Epidural analgesia compared with peripheral nerve blockade after major knee surgery: a systematic review and meta-analysis of randomized trials

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The relative analgesic efficacy and side-effect profile of peripheral nerve blockade (PNB) techniques compared with lumbar epidural analgesia for major knee surgery is unclear. We undertook a systematic review and meta-analysis of all randomized trials comparing epidural analgesia with PNB for major knee surgery. Eight studies were identified that had enrolled a total of 510 patients of whom 464 (91%) had undergone total knee joint replacement. All were small trials and none was blinded (Jadad score 1–3). PNB technique was variable: in addition to a femoral catheter (n=5), femoral single shot (n=2), or lumbar plexus catheter (n=1) techniques, sciatic blockade was performed in three trials. There was no significant difference in pain scores between epidural and PNB at 0–12 or 12–24 h, WMD 0.22 (95% CI: –0.36, 0.81), 0.05 (–1.01, 0.91), respectively, and no clinically significant difference at 24–48 h, WMD –0.35 (–0.64, –0.02). There was also no difference in morphine consumption (mg) at 0–24 h, WMD –6.25 (–18.35, 5.86). Hypotension occurred more frequently among patients who received epidurals [OR 0.19 (0.08, 0.45)], but there was no difference in the incidence of nausea and vomiting. Two studies reported a higher incidence of urinary retention in the epidural group. Patient satisfaction was higher with PNB in two of three studies which measured this, although rehabilitation indices were similar. PNB with a femoral nerve block provides postoperative analgesia which is comparable with that obtained with an epidural technique but with an improved side-effect profile and is less likely to cause a severe neuraxial complication.

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Major knee surgery such as total knee joint replacement (TKJR) and anterior cruciate ligament reconstruction (ACLR) is associated with moderate to severe postoperative pain which can contribute to immobility-related complications, delay in hospital discharge, and interfere with functional outcome.⁸ ³⁷ These lower limb procedures are amenable to regional anaesthesia techniques which reduce neuroendocrine stress responses, central sensitization of the nervous system and muscle spasms which occur in response to pain stimuli.¹ ²⁶

Epidural analgesia has been popular over recent decades as there is evidence for reduced blood loss and fewer thromboembolic complications using neuraxial techniques in orthopaedic surgery.⁴² A recent systematic review comparing lumbar epidural blockade with systemic opioid analgesia reported better dynamic pain scores in the epidural group but no difference in the incidence of side-effects overall.¹² Patients who received epidurals had more frequent hypotension, urinary retention, and pruritis whereas systemic opioids caused more sedation (but no difference was found with respect to respiratory depression or postoperative nausea and vomiting). More importantly, there is evidence that patients undergoing TKJR are at increased risk of serious neurological complications as a result of epidural blockade. This is most likely to be related to degenerative spinal changes and anticoagulant therapy.³²

An alternative regional anaesthesia technique is peripheral nerve blockade (PNB) of one or more major nerves
supplying the lower limb. PNB may provide effective uni-
lateral analgesia with a lower incidence of opioid-related
and autonomic side-effects, less motor block, and fewer
serious neurological complications compared with epidural
analgesia.\textsuperscript{11, 23} In contrast to epidural analgesia, continuous
PNB techniques do appear to provide pain relief superior
to systemic opioid analgesia but with a lower incidence of
side-effects.\textsuperscript{38} Advances in nerve localization such as
ultrasound imaging and continuous catheter technology
have also helped to increase interest in PNB for lower
limb surgery.\textsuperscript{18, 19}

Most previous studies comparing PNB with epidural
analgesia for major knee surgery have demonstrated equivalent
analgesia and an improvement in side-effect profile
associated with PNB. However, the number of patients
enrolled in each study is small and statistical significance is
not reached across all variables and all time periods
measured. The aim of this study was to undertake a systema-
tic review and meta-analysis of randomized trials of epidural
analgesia and PNB in adults undergoing major knee surgery,
including comparison of analgesic efficacy, side-effects/
complications, patient satisfaction, and rehabilitation indices.

