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Analgesic efficacy of single-shot adductor canal block versus adductor canal block combined with intra-articular ropivacaine infusion after total knee arthroplasty

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Aims

Single-shot adductor canal block (ACB) after total knee arthroplasty (TKA) for postoperative analgesia is a common modality. Patients can experience breakthrough pain when the effect of ACB wears off. Local anaesthetic infusion through an intra-articular catheter (IAC) can help manage breakthrough pain after TKA. We hypothesized that combined ACB with ropivacaine infusion through IAC is associated with better pain relief compared to ACB used alone.

Methods

This study was a prospective double-blinded placebo-controlled randomized controlled trial to compare the efficacy of combined ACB+ IAC-ropivacaine infusion (study group, n = 68) versus single-shot ACB+ intra-articular normal saline placebo (control group, n = 66) after primary TKA. The primary outcome was assessment of pain, using the visual analogue scale (VAS) recorded at 6, 12, 24, and 48 hours after surgery. Secondary outcomes included active knee ROM 48 hours after surgery and additional requirement of analgesia for breakthrough pain.

Results

The study group (mean visual analogue scale (VAS) pain score of 5.5 (SD 0.889)) experienced significant reduction in pain 12 hours after surgery compared to the control group (mean VAS 6.62 (SD 1.356); mean difference = 1.12, 95% confidence interval (CI) -1.46 to 0.67; $p < 0.001$), and pain scores on postoperative day (POD) 1 and POD-2 were lower in the study group compared to the control group (mean difference in VAS pain = 1.04 (-1.39 to -0.68, 95% CI, $p < 0.001$). Fewer patients in the study group (0 vs 3 in the control group) required additional analgesia for breakthrough pain, but this was not statistically significant. The study group had significantly increased active knee flexion (mean flexion 86.4° (SD 7.22°)), compared to the control group (mean 73.86° (SD 7.88°), mean difference = 12.54, 95% CI 9.97 to 15.1; $p < 0.014$).

Conclusion

Combined ACB+ ropivacaine infusion via IAC is a safe, reproducible analgesic modality after primary TKA, with superior analgesia compared to ACB alone. Further large volume trials are warranted to generate evidence on clinical significance on analgesia after TKA.

Cite this article: *Bone Jt Open* 2021;2-12:1082–1088.

Keywords: Arthroplasty, Knee, Analgesia, Pain management, Rehabilitation

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doi: 10.1302/2633-1462.212.BJO-2021-0119.R1

Bone Jt Open 2021;2-12:1082–1088.

Introduction

Total knee arthroplasty (TKA) is one of the most commonly performed orthopaedic procedures. Postoperative pain following

TKA is a challenge and can interfere with patient rehabilitation. Inadequate pain control may reduce patient participation in physical therapy, resulting in longer length

