiation of therapy for 						
postoperative initiation of therapy for postoperative pain management 8. Anesthesiologists offering perioperative analgesia services should provide, in collaboration with others as appropriate,	266	51.5*	41.7	5.7	1.1	0.0
patient and family education 9. Perioperative patient education should include instruction in behavioral modalities	268	28.7	56.7*	10.1	3.7	0.8
for control of pain and anxiety IV. Perioperative Techniques for Pain Managem 10. Anesthesiologists who manage perioperative pain should use therapeutic options such as epidural or intrathecal opioids, systemic opioid PCA, and regional techniques after thoughtfully considering the risks and	269 ent	22.7	42.8*	27.1	5.9	1.5
benefits for the individual patient 11. These modalities should be used in preference to intramuscular opioids	269	65.4*	31.2	1.9	1.1	0.4
ordered "as needed" 12. The therapy selected should reflect the individual anesthesiologist's expertise, as well as the capacity for safe application of	269	65.8*	24.9	7.5	1.1	0.7
the modality in each practice setting 13. Special caution should be taken when continuous infusion modalities are used	269	70.6*	26.8	1.9	0.7	0.0
because drug accumulation may contribute to adverse events	268	67.6*	30.2	1.1	1.1	0.0 (continued)

Table 3. Continued

		Pe	ercent Res	ponding to E	ach Item	
	N	Strongly Agree	Agree	Equivocal	Disagree	Strongly Disagree
 V. <i>Multimodal Techniques for Pain Management</i> 14. Whenever possible, anesthesiologists should use multimodal pain management therapy 15. The following drugs should be considered as part of a postoperative multimodal pain 		56.2*	28.1	12.4	2.6	0.7
management regimen: COX-2 selective NSAIDs (COXIBs) Nonselective NSAIDs Acetaminophen Calcium channel α -2- δ antagonists (e.g.	268 267 267	35.8 26.6 41.9	47.4* 57.3* 44.2*	14.2 12.7 12.4	1.9 2.6 1.5	0.7 0.8 0.0
gabapentin, pregabalin) 16. Unless contraindicated, all patients should receive an around-the-clock regimen of	265	15.1	38.5*	38.5	6.8	1.1
NSAIDs, COXIBs, or acetaminophen 17. Regional blockade with local anesthetics	264	24.2	34.1*	25.0	14.4	2.3
should be considered as part of a multimodal approach for pain managemen 18. Dosing regimens should be administered to optimize efficacy while minimizing the	t 264	58.3*	37.1	2.7	1.1	0.8
risk of adverse events 19. The choice of medication, dose, route, and duration of therapy should be	264 d	71.2*	27.3	1.1	0.4	0.0
individualized VI. <i>Patient Subpopulations Pediatric patients</i> 20. Perioperative care for children undergoing painful procedures or surgery requires	266	70.7*	27.1	1.1	1.1	0.0
developmentally appropriate pain assessment and therapy 21. Analgesic therapy should depend upon age, weight, and comorbidity and unless contraindicated should involve a	265	63.4*	35.1	1.5	0.0	0.0
multimodal approach 22. Behavioral techniques, especially importar in addressing the emotional component of		58.6*	34.7	4.5	2.2	0.0
pain, should be applied whenever feasible 23. Because many analgesic medications are synergistic with sedating agents, it is imperative that appropriate monitoring be		34.2	42.5*	21.4	1.5	0.4
used during the procedure and recovery Geriatric Patients 24. Pain assessment and therapy should be	268	69.4*	30.2	0.4	0.0	0.0
integrated into the perioperative care of geriatric patients	268	60.1*	37.7	1.8	0.4	0.0
25. Pain assessment tools appropriate to a patient's cognitive abilities should be used26. Extensive and proactive evaluation and questioning should be conducted to overcome barriers that hinder	I 268	58.6*	39.9	1.1	0.4	0.0
communication regarding unrelieved pain	265	35.9	41.1*	20.0	3.0	0.0 (continued)

Table 3. Continued

	Percent Responding to Each Item								
	N	Strongly Agree	Agree	Equivocal	Disagree	Strongly Disagree			
27. Dose titration should be done to ensure adequate treatment while avoiding adverse effects such as somnolence in this vulnerable group, who may be taking other medications	269	59.5*	39.7	0.4	0.4	0.0			
Other Subpopulations 28. Anesthesiologists should recognize that patients who are critically ill, cognitively impaired, or have communication difficulties may require additional interventions to ensure optimal perioperative pain management	268	53.0*	43.3	2.6	1.1	0.0			
29. Anesthesiologists should consider a therapeutic trial of an analgesic in patients with elevated blood pressure and heart rate or agitated behavior when causes other than pain have been excluded	267	32.6	56.5*	9.4	1.1	0.4			

^{*} Indicates the median.

COX-2 = cyclooxygenase-2; N = number of consultants who responded to each item; NSAID = nonsteroidal antiinflammatory drug; PCA = patient-controlled analgesia.

References

- Coleman SA, Booker-Milburn J: Audit of postoperative pain control: Influence of a dedicated acute pain nurse. Anaesthesia 1996; 51:1093-6
- 2. Harmer M, Davies KA: The effect of education, assessment and a standardised prescription on postoperative pain management. The value of clinical audit in the establishment of acute pain services. Anaesthesia 1998; 53:424-30
- Rose DK, Cohen MM, Yee DA: Changing the practice of pain management. Anesth Analg 1997; 84:764-72
- 4. White CL: Changing pain management practice and impacting on patient outcomes. Clin Nurse Spec 1999; 13:166-72
- Briggs M, Dean KL: A qualitative analysis of the nursing documentation of post-operative pain management. J Clin Nurs 1998; 7:155-63
- Camp LD, O'Sullivan PS: Comparison of medical, surgical and oncology patients' descriptions of pain and nurses' documentation of pain assessments. J Adv Nurs 1987; 12:593–8
- 7. Clarke EB, French B, Bilodeau ML, Capasso VC, Edwards A, Empoliti J: Pain management knowledge, attitudes and clinical practice: The impact of nurses' characteristics and education. J Pain Symptom Manage 1996; 11:18-31
- 8. Davis BD, Billings JR, Ryland RK: Evaluation of nursing process documentation. J Adv Nurs 1994; 19:960-8
- Ehnfors M, Smedby B: Nursing care as documented in patient records. Scand J Caring Sci 1993; 7:209-20
- Idvall E, Ehrenberg A: Nursing documentation of postoperative pain management. J Clin Nurs 2002; 11:734-42
- Salanter ä S, Lauri S, Salmi TT, Aantaa R: Nursing activities and outcomes of care in the assessment, management, and documentation of children's pain. J Pediatr Nurs 1999; 14:408-15
- 12. Bardiau FM, Taviaux NF, Albert A, Boogaerts JG, Stadler M: An intervention study to enhance postoperative pain management. Anesth Analg 2003; 96:179-85
- 13. Gould TH, Crosby DL, Harmer M, Lloyd SM, Lunn JN, Rees