**Methods**

A systematic review of randomized trials comparing epi-
dural analgesia with PNB for postoperative analgesia after
major knee surgery in adults was conducted. The
MEDLINE, EMBASE, Pubmed, and CINAHL databases,
Bandalier, and the Cochrane Controlled Trials Register
(CCTR) were searched from their inception to April 2007
by two of the authors (S.J.F. and S.S.). Abstracts and con-
ference proceedings of the American Society of
Anesthesiologists (ASA), International Anesthesia
Research Society (IARS), and European Society of
Anesthesiology (ESA) were searched manually from 2000
onwards to identify unpublished studies. In addition, we
undertook a Google search and contacted several research-
ers in the field. Prospective randomized trials were
searched using the following combinations of terms: total
knee replacement/major knee surgery/knee/cruciate liga-
ment, plus epidural/extradural/peridural/combined spinal
epidural, plus nerve block/femoral/sciatric/limbar plexus/
local anaesthesia. Language restrictions were not applied.
The reference lists of retrieved papers were scrutinized to
identify further studies for inclusion. Reports were
included if the study was a prospective randomized trial
comparing continuous lumbar epidural analgesia with
local anaesthetic agent compared with any peripheral
nerve block (single shot or continuous administered before
or after operation) in patients undergoing major knee
surgery. Exclusion criteria were opioid-only epidural,
single-shot epidural, and intrathecal-only techniques. A
total of 12 studies were identified but only eight papers
were included in the analysis. Three were clearly multiple
publications of the same research, although the method-
ology reported was variable.\textsuperscript{16, 21, 39} One prospective, ran-
domized study of patients undergoing ACLR was retrieved
from the search of unpublished conference abstracts,\textsuperscript{5} but
little data were presented in this negative study and further
details could not be obtained from the authors. None of
the included studies was double-blind, reflecting the
ethical issues surrounding the use of placebo catheters.

The studies were examined by two of the authors (S.J.F.
and J.S.) for measures that could be meaningfully com-
pared between the studies. If data were not available in
the original paper, the authors were contacted by e-mail to
request further information. The quality of individual trials
was quantified using the Jadad scale\textsuperscript{24} using five criteria:
(i) randomization, (ii) description of randomization, (iii)
blinding, (iv) adequacy of blinding, and (v) withdrawals
documented.

Outcome measures included: (i) analgesia efficacy,
visual analogue dynamic pain scores on the day of surgery
and the first and second days after surgery, morphine con-
sumption in the first 24 h after operation; (ii) adverse
effects, nausea and vomiting, hypotension, pruritis, urinary
retention, sedation, and motor block; (iii) patient satisfac-
tion; and (iv) rehabilitation indices. We also reported other
outcome data, including neurological complications, blood
loss, time to perform regional block, cardiovascular and
surgical complications, and local anaesthetic plasma
levels. All data were independently extracted and verified
by two of the authors (J.S. and S.S.) and differences
resolved by consensus.

Where pain scores were given but it was unclear
whether these were at rest or on movement, the worst pain
score for that time period was used. Where both nausea
and vomiting were reported, we used only the data for
vomiting, assuming that all patients who vomited would
have experienced nausea as well. Where nausea and
vomiting was reported as mild, moderate, or severe, the
worst score was used. Visual analogue pain scores were
converted to a standardized 0–10 scale. Variables which
were not reported numerically were estimated by manual
measurements from the published figures.

When no standard deviation (\(sd\)) was given for continu-
ous data, the \(sd\) was estimated as half the mean value.
When data were presented as 95% confidence intervals
(CI), the \(sd\) was calculated from the formula
\((sd=95\%\ CI/1.96×\sqrt{n})\). When the median and range were reported for
continuous outcomes, the mean and \(sd\) were estimated by
assuming that the mean was equivalent to the median and
that the \(sd\) was a quarter of the range.

Meta-analysis was undertaken using Review Manager
(RevMan for Windows version 4.2.9, The Cochrane
Collaboration, Oxford, UK) when sufficient data existed in
three or more studies. This software calculates the
weighted mean difference and 95% CI (by study size and
\(sd\)) between treatment arms for numerical data in each
study and estimates the overall pooled effect.
For dichotomous data, RevMan calculates odds ratio and 95% CI. If heterogeneity was significant ($P \leq 0.05$), a random effects model was used. If heterogeneity was non-significant ($P > 0.05$), a fixed effects model was used. Those parameters which were unsuitable for meta-analysis or reported in a single study were discussed in the text.

Subgroup analysis was performed for the five studies using femoral blockade alone in an attempt to detect a difference in quality of analgesia compared with combined sciatic and femoral/lumbar plexus block. We were unable to perform a subanalysis comparing quality of analgesia with single-shot vs continuous blocks as there were little data after the first 12 h after operation. Likewise, analysis of TKJR compared with other types of major knee surgery was not possible as we were unable to obtain subgroup data in those papers which included other procedures.

One of the studies compared pain scores and surgical stress hormones only in the very early postoperative period only (0–3 h) but was included in the analysis. In another study, which was also included, both groups received an epidural as the primary anaesthetic technique in the operating theatre and this was replaced at the end of surgery by a femoral nerve catheter in one study group.

**Results**

Of the eight studies included in this systematic review (Table 1), 1 2 6 9 17 27 46 54 five originated from Europe. There were 510 patients in total included in the analysis, of whom 464 (91%) had undergone TKJR. Other procedures were ACLR ($n=28$; 5.5%) and arthrolysis ($n=18$; 3.5%). The peripheral nerve block technique used was variable (Table 2)—the most common technique was a femoral sheath catheter alone (four studies), 6 9 27 46 then a single-shot femoral block in two studies, 1 17 and continuous lumbar plexus blockade in one study. 2 Sciatric nerve blockade was used in three of the eight studies of which two were single-shot techniques 17 and one a continuous catheter. 54 The primary anaesthetic technique was general anaesthesia in six studies and in the remaining two studies, spinal anaesthesia, 9 and epidural/PNB 2 were used.

When all studies were combined, there was no difference in visual analogue pain scores between the epidural and the PNB during the first two time periods (0–12 and 12–24 h) after operation (Figs 1 and 2). Subanalysis of studies with or without sciatic block did not change the results. Lower pain score at 24–48 h in the epidural group reached statistical significance [WMD −0.35 (95% CI: −0.64, −0.02); $P=0.04$] (Fig. 3). Analysis of three studies showed no difference in morphine consumption (mg) at 0–24 h 21 22 24 [WMD −6.25 mg (95% CI: −18.35, 5.86)]. Opiate consumption in two other studies which were not suitable for meta-analysis was also similar between the groups. 9 46

Hypotension occurred more frequently ($P=0.0001$) in patients who received an epidural (Fig. 4). There was no difference between the groups in the rate of nausea and vomiting (Fig. 5). The total number of patients with urinary retention, pruritis, or sedation was low (<50 patients in each treatment arm). However, both studies 46 54 that assessed urinary retention reported that this occurred more often among patients with epidurals [combined OR 0.07 (95% CI: 0.02, 0.27; $P<0.0001$)]. No difference in the incidence of pruritis or sedation was found in two studies. 9 54 There was no apparent bias on funnel plot analysis of adverse event data.

Of the three studies reporting patient satisfaction, 1 2 17 two stated that patient satisfaction was higher in the PNB group. 2 17 Mobilization or rehabilitation was described in six studies. 6 9 17 27 46 54 Rehabilitation indices varied but outcomes were generally similar except in one study which reported that better walking distance (day 0 and day 1) and knee extension (day 2, day 3, and at discharge) were obtained in patients with an epidural as a result of quadriceps weakness associated with PNB. 27 However, the data with respect to motor block on the operative side in two other studies were conflicting. 6 54 There was no difference between the groups in range of knee movement at 6 months.

Only one major neurological complication was reported—a patient who received an epidural developed foot drop and sphincteric disturbance after operation but the aetiology was not stated. 17 In another study, paraesthesia and numbness occurred more frequently in the epidural group but no further details were given. 9 There was no difference between the groups in any of the three studies measuring perioperative blood loss. 2 6 17 nor was there a difference in the number of cardiovascular complications overall 54 In one study, three patients in the PNB group required re-operation for closed manipulation ($n=2$) or wound dehiscence. 27 The study which reported plasma catecholamine and stress hormone levels concluded that epidural analgesia provides the greatest reduction in sympatho-adrenergic stress response in the early postoperative period, although analgesia and adverse effects were judged to be similar. In this study, plasma bupivacaine concentrations in 21 patients who received femoral nerve block using bupivacaine 150 mg remained well below toxic limits 15–180 min after administration. 1

Inadequate or failed block was described in four studies. 6 17 46 54 Overall, there were 10 of 130 (7.7%) patients in the epidural group and four of 133 (3.0%) patients in the PNB group who had treatment failure, although only one study included these patients in the analysis (intention-to-treat). 6 One study found no difference in block insertion times (mean 13 min in epidural and 12.5 min in combined femoral-sciatic single shot block groups; $P=0.92$) and time in the anaesthetic room (29.6 and 7.1 min; $P=0.34$). 17 but in another study there were more catheter problems in the epidural group such as
Table 1 Summary of randomized trials included in the meta-analysis. TKJR, total knee joint replacement; ACLR, anterior cruciate ligament repair; VAS, visual analogue score; GA, general anaesthetic; SE, side-effects; PCA, patient-controlled analgesia; IA, intraarticular.