- GA, Roberts DE, Webster JA: Policy for controlling pain after surgery: Effect of sequential changes in management. BMJ 1992; 305:1187-93
- Mackintosh C, Bowles S: Evaluation of a nurse-led acute pain service. Can clinical nurse specialists make a difference? J Adv Nurs 1997; 25:30-7
- Miaskowski C, Crews J, Ready LB, Paul SM, Ginsberg B: Anesthesia-based pain services improve the quality of postoperative pain management. Pain 1999; 80:23-9
- Pesut B, Johnson J: Evaluation of an acute pain service. Can J Nurs Adm 1997; 10:86-107
- Sartain JB, Barry JJ: The impact of an acute pain service on postoperative pain management. Anaesth Intensive Care 1999; 27:375-80
- Stacey BR, Rudy TE, Nelhaus D: Management of patientcontrolled analgesia: A comparison of primary surgeons and a dedicated pain service. Anesth Analg 1997; 85:130-4
- Stadler M, Schlander M, Braeckman M, Nguyen T, Boogaerts JG: A cost-utility and cost-effectiveness analysis of an acute pain service. J Clin Anesth 2004; 16:159-67
- Tighe SQ, Bie JA, Nelson RA, Skues MA: The acute pain service: Effective or expensive care? Anaesthesia 1998; 53: 397-403
- 21. Furdon SA, Eastman M, Benjamin K, Horgan MJ: Outcome measures after standardized pain management strategies in postoperative patients in the neonatal intensive care unit. J Perinat Neonatal Nurs 1998; 12:58-69
- Anderson EA: Preoperative preparation for cardiac surgery facilitates recovery, reduces psychological distress, and reduces the incidence of acute postoperative hypertension. J Consult Clin Psychol 1987; 55:513-20
- 23. Daltroy LH, Morlino CI, Eaton HM, Poss R, Liang MH: Preoperative education for total hip and knee replacement patients. Arthritis Care Res 1998; 11:469-78
- 24. Doering S, Katzlberger F, Rumpold G, Roessler S, Hofstoetter B, Schatz DS, Behensky H, Krismer M, Luz G, Innerhofer

- P, Benzer H, Saria A, Schuessler G: Videotape preparation of patients before hip replacement surgery reduces stress. Psychosom Med 2000; 62:365-73
- Egbert LD, Battit GE, Welch CE, Bartlett MK: Reduction of postoperative pain by encouragement and instruction of patients. N Engl J Med 1964; 270:825-7
- Elsass P, Eikard B, Junge J, Lykke J, Staun P, Feldt-Rasmussen M: Psychological effect of detailed preanesthetic information. Acta Anaesth Scand 1987; 31:579-83
- 27. Fortin F, Kirouac S: A randomized controlled trial of preoperative patient education. Int J Nurs Stud 1976; 13:11-24
- 28. Griffin MJ, Brennan L, McShane AJ: Preoperative education and outcome of patient controlled analgesia. Can J Anaesth 1998; 45:943-8
- 29. Knoerl DV, Faut-Callahan M, Paice J, Shott S: Preoperative PCA teaching program to manage postoperative pain. Medsurg Nurs 1999; 8:25-33
- Lam KK, Chan MT, Chen PP, Ngan Kee WD: Structured preoperative patient education for patient-controlled analgesia. J Clin Anesth 2001; 13:465-9
- 31. Lilja Y, Rydén S, Fridlund B: Effects of extended preoperative information on perioperative stress: An anaesthetic nurse intervention for patients with breast cancer and total hip replacement. Intensive Crit Care Nurs 1998; 14:276-82
- 32. McDonald DD, Freeland M, Thomas G, Moore J: Testing a preoperative pain management intervention for elders. Res Nurs Health 2001; 24:402-9
- Shuldham CM, Fleming S, Goodman H: The impact of preoperative education on recovery following coronary artery bypass surgery. A randomized controlled clinical trial. Eur Heart J 2002; 23:666-74
- 34. Watt-Watson J, Stevens B, Costello J, Katz J, Reid G: Impact of preoperative education on pain management outcomes after coronary artery bypass graft surgery: A pilot. Can J Nurs Res 2000; 31:41–56
- 35. Wilson JF: Behavioral preparation for surgery: Benefit or harm? J Behav Med 1981; 4:79-102
- 36. Banning AM, Schmidt JF, Chraemmer-Jørgensen B, Risbo A: Comparison of oral controlled release morphine and epidural morphine in the management of postoperative pain. Anesth Analg 1986; 65:385-8
- 37. Fitzpatrick GJ, Moriarty DC: Intrathecal morphine in the management of pain following cardiac surgery. A comparison with morphine I.V. Br J Anaesth 1988; 60:639-44
- 38. Tsui SL, Chan CS, Chan AS, Wong SJ, Lam CS, Jones RD: Postoperative analgesia for oesophageal surgery: A comparison of three analgesic regimens. Anaesth Intensive Care 1991; 19:329-37
- 39. Abboud TK, Dror A, Mosaad P, Zhu J, Mantilla M, Swart F, Gangolly J, Silao P, Makar A, Moore J, Davis H, Lee J: Mini-dose intrathecal morphine for the relief of post-cesarean section pain: Safety, efficacy, and ventilatory responses to carbon dioxide. Anesth Analg 1988; 67:137-43
- 40. Gall O, Aubineau JV, Bernière J, Desjeux L, Murat I: Analgesic effect of low-dose intrathecal morphine after spinal fusion in children. Anesthesiology 2001; 94:447-52
- Vanstrum GS, Bjornson KM, Ilko R: Postoperative effects of intrathecal morphine in coronary artery bypass surgery. Anesth Analg 1988; 67:261-7
- Broekema AA, Kuizenga K, Hennis PJ: Does epidural sufentanil provide effective analgesia per- and postoperatively for abdominal aortic surgery? Acta Anaesthesiol Scand 1996; 40:20-5
- 43. Kundra P, Gurnani A, Bhattacharya A: Preemptive epidural morphine for postoperative pain relief after lumbar laminectomy. Anesth Analg 1997; 85:135-8
- 44. Subramaniam B, Pawar DK, Kashyap L: Pre-emptive analgesia with epidural morphine or morphine and bupivacaine. Anaesth Intensive Care 2000; 28:392-8

- 45. Katz J, Kavanagh BP, Sandler AN, Nierenberg H, Boylan JF, Friedlander M, Shaw BF: Preemptive analgesia. Clinical evidence of neuroplasticity contributing to postoperative pain. Anesthesiology 1992; 77:439-46
- Cullen ML, Staren ED, el-Ganzouri A, Logas WG, Ivankovich AD, Economou SG: Continuous epidural infusion for analgesia after major abdominal operations: A randomized, prospective, double-blind study. Surgery 1985; 98: 718-28
- 47. Jacobson L, Chabal C, Brody MC: A dose-response study of intrathecal morphine: Efficacy, duration, optimal dose, and side effects. Anesth Analg 1988; 67:1082-8
- 48. Kawana Y, Sato H, Shimada H, Fujita N, Ueda Y, Hayashi A, Araki Y: Epidural ketamine for postoperative pain relief after gynecologic operations: A double-blind study and comparison with epidural morphine. Anesth Analg 1987; 66:735-8
- Logas WG, el-Baz N, el-Ganzouri A, Cullen M, Staren E, Faber LP, Ivankovich AD: Continuous thoracic epidural analgesia for postoperative pain relief following thoracotomy: A randomized prospective study. Anesthesiology 1987; 67:787-91
- Ross DA, Drasner K, Weinstein PR, Flaherty JF, Barbaro NM: Use of intrathecally administered morphine in the treatment of postoperative pain after lumbar spinal surgery: A prospective, double-blind, placebo-controlled study. Neurosurgery 1991; 28:700-4
- 51. Sarma VJ, Boström UV: Intrathecal morphine for the relief of post-hysterectomy pain-a double-blind, dose-response study. Acta Anaesth Scand 1993; 37:223-7
- Waikakul W, Chumniprasas K: Direct epidural morphine injection during lumbar discectomy for postoperative analgesia. J Med Assoc Thai 1992; 75:428-33
- 53. Writer WD, Hurtig JB, Evans D, Needs RE, Hope CE, Forrest JB: Epidural morphine prophylaxis of postoperative pain: Report of a double-blind multicentre study. Can Anaesth Soc J 1985; 32:330-8
- 54. Yamaguchi H, Watanabe S, Harukuni I, Hamaya Y: Effective doses of epidural morphine for relief of postcholecystectomy pain. Anesth Analg 1991; 72:80-3
- Bourke DL, Spatz E, Motara R, Ordia JI, Reed J, Hlavacek JM: Epidural opioids during laminectomy surgery for postoperative pain. J Clin Anesth 1992; 4:277-81
- Harrison DM, Sinatra R, Morgese L, Chung JH: Epidural narcotic and patient-controlled analgesia for post-cesarean section pain relief. Anesthesiology 1988; 68:454-7
- Klinck JR, Lindop MJ: Epidural morphine in the elderly. A controlled trial after upper abdominal surgery. Anaesthesia 1982; 37:907-12
- 58. Thind GS, Wells JC, Wilkes RG: The effects of continuous intravenous naloxone on epidural morphine analgesia. Anaesthesia 1986; 41:582-5
- 59. Youngstrom PC, Cowan RI, Sutheimer C, Eastwood DW, Yu JC: Pain relief and plasma concentrations from epidural and intramuscular morphine in post-cesarean patients. Anesthesiology 1982; 57:404-9
- Chan JH, Heilpern GN, Packham I, Trehan RK, Marsh GD, Knibb AA: A prospective randomized double-blind trial of the use of intrathecal fentanyl in patients undergoing lumbar spinal surgery. Spine 2006; 31:2529-33
- 61. Binsted RJ: Epidural morphine after caesarean section. Anaesth Intensive Care 1983; 11:130-4
- 62. Chabas E, Gomar C, Villalonga A, Sala X, Taura P: Postoperative respiratory function in children after abdominal surgery. A comparison of epidural and intramuscular morphine analgesia. Anaesthesia 1998; 53:393-7
- Chambers WA, Mowbray A, Wilson J: Extradural morphine for the relief of pain following caesarean section. Br J Anaesth 1983; 55:1201-3

- Daley MD, Sandler AN, Turner KE, Vosu H, Slavchenko P: A comparison of epidural and intramuscular morphine in patients following cesarean section. Anesthesiology 1990; 72: 289-94
- 65. Farag H, Naguib M: Caudal morphine for pain relief following anal surgery. Ann R Coll Surg Engl 1985; 67:257-8
- 66. Gustafsson LL, Friberg-Nielsen S, Garle M, Mohall A, Rane A, Schildt B, Symreng T: Extradural and parenteral morphine: Kinetics and effects in postoperative pain. A controlled clinical study. Br J Anaesth 1982; 54:1167-74
- 67. Rawal N, Sjöstrand U, Christoffersson E, Dahlström B, Arvill A, Rydman H: Comparison of intramuscular and epidural morphine for postoperative analgesia in the grossly obese: Influence on postoperative ambulation and pulmonary function. Anesth Analg 1984; 63:583–92
- 68. Reiz S, Ahlin J, Ahrenfeldt B, Andersson M, Andersson S: Epidural morphine for postoperative pain relief. Acta Anaesth Scand 1981; 25:111-4
- Rosen MA, Hughes SC, Shnider SM, Abboud TK, Norton M, Dailey PA, Curtis JD: Epidural morphine for the relief of postoperative pain after cesarean delivery. Anesth Analg 1983; 62:666-72
- Ibrahim AW, Farag H, Naguib M: Epidural morphine for pain relief after lumbar laminectomy. Spine 1986; 11: 1024-6
- Ellis DJ, Millar WL, Reisner LS: A randomized double-blind comparison of epidural *versus* intravenous fentanyl infusion for analgesia after cesarean section. Anesthesiology 1990; 72:981-6
- 72. Inagaki Y, Mashimo T, Yoshiya I: Segmental analgesic effect and reduction of halothane MAC from epidural fentanyl in humans. Anesth Analg 1992; 74:856-64
- Salomäki TE, Laitinen JO, Nuutinen LS: A randomized double-blind comparison of epidural *versus* intravenous fentanyl infusion for analgesia after thoracotomy. Anesthesiology 1991; 75:790-5
- 74. Sandler AN, Stringer D, Panos L, Badner N, Friedlander M, Koren G, Katz J, Klein J: A randomized, double-blind comparison of lumbar epidural and intravenous fentanyl infusions for postthoracotomy pain relief: Analgesic, pharmacokinetic, and respiratory effects. Anesthesiology 1992; 77: 626-34
- 75. Guinard JP, Mavrocordatos P, Chiolero R, Carpenter RL: A randomized comparison of intravenous *versus* lumbar and thoracic epidural fentanyl for analgesia after thoracotomy. Anesthesiology 1992; 77:1108-15
- 76. van Lersberghe C, Camu F, de Keersmaecker E, Sacré S: Continuous administration of fentanyl for postoperative pain: A comparison of the epidural, intravenous, and transdermal routes. J Clin Anesth 1994; 6:308-14
- 77. Boldt J, Thaler E, Lehmann A, Papsdorf M, Isgro F: Pain management in cardiac surgery patients: Comparison between standard therapy and patient-controlled analgesia regimen. J Cardiothorac Vasc Anesth 1998; 12:654-8
- Murphy DF, Graziotti P, Chalkiadis G, McKenna M: Patientcontrolled analgesia: A comparison with nurse-controlled intravenous opioid infusions. Anaesth Intensive Care 1994; 22:589-92
- Myles PS, Buckland MR, Cannon GB, Bujor MA, Langley M, Breaden A, Salamonsen RF, Davis BB: Comparison of patient-controlled analgesia and nurse-controlled infusion analgesia after cardiac surgery. Anaesth Intensive Care 1994; 22:672-8
- 80. O'Halloran P, Brown R: Patient-controlled analgesia compared with nurse-controlled infusion analgesia after heart surgery. Intensive Crit Care Nurs 1997; 13:126-9
- 81. Berde CB, Lehn BM, Yee JD, Sethna NF, Russo D: Patient-controlled analgesia in children and adolescents: A randomized, prospective comparison with intramuscular adminis-