<table>
<thead>
<tr>
<th>Study/Country</th>
<th>Type of surgery (Jadad score)</th>
<th>No. of patients/groups</th>
<th>Techniques</th>
<th>Duration</th>
<th>Other group</th>
<th>Additional analgesics</th>
</tr>
</thead>
<tbody>
<tr>
<td>Zarić and colleagues\textsuperscript{54}/Denmark</td>
<td>TKJR (3)</td>
<td>49/2</td>
<td>GA</td>
<td>Ropivacaine 0.75% boluses to block level T10 preop+infusion ropivacaine 0.25% with sufentanil postop</td>
<td>Femoral and sciatic catheters: preop bolus ropivacaine 0.75% 30 ml each catheter+infusion ropivacaine 0.2% with sufentanil (femoral) and ropivacaine 0.05% (sciatic) postop</td>
<td>55 h postop</td>
</tr>
<tr>
<td>Barrington and colleagues\textsuperscript{6}/Australia</td>
<td>TKJR (3)</td>
<td>108/2</td>
<td>Spinal</td>
<td>Ropivacaine 0.2% with fentanyl infusion postop</td>
<td>Femoral catheter: preop bolus ropivacaine 0.25% with adrenaline 25 ml+infusion ropivacaine 0.2% with PCA bolus postop</td>
<td>Postop D3</td>
</tr>
<tr>
<td>Davies and colleagues\textsuperscript{17}/UK</td>
<td>TKJR (3)</td>
<td>59/2</td>
<td>GA</td>
<td>Bupivacaine 0.5% bolus preop+infusion bupivacaine 0.25% postop</td>
<td>Single-shot femoral and sciatic blocks: preop bolus bupivacaine 0.375% 30 ml (femoral)+25 ml (sciatic) limited to 3 mg kg\textsuperscript{-1}</td>
<td>Postop D2</td>
</tr>
<tr>
<td>Singelyn and colleagues\textsuperscript{46}/Belgium</td>
<td>TKJR (2)</td>
<td>45/3</td>
<td>GA</td>
<td>Bupivacaine 0.25% with epinephrine and sufentanil bolus preop+infusion bupivacaine 0.125% with sufentanil and clonidine postop</td>
<td>Femoral catheter: preop bolus bupivacaine 0.25% with adrenaline 37 ml+infusion bupivacaine 0.125% with sufentanil and clonidine postop</td>
<td>48 h postop</td>
</tr>
<tr>
<td>Adams and colleagues\textsuperscript{1}/Germany</td>
<td>TKJR (1)</td>
<td>63/3</td>
<td>GA</td>
<td>Bupivacaine 0.375% bolus postop</td>
<td>Single-shot femoral block: postop bolus bupivacaine 0.375%</td>
<td>3 h postop</td>
</tr>
<tr>
<td>Capdevila and colleagues\textsuperscript{9}/France</td>
<td>TKJR and arthrolysis/3 (1)</td>
<td>38 TKJR and 18 arthrolysis/3</td>
<td>GA</td>
<td>Lidocaine 2% with epinephrine and morphine 2 mg to block level T10 bolus postop+infusion 1% lidocaine with clonidine and morphine</td>
<td>Femoral catheter: postop bolus 2% lidocaine with epinephrine 25 ml and morphine 2 mg+infusion 1% lidocaine with clonidine and morphine</td>
<td>Postop D3</td>
</tr>
<tr>
<td>Aldahish and colleagues\textsuperscript{2}/Egypt</td>
<td>TKJR and ACLR/2</td>
<td>32 TKJR and 28 ACLR/2</td>
<td>Regional</td>
<td>Bupivacaine 0.5% bolus preop+infusion bupivacaine 0.125% postop</td>
<td>Lumbar plexus catheter and single-shot sciatic block: preop bolus bupivacaine 0.5%+infusion bupivacaine 0.125% postop (lumbar plexus only)</td>
<td>48 h postop</td>
</tr>
<tr>
<td>Long and colleagues\textsuperscript{27}/USA</td>
<td>TKJR (2)</td>
<td>70/2</td>
<td>GA+Epidural</td>
<td>Ropivacaine 1% bolus preop+infusion ropivacaine 0.2% postop</td>
<td>Femoral catheter: postop bolus+infusion ropivacaine 0.2%</td>
<td>36 h postop</td>
</tr>
</tbody>
</table>