- tration of morphine for postoperative analgesia. J Pediatr 1991; 118:460-6
- 82. Bollish SJ, Collins CL, Kirking DM, Bartlett RH: Efficacy of patient-controlled *versus* conventional analgesia for postoperative pain. Clin Pharm 1985; 4:48-52
- 83. Chan VW, Chung F, McQuestion M, Gomez M: Impact of patient-controlled analgesia on required nursing time and duration of postoperative recovery. Reg Anesth 1995; 20: 506-14
- 84. Choinière M, Rittenhouse BE, Perreault S, Chartrand D, Rousseau P, Smith B, Pepler C: Efficacy and costs of patient-controlled analgesia *versus* regularly administered intramuscular opioid therapy. Anesthesiology 1998; 89:1377–88
- Egbert AM, Parks LH, Short LM, Burnett ML: Randomized trial of postoperative patient-controlled analgesia vs intramuscular narcotics in frail elderly men. Arch Intern Med 1990; 150:1897-903
- 86. Passchier J, Rupreht J, Koenders ME, Olree M, Luitwieler RL, Bonke B: Patient-controlled analgesia (PCA) leads to more postoperative pain relief, but also to more fatigue and less vigour. Acta Anaesthesiol Scand 1993; 37:659-63
- 87. Sanansilp V, Lertakyamanee J, Udompunturak S: Cost-effectiveness analysis of patient-controlled analgesia, intramuscular q.i.d. injection and p.r.n. injection for postoperative pain relief. J Med Assoc Thai 1995; 78:600 4
- 88. Wheatley RG, Shepherd D, Jackson IJ, Madej TH, Hunter D: Hypoxaemia and pain relief after upper abdominal surgery: Comparison of i.m. and patient-controlled analgesia. Br J Anaesth 1992; 69:558-61
- 89. Grant RP, Dolman JF, Harper JA, White SA, Parsons DG, Evans KG, Merrick CP: Patient-controlled lumbar epidural fentanyl compared with patient-controlled intravenous fentanyl for post-thoracotomy pain. Can J Anaesth 1992; 39: 214-9
- 90. Ngan Kee WD, Lam KK, Chen PP, Gin T: Comparison of patient-controlled epidural analgesia with patient-controlled intravenous analgesia using pethidine or fentanyl. Anaesth Intensive Care 1997; 25:126-32
- Paech MJ, Moore JS, Evans SF: Meperidine for patient-controlled analgesia after cesarean section. Intravenous versus epidural administration. Anesthesiology 1994; 80: 1268-76
- 92. Stoddart PA, Cooper A, Russell R, Reynolds F: A comparison of epidural diamorphine with intravenous patient-controlled analgesia using the Baxter infusor following caesarean section. Anaesthesia 1993; 48:1086-90
- 93. Welchew EA, Breen DP: Patient-controlled on-demand epidural fentanyl: A comparison of patient-controlled on-demand fentanyl delivered epidurally or intravenously. Anaesthesia 1991; 46:438-41
- 94. Guler T, Unlugenc H, Gundogan Z, Ozalevli M, Balcioglu O, Topcuoglu MS: A background infusion of morphine enhances patient-controlled analgesia after cardiac surgery. Can J Anaesth 2004; 51:718-22
- Dal D, Kanbak M, Caglar M, Aypar U: A background infusion of morphine does not enhance postoperative analgesia after cardiac surgery. Can J Anaesth 2003; 50:476-9
- 96. Doyle E, Harper I, Morton NS: Patient-controlled analgesia with low dose background infusions after lower abdominal surgery in children. Br J Anaesth 1993; 71:818-22
- 97. Doyle E, Robinson D, Morton NS: Comparison of patient-controlled analgesia with and without a background infusion after lower abdominal surgery in children. Br J Anaesth 1993; 71:670-3
- 98. McNeely JK, Trentadue NC: Comparison of patient-controlled analgesia with and without nighttime morphine infusion following lower extremity surgery in children. J Pain Symptom Manage 1997; 13:268-73

- 99. Owen H, Szekely SM, Plummer JL, Cushnie JM, Mather LE: Variables of patient-controlled analgesia: 2. Concurrent infusion. Anaesthesia 1989; 44:11-3
- 100. Parker RK, Holtmann B, White PF: Effects of a nighttime opioid infusion with PCA therapy on patient comfort and analgesic requirements after abdominal hysterectomy. Anesthesiology 1992; 76:362-7
- 101. Russell AW, Owen H, Ilsley AH, Kluger MT, Plummer JL: Background infusion with patient-controlled analgesia: Effect on postoperative oxyhaemoglobin saturation and pain control. Anaesth Intensive Care 1993; 21:174-9
- 102. Sinatra R, Chung KS, Silverman DG, Brull SJ, Chung J, Harrison DM, Donielson D, Weinstock A: An evaluation of morphine and oxymorphone administered *via* patient-controlled analgesia (PCA) or PCA plus basal infusion in postcesarean-delivery patients. ANESTHESIOLOGY 1989; 71:502-7
- 103. Smythe MA, MB Zak, O'Donnell MP, Schad RF, Dmuchowski CF: Patient-controlled analgesia versus patient-controlled analgesia plus continuous infusion after hip replacement surgery. Ann Pharmacother 1996; 30:224-7
- 104. Eng J, Sabanathan S: Continuous extrapleural intercostal nerve block and post-thoracotomy pulmonary complications. Scand J Thorac Cardiovasc Surg 1992; 26:219-23
- 105. Rademaker BM, Sih IL, Kalkman CJ, Henny CP, Filedt Kok JC, Endert E, Zuurmond WW: Effects of interpleurally administered bupivacaine 0.5% on opioid analgesic requirements and endocrine response during and after cholecystectomy: A randomized double-blind controlled study. Acta Anaesth Scand 1991; 35:108-12
- 106. Barron DJ, Tolan MJ, Lea RE: A randomized controlled trial of continuous extra-pleural analgesia post-thoracotomy: Efficacy and choice of local anaesthetic. Eur J Anaesthesiol 1999; 16:236-45
- 107. Deneuville M, Bisserier A, Regnard JF, Chevalier M, Levasseur P, Hervé P: Continuous intercostal analgesia with 0.5% bupivacaine after thoracotomy: A randomized study. Ann Thorac Surg 1993; 55:381-5
- 108. Knowles P, Hancox D, Letheren M, Eddleston J: An evaluation of intercostal nerve blockade for analgesia following renal transplantation. Eur J Anaesthesiol 1998; 15:457-61
- Lee A, Boon D, Bagshaw P, Kempthorne P: A randomised double-blind study of interpleural analgesia after cholecystectomy. Anaesthesia 1990; 45:1028-31
- 110. Chan VW, Chung F, Cheng DC, Seyone C, Chung A, Kirby TJ: Analgesic and pulmonary effects of continuous intercostal nerve block following thoracotomy. Can J Anaesth 1991; 38:733-9
- Dryden CM, McMenemin I, Duthie DJ: Efficacy of continuous intercostal bupivacaine for pain relief after thoracotomy. Br J Anaesth 1993; 70:508-10
- 112. Mann LJ, Young GR, Williams JK, Dent OF, McCaughan BC: Intrapleural bupivacaine in the control of postthoracotomy pain. Ann Thorac Surg 1992; 53:449-54
- 113. Mozell EJ, Sabanathan S, Mearns AJ, Bickford-Smith PJ, Majid MR, Zografos G: Continuous extrapleural intercostal nerve block after pleurectomy. Thorax 1991; 46:21-4
- 114. Sabanathan S, Mearns AJ, Bickford Smith PJ, Eng J, Berrisford RG, Bibby SR, Majid MR: Efficacy of continuous extrapleural intercostal nerve block on post-thoracotomy pain and pulmonary mechanics. Br J Surg 1990; 77:221-5
- 115. Schneider RF, Villamena PC, Harvey J, Surick BG, Surick IW, Beattie EJ: Lack of efficacy of intrapleural bupivacaine for postoperative analgesia following thoracotomy. Chest 1993; 103:414-6
- 116. Symreng T, Gomez MN, Rossi N: Intrapleural bupivacaine versus saline after thoracotomy: Effects on pain and lung function: A double-blind study. J Cardiothorac Anesth 1989; 3:144-9

- 117. VadeBoncouer TR, Riegler FX, Gautt RS, Weinberg GL: A randomized, double-blind comparison of the effects of interpleural bupivacaine and saline on morphine requirements and pulmonary function after cholecystectomy. ANESTHESIOLOGY 1989; 71:339 - 43
- 118. Al-Kaisy A, McGuire G, Chan VW, Bruin G, Peng P, Miniaci A, Perlas A: Analgesic effect of interscalene block using low-dose bupivacaine for outpatient arthroscopic shoulder surgery. Reg Anesth Pain Med 1998; 23:469-73
- 119. Aunac S, Carlier M, Singelyn F, De Kock M: The analgesic efficacy of bilateral combined superficial and deep cervical plexus block administered before thyroid surgery under general anesthesia. Anesth Analg 2002; 95:746-50
- 120. Ding Y, White PF: Post-herniorrhaphy pain in outpatients after pre-incision ilioinguinal-hypogastric nerve block during monitored anaesthesia care. Can J Anaesth 1995; 42: 12-5
- 121. Toivonen J, Permi J, Rosenberg PH: Effect of preincisional ilioinguinal and iliohypogastric nerve block on postoperative analgesic requirement in day-surgery patients undergoing herniorrhaphy under spinal anaesthesia. Acta Anaesthesiol Scand 2001; 45:603–7
- 122. Langer JC, Shandling B, Rosenberg M: Intraoperative bupivacaine during outpatient hernia repair in children: A randomized double blind trial. J Pediatr Surg 1987; 22:267-70
- 123. Brunat G, Pouzeratte Y, Mann C, Didelot JM, Rochon JC, Eledjam JJ: Posterior perineal block with ropivacaine 0.75% for pain control during and after hemorrhoidectomy. Reg Anesth Pain Med 2003; 28:228-32
- 124. Huffnagle HJ, Norris MC, Leighton BL, Arkoosh VA: Ilioinguinal iliohypogastric nerve blocks-before or after cesarean delivery under spinal anesthesia? Anesth Analg 1996; 82:8-12
- 125. Stevens RD, Van Gessel E, Flory N, Fournier R, Gamulin Z: Lumbar plexus block reduces pain and blood loss associated with total hip arthroplasty. Anesthesiology 2000; 93: 115-21
- 126. Singelyn FJ, Lhotel L, Fabre B: Pain relief after arthroscopic shoulder surgery: A comparison of intraarticular analgesia, suprascapular nerve block, and interscalene brachial plexus block. Anesth Analg 2004; 99:589-92
- Tree-Trakarn T, Pirayavaraporn S: Postoperative pain relief for circumcision in children: Comparison among morphine, nerve block, and topical analgesia. Anesthesiology 1985; 62:519-22
- 128. Bogoch ER, Henke M, Mackenzie T, Olschewski E, Mahomed NN: Lumbar paravertebral nerve block in the management of pain after total hip and knee arthroplasty: A randomized controlled clinical trial. J Arthroplasty 2002; 17:398-401
- 129. Dieudonne N, Gomola A, Bonnichon P, Ozier YM: Prevention of postoperative pain after thyroid surgery: A double-blind randomized study of bilateral superficial cervical plexus blocks. Anesth Analg 2001; 92:1538-42
- 130. Fredman B, Zohar E, Ganim T, Shalev M, Jedeikin R: Bupivacaine infiltration into the neurovascular bundle of the prostatic nerve does not improve postoperative pain or recovery following transvesical prostatectomy. J Urol 1998; 159:154-6
- 131. Hinkle AJ: Percutaneous inguinal block for the outpatient management of post-herniorrhaphy pain in children. ANESTHESIOLOGY 1987; 67:411-3
- 132. McLoughlin J, Kelley CJ: Study of the effectiveness of bupivicaine infiltration of the ilioinguinal nerve at the time of hernia repair for post-operative pain relief. Br J Clin Pract 1989; 43:281-3
- 133. Brandsson S, Karlsson J, Morberg P, Rydgren B, Eriksson BI, Hedner T: Intraarticular morphine after arthroscopic ACL reconstruction: A double-blind placebo-controlled study of 40 patients. Acta Orthop Scand 2000; 71:280-5

- 134. Heard SO, Edwards WT, Ferrari D, Hanna D, Wong PD, Liland A, Willock MM: Analgesic effects of intraarticular bupivacaine or morphine after arthroscopic knee surgery: A randomized, prospective, double-blind study. Anesth Analg 1992; 74:822-6
- 135. Kanbak M, Akpolat N, Ocal T, Doral MN, Ercan M, Erdem K: Intraarticular morphine administration provides pain relief after knee arthroscopy. Eur J Anaesthesiol 1997; 14:153-6
- 136. Raja SN, Dickstein RE, Johnson CA: Comparison of postoperative analgesic effects of intraarticular bupivacaine and morphine following arthroscopic knee surgery. Anesthesiology 1992; 77:1143-7
- 137. Rosseland LA, Stubhaug A, Skoglund A, Breivik H: Intraarticular morphine for pain relief after knee arthroscopy. Acta Anaesth Scand 1999; 43:252-7
- 138. Chirwa SS, MacLeod BA, Day B: Intraarticular bupivacaine (Marcaine) after arthroscopic meniscectomy: A randomized double-blind controlled study. Arthroscopy 1989; 5:33-5
- 139. Henderson RC, Campion ER, DeMasi RA, Taft TN: Postarthroscopy analgesia with bupivacaine: A prospective, randomized blinded evaluation. Am J Sports Med 1990; 18: 614-7
- 140. Fong SY, Pavy TJ, Yeo ST, Paech MJ, Gurrin LC: Assessment of wound infiltration with bupivacaine in women undergoing day-case gynecological laparoscopy. Reg Anesth Pain Med 2001; 26:131-6
- 141. Goldsher M, Podoshin L, Fradis M, Malatskey S, Gerstel R, Vaida S, Gaitini L: Effects of peritonsillar infiltration on post-tonsillectomy pain. A double-blind study. Ann Otol Rhinol Laryngol 1996; 105:868-70
- 142. Jebeles JA, Reilly JS, Gutierrez JF, Bradley EL Jr, Kissin I: Tonsillectomy and adenoidectomy pain reduction by local bupivacaine infiltration in children. Int J Pediatr Otorhinolaryngol 1993; 25:149-54
- 143. Jebeles JA, Reilly JS, Gutierrez JF, Bradley EL Jr, Kissin I: The effect of pre-incisional infiltration of tonsils with bupivacaine on the pain following tonsillectomy under general anesthesia. Pain 1991; 47:305-8
- 144. Johansen M, Harbo G, Illum P: Preincisional infiltration with bupivacaine in tonsillectomy. Arch Otolaryngol Head Neck Surg 1996; 122:261-3
- 145. Ke RW, Portera SG, Bagous W, Lincoln SR: A randomized, double-blinded trial of preemptive analgesia in laparoscopy. Obstet Gynecol 1998; 92:972-5
- 146. Marsh GD, Huddy SP, Rutter KP: Bupivacaine infiltration after haemorrhoidectomy. J R Coll Surg Edinb 1993; 38: 41-2
- 147. Molliex S, Haond P, Baylot D, Prades JM, Navez M, Elkhoury Z, Auboyer C: Effect of pre- vs postoperative tonsillar infiltration with local anesthetics on postoperative pain after tonsillectomy. Acta Anaesthesiol Scand 1996; 40:1210-5
- 148. Vasan NR, Stevenson S, Ward M: Preincisional bupivacaine in posttonsillectomy pain relief: A randomized prospective study. Arch Otolaryngol Head Neck Surg 2002; 128:145-9
- 149. Eriksson-Mjöberg M, Kristiansson M, Carlström K, Eklund J, Gustafsson LL, Olund A: Preoperative infiltration of bupivacaineeffects on pain relief and trauma response (cortisol and interleukin-6). Acta Anaesthesiol Scand 1997; 41:466-72
- 150. Hannibal K, Galatius H, Hansen A, Obel E, Ejlersen E: Preoperative wound infiltration with bupivacaine reduces early and late opioid requirement after hysterectomy. Anesth Analg 1996; 83:376-81
- 151. Christie JM, Chen GW: Secondary hyperalgesia is not affected by wound infiltration with bupivacaine. Can J Anaesth 1993; 40:1034-7
- 152. Dierking GW, Ostergaard E, Ostergård HT, Dahl JB: The effects of wound infiltration with bupivacaine *versus* saline on postoperative pain and opioid requirements after herniorrhaphy. Acta Anaesthesiol Scand 1994; 38:289-92