Continued
<table>
<thead>
<tr>
<th>Study/Country</th>
<th>Follow-up</th>
<th>Summary of results</th>
<th>Patient satisfaction</th>
<th>Rehab indices</th>
<th>Technical</th>
<th>Blood loss</th>
<th>Neurological complications</th>
<th>LA plasma conc</th>
</tr>
</thead>
<tbody>
<tr>
<td>Zaric and colleagues(^5)/Denmark</td>
<td>72 h postop+ day 7+ 6 week visit</td>
<td>Similar analgesia and rehab indices in both groups; less urinary retention and overall SE with PNB; more motor block in operated limb with PNB</td>
<td>No</td>
<td>Yes</td>
<td>Yes</td>
<td>No</td>
<td>None</td>
<td>No</td>
</tr>
<tr>
<td>Barrington and colleagues(^6)/Australia</td>
<td>D2 postop+ rehab</td>
<td>Similar analgesia and rehab indices in both groups but oxycodone and rofecoxib consumption higher with PNB; less PONV with PNB</td>
<td>No</td>
<td>Yes</td>
<td>Yes</td>
<td>No difference</td>
<td>None</td>
<td>No</td>
</tr>
<tr>
<td>Davies and colleagues(^7)/UK</td>
<td>D2 postop+rehab</td>
<td>Lower pain scores at 24 h and higher patient satisfaction at 48 h with PNB and no difference in SE profile</td>
<td>Yes</td>
<td>Yes</td>
<td>No difference</td>
<td>No difference</td>
<td>One patient in epidural group (see text)</td>
<td>No</td>
</tr>
<tr>
<td>Singelyn and colleagues(^8)/Belgium</td>
<td>D2 postop+ 6 weeks+ 3 months</td>
<td>Lower pain scores at 4 and 24 h+more knee flexion until weeks postop+shorter hospital stay with epidural and PNB vs PCA; lower mean pain scores at 4 h but urinary retention and catheter problems with epidural vs PNB</td>
<td>No</td>
<td>Yes</td>
<td>More catheter problems in epidural group</td>
<td>No</td>
<td>None</td>
<td>No</td>
</tr>
<tr>
<td>Adams and colleagues(^9)/Germany</td>
<td>D1 postop</td>
<td>Similar analgesia and SE profile in all groups; epidural best reduction in neuro-humoral stress response</td>
<td>Yes</td>
<td>No</td>
<td>No</td>
<td>No</td>
<td>Not known</td>
<td>Yes</td>
</tr>
<tr>
<td>Capdevila and colleagues(^4)/France</td>
<td>48 h postop+ rehab</td>
<td>Lower pain scores with better mobilization at 24 and 48 h+more knee flexion at discharge+shorter rehab centre stay with epidural and PNB vs PCA; less pain at rest 6–12 h postop but more hypotension, urinary retention with epidural vs PNB</td>
<td>No</td>
<td>Yes</td>
<td>No</td>
<td>No</td>
<td>Dysaesthesia common in epidural group ?signif</td>
<td>No</td>
</tr>
<tr>
<td>Aldahish and colleagues(^7)/Egypt</td>
<td>48 h postop</td>
<td>Similar analgesia in both groups; patient satisfaction higher in PNB group; more intraop hypotension with epidural</td>
<td>Yes</td>
<td>No</td>
<td>No</td>
<td>No difference</td>
<td>None</td>
<td>No clinical toxicity</td>
</tr>
<tr>
<td>Long and colleagues(^2)/USA</td>
<td>D3 postop+ discharge+ 6 months</td>
<td>Lower pain scores (D0 and D1) and lower opiate consumption (D1) with PNB but quad weakness resulting in better walking distance (D0 and D1) and better knee extension (D2, D3, discharge) with epidural</td>
<td>No</td>
<td>Yes</td>
<td>No</td>
<td>No</td>
<td>None</td>
<td>No</td>
</tr>
</tbody>
</table>
Discussion

The principal finding of this systematic review is that a PNB technique using femoral nerve blockade provides analgesia which is comparable with that obtained with an epidural but with a lower incidence of hypotension. This is consistent with the findings of two previous meta-analyses which compared systemic opioid analgesia with epidural analgesia\textsuperscript{12} and continuous PNB,\textsuperscript{38} respectively, and also with a previous report in patients undergoing hip surgery.\textsuperscript{50} We did not detect any benefit from the addition of a sciatic block to a femoral nerve blockade at 0–24 h after operation. The marginal improvement in pain score at 24–48 h in patients receiving epidural analgesia is unlikely to be clinically significant. Although the number of patients in the analysis was small (n=79), there was a reduction in the incidence of urinary retention using PNB (P<0.0001). This finding is consistent with previous systematic reviews\textsuperscript{12,38} and also with the pharmacology of local anaesthetic administered into the epidural space.

Limitations of the study

There were no large (n≥1000) randomized trials comparing PNB with epidural analgesia. We identified eight relevant studies so that the meta-analysis included 510 patients but only two studies defined a primary outcome measure. Variable reporting of end points and inconsistent definitions meant that we were unable to include every study for each outcome, despite attempting to contact authors. Although pooling of results increases statistical power, the findings need to be interpreted with caution as there were only a small number of patients identified for some variables. For example, any difference in the rate of pruritus and sedation would be difficult to detect as there were less than 50 patients in each arm of the meta-analysis.

There was also clinical heterogeneity among the studies. For example, there was wide variation in the type of PNB and different techniques such as femoral, combined femoral–sciatic, lumbar plexus, single-shot, and continuous blocks were all treated as one group. There was also variation in the drug, additives, and doses used in both arms of this systematic review (Table 1).

It is important to note that none of the studies reported improved long-term outcome with PNB, which is likely as the choice of analgesic technique is only one component of quality postoperative care, which must include expert physiotherapy and nursing care. Routine hospital discharge

<table>
<thead>
<tr>
<th>Table 2 Summary of peripheral block techniques employed in the eight included studies</th>
</tr>
</thead>
<tbody>
<tr>
<td>No. of studies</td>
</tr>
<tr>
<td>----------------</td>
</tr>
<tr>
<td>No sciatic block</td>
</tr>
<tr>
<td>Sciatic single-shot</td>
</tr>
<tr>
<td>Sciatic catheter</td>
</tr>
<tr>
<td>Total</td>
</tr>
</tbody>
</table>

Fig 1 A meta-analysis of trials comparing PNB with epidural on visual analogue pain scores at 0–12 h including sub-analysis with and without sciatic block.
and rehabilitation after major knee surgery is very variable between countries and institutions. A limitation of all systematic reviews is that negative studies are less likely to be submitted or accepted for publication. Despite these weaknesses, meta-analysis is considered a reliable source of evidence.

Comparison with other studies
The findings of several larger non-randomized series comparing PNB with epidural analgesia for lower limb surgery have been published which are consistent with the results of our study.\cite{11, 47, 48} Singelyn and Gouverneur\cite{47} reported that 540 patients undergoing major knee surgery who received either continuous ‘3-in-1’ block (n=415) or epidural analgesia (n=125) had similar dynamic pain scores with fewer side-effects and block failures in the PNB group. In another report, these authors reiterated that PNB was the technique of choice.\cite{48} Chelly and colleagues compared i.v. PCA morphine, continuous femoral with single-shot sciatic blockade and epidural analgesia in a cohort study of 92 patients undergoing TKJR. Patients who received PNB had improved pain control, less cardiovascular instability during surgery, and less nausea and vomiting compared with those who had epidurals or PCA. PNB also allowed better toleration of continuous passive motion with earlier mobilization and hospital discharge.\cite{11} Hebl and colleagues\cite{22} and Horlocker and colleagues\cite{23} described a ‘total joint regional anaesthesia clinical pathway’ used at their institution which emphasizes continuous lumbar plexus block along with single-shot sciatic block as part of a multimodal analgesia regimen after TKJR and have reported improved outcome compared with matched historical controls. Interestingly, PNB may also improve the quality of analgesia when added to epidural blockade after TKJR.\cite{53, 55}

**PNB of the lower limb**
The femoral nerve along with contributions from the sciatic and obturator nerves at the posterior and medial aspects, respectively, provide sensory innervation of the knee and these are the three terminal nerves are targeted by PNB techniques for major knee surgery.

With the exception of one, all the studies included in our meta-analysis used an anterior infra-inguinal femoral nerve block which is easily performed with the patient in the supine position. This approach is less invasive than lumbar plexus block and is correspondingly associated with fewer serious complications.\cite{4, 18, 45} Although the obturator nerve is far more consistently involved with lumbar plexus block than with either infra-inguinal technique,\cite{18} it is not clear that obturator block translates into improved patient recovery after TKJR.\cite{25, 28, 31} MRI studies suggest that the local anaesthetic solution spreads predominantly caudally after ‘3-in-1’ block using a peripheral nerve stimulator, blocking the femoral and lateral femoral cutaneous nerves and the anterior branch of the obturator nerve,\cite{30} although there is some evidence that the latter nerves are more reliably blocked with the more lateral, blind, ‘double-pop’ fascia iliaca block.\cite{33}
Of the three studies which used sciatic blockade, two performed the blocks using the classical Labat approach and one with an anterior approach. Since the earlier publication of one small negative study,\textsuperscript{3} trials have shown that the addition of sciatic nerve block improves the quality of analgesia by reducing posterior knee and calf pain after major knee surgery, corresponding to the area innervated by the nerve.\textsuperscript{71 31 53 45 2} However, our analysis of patients without sciatic block failed to demonstrate inferior analgesia in this subgroup between 0 and 24 h after operation.