- 153. Klein JR, Heaton JP, Thompson JP, Cotton BR, Davidson AC, Smith G: Infiltration of the abdominal wall with local anaesthetic after total abdominal hysterectomy has no opioidsparing effect. Br J Anaesth 2000; 84:248-9
- 154. Kountakis SE: Effectiveness of perioperative bupivacaine infiltration in tonsillectomy patients. Am J Otolaryngol 2002; 23:76-80
- 155. Owen H, Galloway DJ, Mitchell KG: Analgesia by wound infiltration after surgical excision of benign breast lumps. Ann R Coll Surg Engl 1985; 67:114-5
- 156. Partridge BL, Stabile BE: The effects of incisional bupivacaine on postoperative narcotic requirements, oxygen saturation and length of stay in the post-anesthesia care unit. Acta Anaesth Scand 1990; 34:486-91
- 157. Patel JM, Lanzafame RJ, Williams JS, Mullen BV, Hinshaw JR: The effect of incisional infiltration of bupivacaine hydrochloride upon pulmonary functions, atelectasis and narcotic need following elective cholecystectomy. Surg Gynecol Obstet 1983; 157:338-40
- 158. Russell WC, Ramsay AH, Fletcher DR: The effect of incisional infiltration of bupivacaine upon pain and respiratory function following open cholecystectomy. Aust N Z J Surg 1993: 63:756-9
- 159. Trotter TN, Hayes-Gregson P, Robinson S, Cole L, Coley S, Fell D: Wound infiltration of local anaesthetic after lower segment caesarean section. Anaesthesia 1991; 46:404-7
- 160. Wright JE: Controlled trial of wound infiltration with bupivacaine for postoperative pain relief after appendicectomy in children. Br J Surg 1993; 80:110-1
- 161. Bourget JL, Clark J, Joy N: Comparing preincisional with postincisional bupivacaine infiltration in the management of postoperative pain. Arch Surg 1997; 132:766-9
- 162. Cnar SO, Kum U, Cevizci N, Kayaoglu S, Oba S: Effects of levobupivacaine infiltration on postoperative analgesia and stress response in children following inguinal hernia repair. Eur J Anaesthesiol 2009; 26:430-4
- 163. Dahl V, Raeder JC, Ernø PE, Kovdal A: Pre-emptive effect of pre-incisional versus post-incisional infiltration of local anaesthesia on children undergoing hernioplasty. Acta Anaesthesiol Scand 1996; 40:847-51
- 164. O'Hanlon DM, Colbert ST, Keane PW, Given FH: Preemptive bupivacaine offers no advantages to postoperative wound infiltration in analgesia for outpatient breast biopsy. Am J Surg 2000; 180:29-32
- 165. Gemma M, Piccioni LO, Gioia L, Beretta L, Bussi M: Ropivacaine peritonsillar infiltration for analgesia after adenotonsillectomy in children: A randomized, double-blind, placebo-controlled study. Ann Otol Rhinol Laryngol 2009; 118: 227-31
- 166. Giannoni C, White S, Enneking FK, Morey T: Ropivacaine with or without clonidine improves pediatric tonsillectomy pain. Arch Otolaryngol Head Neck Surg 2001; 127:1265-70
- Johansson A, Axelson J, Ingvar C, Luttropp H-H, Lundberg J: Preoperative ropivacaine infiltration in breast surgery. Acta Anaesthesiol Scand 2000; 44:1093-8
- 168. Johansson B, Hallerbäck B, Stubberöd A, Janbu T, Edwin B, Glise H, Solhaug JH: Preoperative local infiltration with ropivacaine for postoperative pain relief after inguinal hernia repair. A randomised controlled trial. Eur J Surg 1997; 163:371-8
- 169. Kato J, Ogawa S, Katz J, Nagai H, Kashiwazaki M, Saeki H, Suzuki H: Effects of presurgical local infiltration of bupivacaine in the surgical field on postsurgical wound pain in laparoscopic gynecologic examinations: A possible preemptive analgesic effect. Clin J Pain 2000; 16:12-7
- 170. Papaziogas B, Argiriadou H, Papagiannopoulou P, Pavlidis T, Georgiou M, Sfyra E, Papaziogas T: Preincisional intravenous low-dose ketamine and local infiltration with ropivacaine reduces postoperative pain after laparoscopic cholecystectomy. Surg Endosc 2001; 15:1030-3

- 171. Vinson-Bonnet B, Coltat JC, Fingerhut A, Bonnet F: Local infiltration with ropivacaine improves immediate postoperative pain control after hemorrhoidal surgery. Dis Colon Rectum 2002; 45:104-8
- 172. Asantila R, Eklund P, Rosenberg PH: Continuous epidural infusion of bupivacaine and morphine for postoperative analgesia after hysterectomy. Acta Anaesth Scand 1991; 35:513-7
- 173. Crews JC, Hord AH, Denson DD, Schatzman C: A comparison of the analgesic efficacy of 0.25% levobupivacaine combined with 0.005% morphine, 0.25% levobupivacaine alone, or 0.005% morphine alone for the management of postoperative pain in patients undergoing major abdominal surgery. Anesth Analg 1999; 89:1504-9
- 174. Douglas MJ, McMorland GH, Janzen JA: Influence of bupivacaine as an adjuvant to epidural morphine for analgesia after cesarean section. Anesth Analg 1988; 67:1138-41
- 175. Hesselgard K, Strömblad LG, Reinstrup P: Morphine with or without a local anaesthetic for postoperative intrathecal pain treatment after selective dorsal rhizotomy in children. Paediatr Anaesth 2001; 11:75-9
- 176. Liu SS, Carpenter RL, Mackey DC, Thirlby RC, Rupp SM, Shine TS, Feinglass NG, Metzger PP, Fulmer JT, Smith SL: Effects of perioperative analgesic technique on rate of recovery after colon surgery. Anesthesiology 1995; 83: 757-65
- 177. Benzon HT, Wong CA, Wong HY, Brooke C, Wade L: The effect of low-dose bupivacaine on postoperative epidural fentanyl analgesia and thrombelastography. Anesth Analg 1994; 79:911-7
- 178. Cohen S, Lowenwirt I, Pantuck CB, Amar D, Pantuck EJ: Bupivacaine 0.01% and/or epinephrine 0.5 microg/ml improve epidural fentanyl analgesia after cesarean section. Anesthesiology 1998; 89:1354-61
- 179. Cooper DW, Ryall DM, McHardy FE, Lindsay SL, Eldabe SS: Patient-controlled extradural analgesia with bupivacaine, fentanyl, or a mixture of both, after Caesarean section. Br J Anaesth 1996; 76:611-5
- 180. Cooper DW, Turner G: Patient-controlled extradural analgesia to compare bupivacaine, fentanyl and bupivacaine with fentanyl in the treatment of postoperative pain. Br J Anaesth 1993; 70:503-7
- 181. George KA, Chisakuta AM, Gamble JA, Browne GA: Thoracic epidural infusion for postoperative pain relief following abdominal aortic surgery: Bupivacaine, fentanyl or a mixture of both? Anaesthesia 1992; 47:388-94
- 182. George KA, Wright PM, Chisakuta A: Continuous thoracic epidural fentanyl for post-thoracotomy pain relief: With or without bupivacaine? Anaesthesia 1991; 46:732-6
- 183. Kostamovaara PA, Laurila JJ, Alahuhta S, Salomåki TE: Ropivacaine 1 mg x ml(-1) does not decrease the need for epidural fentanyl after hip replacement surgery. Acta Anaesthesiol Scand 2001; 45:489-94
- 184. Mahon SV, Berry PD, Jackson M, Russell GN, Pennefather SH: Thoracic epidural infusions for post-thoracotomy pain: A comparison of fentanyl-bupivacaine mixtures *vs.* fentanyl alone. Anaesthesia 1999; 54:641-6
- 185. Paech MJ, Westmore MD: Postoperative epidural fentanyl infusion-is the addition of 0.1% bupivacaine of benefit? Anaesth Intensive Care 1994; 22:9-14
- 186. Reinoso-Barbero F, Saavedra B, Hervilla S, de Vicente J, Tabarés B, Gómez-Criado MS: Lidocaine with fentanyl, compared to morphine, marginally improves postoperative epidural analgesia in children. Can J Anaesth 2002; 49:67-71
- 187. Salomäki TE, Laitinen JO, Vainionpää V, Nuutinen LS: 0.1% bupivacaine does not reduce the requirement for epidural fentanyl infusion after major abdominal surgery. Reg Anesth 1995; 20:435-43
- 188. Torda TA, Hann P, Mills G, De Leon G, Penman D: Comparison of extradural fentanyl, bupivacaine and two fenta-