**Neurological complications**

It was not a goal of our systematic review to calculate an incidence figure for serious neurological complications as much larger numbers of patients are needed to contribute meaningful data. However, we did identify one case of a unilateral foot drop with sphincteric disturbance in approximately 250 patients who received an epidural. No neurological complications were reported in the PNB group.

Spinal haematoma after epidural blockade is more likely when patient ‘red flags’ are present (e.g. older age, degenerative spinal disorders, anticoagulation) or after traumatic insertion,\textsuperscript{51} a common scenario in older patients undergoing lower limb orthopaedic surgery. The incidence of intravertebral haematoma in female patients undergoing TKJR with an epidural in Sweden during the 1990s was 1:3600.\textsuperscript{32} This is in close agreement with the estimated reporting rate in the USA during the mid-1990s,\textsuperscript{41} despite the utilization of lower routine doses of anticoagulation.\textsuperscript{49} Hypotension associated with epidural analgesia may contribute to end organ ischaemia or infarction in this group of patients if left untreated.\textsuperscript{35} Because of the problems which may occur, epidural analgesia is not appropriate in the ward setting in every institution. If general anaesthesia is not desirable, single-shot spinal anaesthesia can be combined with PNB\textsuperscript{44} and should be considered as the incidence of neuraxial complications is lower using this technique than with an epidural in this subpopulation.\textsuperscript{32}

A major benefit of PNB is that the neurological complications are usually less disastrous than those associated with neuraxial blockade. In the most comprehensive prospective study published to date,\textsuperscript{7} peripheral neuropathy occurred at a rate of 1:3763 femoral or sciatic block procedures (95% CI 0–8/10 000). Symptoms persisted after 6 months in more than half of patients who had a nerve injury, and the actual rate may be higher as many practitioners were regional anaesthesia enthusiasts (volunteer bias). A recent editorial estimated that the true rate of neuropathy attributable to PNB is probably around

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**Fig 3** A meta-analysis of trials comparing PNB with epidural on visual analogue pain scores at 24–48 h.

**Fig 4** A meta-analysis of trials comparing PNB with epidural on rate of hypotension after surgery expressed as odds ratio (OR).
Two series of continuous PNB techniques including 21114 and 77710 femoral catheters have been published which reported success rates and technical, infectious, and neurological complications. The rate of neuropraxia after femoral catheter techniques was very similar in these studies: 1/21114 and 3/683.10 Although neuropraxia occurred frequently, most of these injuries were patchy sensory deficits and all except one had resolved within a year. In these two studies, 28.6–56% of femoral catheters had bacterial colonization at C2148. There were no clinically significant local or surgical infectious complication in the first study,14 but in the second study, a psoas abscess was diagnosed in a diabetic woman after a femoral catheter.10 Placement was difficult in 9.5% of 211 catheters in the first study 14 (with vascular puncture occurring in 5.6%) whereas in the other study, 10 catheter-related problems (e.g. kinking, leakage, blockage, and dislodgement) occurred in 17.9% of 1416 patients. Salinas and colleagues40 reported better analgesia but a higher rate of nerve injury and infectious complications and no improvement in overall outcome with continuous catheters compared with single-shot femoral blocks. Unfortunately, we were unable to compare catheter vs single-shot techniques; therefore, it is not possible to state which is preferable.