- nyl-bupivacaine mixtures of pain relief after abdominal surgery. Br J Anaesth 1995; 74:35-40
- 189. Cullen ML, Staren ED, El-Ganzouri A, Logas WG, Ivankovich AD, Economou SG: Continuous epidural infusion for analgesia after major abdominal operations: A randomized, prospective, double-blind study. Surgery 1985; 98:718-28
- 190. Jørgensen H, Fomsgaard JS, Dirks J, Wetterslev J, Andreasson B, Dahl JB: Effect of epidural bupivacaine vs combined epidural bupivacaine and morphine on gastrointestinal function and pain after major gynaecological surgery. Br J Anaesth 2001; 87:727-32
- 191. Martin LV: Postoperative analgesia after circumcision in children. Br J Anaesth 1982; 54:1263-6
- 192. Scott NB, Mogensen T, Bigler D, Lund C, Kehlet H: Continuous thoracic extradural 0.5% bupivacaine with or without morphine: Effect on quality of blockade, lung function and the surgical stress response. Br J Anaesth 1989; 62:253-7
- 193. Wolf AR, Hughes D, Hobbs AJ, Prys-Roberts C: Combined morphine-bupivacaine caudals for reconstructive penile surgery in children: Systemic absorption of morphine and postoperative analgesia. Anaesth Intensive Care 1991; 19: 17-21
- 194. Wolf AR, Hughes D, Wade A, Mather SJ, Prys-Roberts C: Postoperative analgesia after paediatric orchidopexy: Evaluation of a bupivacaine-morphine mixture. Br J Anaesth 1990; 64:430-5
- 195. Campbell FA, Yentis SM, Fear DW, Bissonnette B: Analgesic efficacy and safety of a caudal bupivacaine-fentanyl mixture in children. Can J Anaesth 1992; 39:661-4
- 196. Lauretti GR, Mattos AL, Reis MP, Pereira NL: Combined intrathecal fentanyl and neostigmine: Therapy for postoperative abdominal hysterectomy pain relief. J Clin Anesth 1998; 10:291-6
- 197. Løvstad RZ, Støen R: Postoperative epidural analgesia in children after major orthopaedic surgery: A randomised study of the effect on PONV of two anaesthetic techniques: Low and high dose I.V. fentanyl and epidural infusions with and without fentanyl. Acta Anaesthesiol Scand 2001; 45: 482-8
- 198. Berti M, Casati A, Fanelli G, Albertin A, Palmisano S, Danelli G, Comotti L, Torri G: 0.2% ropivacaine with or without fentanyl for patient-controlled epidural analgesia after major abdominal surgery: A double-blind study. J Clin Anesth 2000; 12:292–7
- 199. Buggy DJ, Hall NA, Shah J, Brown J, Williams J: Motor block during patient-controlled epidural analgesia with ropivacaine or ropivacaine/fentanyl after intrathecal bupivacaine for caesarean section. Br J Anaesth 2000; 85: 468-70
- 200. Finucane BT, Ganapathy S, Carli F, Pridham JN, Ong BY, Shukla RC, Kristoffersson AH, Huizar KM, Nevin K, Ahlén KG, Canadian Ropivacaine Research Group: Prolonged epidural infusions of ropivacaine (2 mg/ml) after colonic surgery: The impact of adding fentanyl. Anesth Analg 2001; 92:1276-85
- 201. Scott DA, Blake D, Buckland M, Etches R, Halliwell R, Marsland C, Merridew G, Murphy D, Paech M, Schug SA, Turner G, Walker S, Huizar K, Gustafsson U: A comparison of epidural ropivacaine infusion alone and in combination with 1, 2, and 4 microg/ml fentanyl for seventy-two hours of postoperative analgesia after major abdominal surgery. Anesth Analg 1999; 88:857-64
- 202. Hübler M, Litz RJ, Sengebusch KH, Kreinecker I, Frank MD, Hakenberg OW, Albrecht DM: A comparison of five solutions of local anaesthetics and/or sufentanil for continuous, postoperative epidural analgesia after major urological surgery. Eur J Anaesthesiol 2001; 18:450-7
- 203. Kampe S, Weigand C, Kaufmann J, Klimek M, König DP,