The overall effect of quadriceps weakness associated with femoral nerve block on the operative side on recovery and patient satisfaction is unclear from the studies included in this systematic review. One group reported better mobilization with epidural analgesia,27 and in our own experience some patients find the leg weakness from femoral block unsettling. However, others comment that reduction of pain from significant quadriceps spasm which occurs after TKJR8 is central to the efficacy of PNB6 and improves toleration of continuous passive motion.9 Rehabilitation indices were similar in both groups, although patient satisfaction appears to be higher with PNB. Ropivacaine appears attractive as an agent because of reduced cardiotoxicity and motor block compared with bupivacaine. The optimal concentration appears to be around 0.2–0.25%.20 36

**Success rates**

The estimated success rate of 97% for PNB among the four studies6 17 46 54 included in this systematic review which reported block failures may reflect the expertise of the investigators and may not be reproducible if used by practitioners with less experience. Although this success rate is in close agreement with that reported by Capdevila and colleagues10 in a series of 1416 continuous PNB catheters, it is at the upper end of previously published success rates for femoral (86–100%) and sciatic block (94–100%).18 Similarly, although the mean insertion time for combined blocks of 12.5 min reported by Davies and colleagues17 is in close agreement with the insertion time reported by Hebl’s group,22 this may not be generally reproducible. Identification of the femoral and sciatic nerves using ultrasound guidance is relatively easy and rapid onset of block with success rates near 100% have been reported in experienced hands.29

### Conclusion

A PNB technique which includes femoral block represents the best balance between analgesia and side-effects as a choice of postoperative analgesic technique for major knee surgery such as TKJR, especially as the risk of injury to the neuraxis is negligible. Data are urgently required comparing efficacy and morbidity of single-shot blocks compared with perineural catheter techniques, preferably with a large randomized controlled trial so that a meaningful comparison of less common complications can be undertaken. More work is also needed to prove that newer techniques offer important advantages such as faster return to normal daily activities, decreased morbidity, and improved patient satisfaction. As with all anaesthetic procedures, complication rates reflect in part the skills and judgement of the operator and the risk of system error. It is important to consider risk–benefit on a patient-by-patient basis and tailor the analgesic technique accordingly. Overall, however, we believe that there is now sufficient evidence

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### Table: Comparison of studies

<table>
<thead>
<tr>
<th>Study or subcategory</th>
<th>PNB n/N</th>
<th>Epidual n/N</th>
<th>OR (fixed)</th>
<th>Weight %</th>
<th>OR (fixed)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Adams and colleagues</td>
<td>3/21</td>
<td>5/21</td>
<td>33.11</td>
<td>0.53 (0.11, 2.59)</td>
<td></td>
</tr>
<tr>
<td>Aldahish and colleagues</td>
<td>0/30</td>
<td>1/30</td>
<td>11.40</td>
<td>0.32 (0.01, 8.24)</td>
<td></td>
</tr>
<tr>
<td>Capdevilla and colleagues</td>
<td>5/20</td>
<td>1/17</td>
<td>12.18</td>
<td>0.27 (0.01, 7.59)</td>
<td></td>
</tr>
<tr>
<td>Jingsen and colleagues</td>
<td>5/15</td>
<td>4/15</td>
<td>20.60</td>
<td>1.98 (0.29, 6.60)</td>
<td></td>
</tr>
<tr>
<td>Zaric and colleagues</td>
<td>2/26</td>
<td>3/23</td>
<td>22.70</td>
<td>0.96 (0.08, 3.66)</td>
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</tr>
<tr>
<td>Total (95% CI)</td>
<td>112</td>
<td>106</td>
<td>100.00</td>
<td>0.66 (0.28, 1.55)</td>
<td></td>
</tr>
</tbody>
</table>

Test for heterogeneity: X²=1.42, df=4 (P=0.84), F=0%

Test for overall effect: Z=0.36 (P=0.34)
that lumbar epidural analgesia should not be used routinely and that PNB is appropriate for multimodal analgesia care after routine major knee surgery.

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References
36 Ng HP, Cheong KF, Lim A, Lim J, Puhaindran ME. Intraoperative single-shot ‘3-in-1’ femoral nerve block with ropivacaine 0.25%, ropivacaine 0.5% or bupivacaine 0.25% provides comparable 48-hr analgesia after unilateral total knee replacement. Can J Anaesth 2001; 48: 1102–8
43 Schug SA. Peripheral nerve blockade carries only a minimal risk of permanent neurological complications. Anaest Intensive Care 2007; 35: 11–2
44 Simanski C, Fischer B. Procedure specific postoperative pain management (PROSPECT) recommendations for total knee arthroplasty. 2007; http://www.postoppain.org
48 Soubiron L, Singelyn FJ. Postoperative analgesia after elective lower limb orthopaedic surgery: IV PCA, continuous peripheral nerve blocks or continuous epidural analgesia? Br J Anaesth 1997; 78: A415
49 Tryba M, Wedel D. Central neuraxial block and low molecular weight heparin (enoxaparin): lessons learned from different dosage regimes in two continents. Acta Anaesthesiol Scand 1997; 41: 100–3