- Lynch J: Postoperative analgesia with no motor block by continuous epidural infusion of ropivacaine 0.1% and sufentanil after total hip replacement. Anesth Analg 1999; 89: 395-8
- 204. Lorenzini C, Moreira LB, Ferreira MB: Efficacy of ropivacaine compared with ropivacaine plus sufentanil for postoperative analgesia after major knee surgery. Anaesthesia 2002; 57:424-8
- 205. Pouzeratte Y, Delay JM, Brunat G, Boccara G, Vergne C, Jaber S, Fabre JM, Colson P, Mann C: Patient-controlled epidural analgesia after abdominal surgery: Ropivacaine versus bupivacaine. Anesth Analg 2001; 93:1587-92
- 206. Wiebalck A, Brodner G, Van Aken H: The effects of adding sufentanil to bupivacaine for postoperative patient-controlled epidural analgesia. Anesth Analg 1997; 85:124-9
- Carabine UA, Milligan KR, Mulholland D, Moore J: Extradural clonidine infusions for analgesia after total hip replacement. Br J Anaesth 1992; 68:338-43
- 208. Motsch J, Gräber E, Ludwig K: Addition of clonidine enhances postoperative analgesia from epidural morphine: A double-blind study. Anesthesiology 1990; 73:1067-73
- 209. Rockemann MG, Seeling W, Brinkmann A, Goertz AW, Hauber N, Junge J, Georgieff M: Analgesic and hemodynamic effects of epidural clonidine, clonidine/morphine, and morphine after pancreatic surgery-a double-blind study. Anesth Analg 1995; 80:869-74
- 210. van Essen EJ, Bovill JG, Ploeger EJ: Extradural clonidine does not potentiate analgesia produced by extradural morphine after meniscectomy. Br J Anaesth 1991; 66:237-41
- 211. Vercauteren MP, Saldien V, Bosschaerts P, Adriaensen HA: Potentiation of sufentanil by clonidine in PCEA with or without basal infusion. Eur J Anaesthesiol 1996; 13:571-6
- 212. Vercauteren MP, Vandeput DM, Meert TF, Adriaensen HA: Patient-controlled epidural analgesia with sufentanil following caesarean section: The effect of adrenaline and clonidine admixture. Anaesthesia 1994; 49:767-71
- 213. Burns JW, Aitken HA, Bullingham RE, McArdle CS, Kenny GN: Double-blind comparison of the morphine sparing effect of continuous and intermittent I.M. administration of ketorolac. Br J Anaesth 1991; 67:235–8
- 214. Cataldo PA, Senagore AJ, Kilbride MJ: Ketorolac and patient controlled analgesia in the treatment of postoperative pain. Surg Gynecol Obstet 1993; 176:435-8
- 215. Munro HM, Walton SR, Malviya S, Merkel S, Voepel-Lewis T, Loder RT, Farley FA: Low-dose ketorolac improves analgesia and reduces morphine requirements following posterior spinal fusion in adolescents. Can J Anaesth 2002; 49:461-6
- 216. Reuben SS, Connelly NR, Lurie S, Klatt M, Gibson CS: Dose-response of ketorolac as an adjunct to patient-controlled analgesia morphine in patients after spinal fusion surgery. Anesth Analg 1998; 87:98-102
- 217. Reuben SS, Connelly NR, Steinberg R: Ketorolac as an adjunct to patient-controlled morphine in postoperative spine surgery patients. Reg Anesth 1997; 22:343-6
- 218. Sevarino FB, Sinatra RS, Paige D, Silverman DG: Intravenous ketorolac as an adjunct to patient-controlled analgesia (PCA) for management of postgynecologic surgical pain. J Clin Anesth 1994; 6:23-7
- 219. Sutters KA, Shaw BA, Gerardi JA, Hebert D: Comparison of morphine patient-controlled analgesia with and without ketorolac for postoperative analgesia in pediatric orthopedic surgery. Am J Orthop 1999; 28:351-8
- 220. Vetter TR, Heiner EJ: Intravenous ketorolac as an adjuvant to pediatric patient-controlled analgesia with morphine. J Clin Anesth 1994; 6:110-3
- 221. Adriaenssens G, Vermeyen KM, Hoffmann VL, Mertens E, Adriaensen HF: Postoperative analgesia with i.v. patient-

- controlled morphine: Effect of adding ketamine. Br J Anaesth 1999; 83:393-6
- 222. Edwards ND, Fletcher A, Cole JR, Peacock JE: Combined infusions of morphine and ketamine for postoperative pain in elderly patients. Anaesthesia 1993; 48:124-7
- 223. Javery KB, Ussery TW, Steger HG, Colclough GW: Comparison of morphine and morphine with ketamine for postoperative analgesia. Can J Anaesth 1996; 43:212-5
- 224. Michelet P, Guervilly C, Hélaine A, Avaro JP, Blayac D, Gaillat F, Dantin T, Thomas P, Kerbaul F: Adding ketamine to morphine for patient-controlled analgesia after thoracic surgery: Influence on morphine consumption, respiratory function, and nocturnal desaturation. Br J Anaesth 2007; 99:396-403
- 225. Reeves M, Lindholm DE, Myles PS, Fletcher H, Hunt JO: Adding ketamine to morphine for patient-controlled analgesia after major abdominal surgery: A double-blinded, randomized controlled trial. Anesth Analg 2001; 93:116–20
- 226. Sveticic G, Farzanegan F, Zmoos P, Zmoos S, Eichenberger U, Curatolo M: Is the combination of morphine with ketamine better than morphine alone for postoperative intravenous patient-controlled analgesia? Anesth Analg 2008; 106:287-93
- 227. Huang YM, Wang CM, Wang CT, Lin WP, Horng LC, Jiang CC: Perioperative celecoxib administration for pain management after total knee arthroplasty: A randomized, controlled study. BMC Musculoskelet Disord 2008; 9:77
- 228. Plummer JL, Owen H, Ilsley AH, Tordoff K: Sustained-release ibuprofen as an adjunct to morphine patient-controlled analgesia. Anesth Analg 1996; 83:92-6
- 229. Serpell MG, Thomson MF: Comparison of piroxicam with placebo in the management of pain after total hip replacement. Br J Anaesth 1989; 63:354-6
- 230. Schug SA, Sidebotham DA, McGuinnety M, Thomas J, Fox L: Acetaminophen as an adjunct to morphine by patient-controlled analgesia in the management of acute postoperative pain. Anesth Analg 1998; 87:368-72
- 231. Agarwal A, Gautam S, Gupta D, Agarwal S, Singh PK, Singh U: Evaluation of a single preoperative dose of pregabalin for attenuation of postoperative pain after laparoscopic cholecystectomy. Br J Anaesth 2008; 101:700 4
- 232. Al-Mujadi H, A-Refai AR, Katzarov MG, Dehrab NA, Batra YK, Al-Qattan AR: Preemptive gabapentin reduces postoperative pain and opioid demand following thyroid surgery. Can J Anaesth 2006; 53:268-73
- 233. Clarke H, Pereira S, Kennedy D, Gilron I, Katz J, Gollish J, Kay J: Gabapentin decreases morphine consumption and improves functional recovery following total knee arthroplasty. Pain Res Manag 2009; 14:217–22
- 234. Dirks J, Fredensborg BB, Christensen D, Fomsgaard JS, Flyger H, Dahl JB: A randomized study of the effects of single-dose gabapentin *versus* placebo on postoperative pain and morphine consumption after mastectomy. ANESTHESIOLOGY 2002; 97:560-4
- 235. Grover VK, Mathew PJ, Yaddanapudi S, Sehgal S: A single dose of preoperative gabapentin for pain reduction and requirement of morphine after total mastectomy and axillary dissection: Randomized placebo-controlled doubleblind trial. J Postgrad Med 2009; 55:257-60
- 236. Pandey CK, Singhal V, Kumar M, Lakra A, Ranjan R, Pal R, Raza M, Singh U, Singh PK: Gabapentin provides effective postoperative analgesia whether administered pre-emptively or post-incision. Can J Anaesth 2005; 52:827-31
- 237. Pandey CK, Sahay S, Gupta D, Ambesh SP, Singh RB, Raza M, Singh U, Singh PK: Preemptive gabapentin decreases postoperative pain after lumbar discoidectomy. Can J Anaesth 2004; 51:986-9
- 238. Radhakrishnan M, Bithal PK, Chaturvedi A: Effect of preemptive gabapentin on postoperative pain relief and mor-

- phine consumption following lumbar laminectomy and discectomy: A randomized, double-blinded, placebo-controlled study. J Neurosurg Anesthesiol 2005; 17:125-8
- 239. Rapchuk II., O'Connell L, Liessmann CD, Cornelissen HR, Fraser JF: Effect of gabapentin on pain after cardiac surgery: A randomised, double-blind, placebo-controlled trial. Anaesth Intensive Care 2010; 38:445-51
- 240. Srivastava U, Kumar A, Saxena S, Mishra AR, Saraswat N, Mishra S: Effect of preoperative gabapentin on postoperative pain and tramadol consumption after minilap open
- cholecystectomy: A randomized double-blind, placebo-controlled trial. Eur J Anaesthesiol 2010; 27:331-5
- 241. Dierking G, Duedahl TH, Rasmussen ML, Fomsgaard JS, Møiniche S, Rømsing J, Dahl JB: Effects of gabapentin on postoperative morphine consumption and pain after abdominal hysterectomy: A randomized, double-blind trial. Acta Anaesthesiol Scand 2004; 48:322-7
- 242. Elander G, Hellström G: Analgesic administration in children and adults following open heart surgery. Scand J Caring Sci 1992; 6:17-21