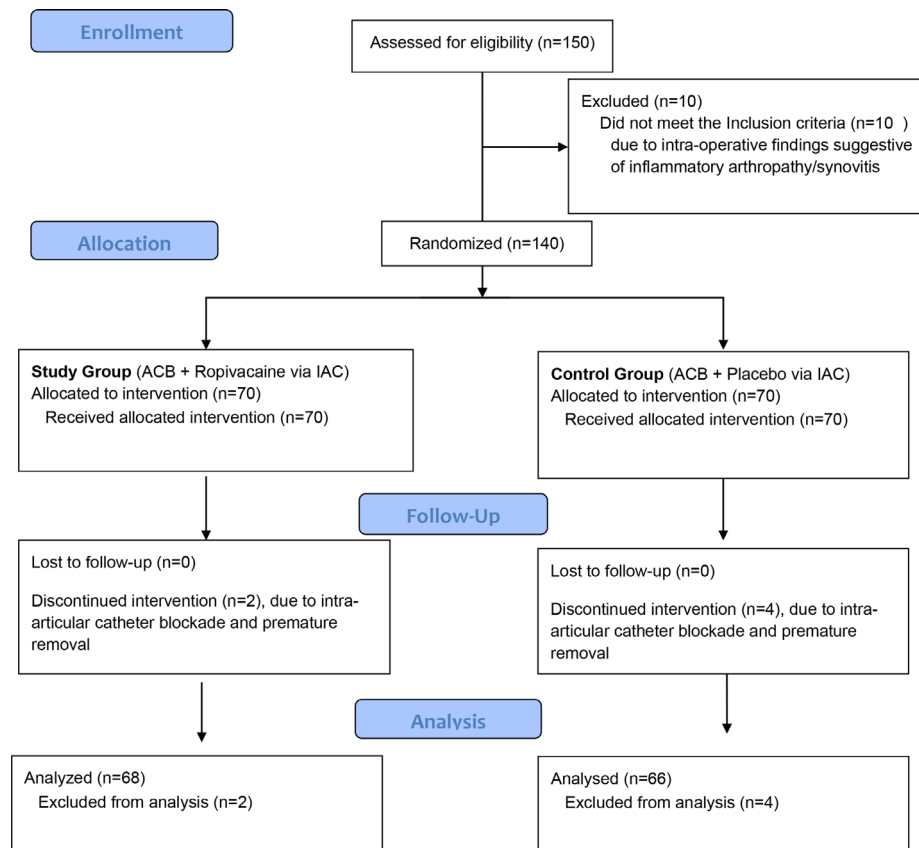


Fig. 1

Consolidated Standards of Reporting Trials flow diagram.

of stay (LOS) and poorer outcomes. Pain management strategies have evolved from systemic opioids to epidural anaesthesia to peripheral nerve blockade, trending towards a more local approach which has shown promising results.¹⁻³ Multimodal analgesia has been shown to be superior to monotherapy, subsequently reducing opiate use after surgery.

Peripheral nerve blocks and local anaesthetic agents have several advantages when compared to regional or systemic methods. These include less toxicity and early mobilization.^{2,3} Problems with epidural anaesthesia are mainly related to delayed rehabilitation, and bladder and bowel problems.⁴ There have been reports of quadriceps weakness with femoral nerve block, leading to delayed mobilization.⁵ Patient-controlled analgesia often leads to under-dosage or over-dosage of the drug, requiring frequent monitoring and dose adjustment for adequate pain relief, to minimize opioid-related side effects.⁶

Adductor canal block (ACB) is a peripheral nerve blockade technique which provides pain relief without compromising quadriceps strength and allows early rehabilitation.^{1,7} However, the majority of patients experience block resolution and breakthrough pain on the day of surgery. Andersen et al⁸ reported a mean duration of only 10.5 hours of pain relief with adductor canal block.

We hypothesize that local anaesthetic infusion through a catheter can help manage breakthrough pain and provide prolonged analgesia. Intra-articular catheter (IAC) with infusion of a local anaesthetic (ropivacaine) has also been reported to provide adequate analgesia following surgery without any systemic effects.⁹ Dannana et al¹⁰ studied the efficacy of ACB+ multimodal periarticular infiltration (MPI) with and without local anaesthetic infusion via IAC, reporting increased pain relief in the group with additional local anaesthetic infusion. However, this was a non-randomized prospective study. To our knowledge, there are no placebo-controlled double-blind randomized controlled trials (RCTs) which studied the efficacy of local anaesthetic infusion via IAC in combination with single-shot ACB.

The aim of this study was to compare the post-operative analgesic efficacy of single-shot adductor canal blockade (ACB) in isolation (in combination with placebo normal saline infusion through IAC) versus single-shot ACB in combination with intra-articular ropivacaine infusion after primary TKA. Secondary objectives were to compare active knee range of motion (ROM) and analgesia-related complications between the two groups.

Table 1. Baseline information of study participants.

Variable	Study group (n = 68)	Control group (n = 66)
Mean age, yrs (SD)	66.4 (9.4)	66.6 (8.6)
Sex, n (%)		
Male	28 (41.1)	31 (46.9)
Female	40 (58.8)	35 (53.03)
Mean height, cm (SD)	157 (8.5)	163 (7.4)
Mean weight, kg, (SD)	68 (11.4)	72 (9.6)
Mean BMI, kg/m ² (SD)	27.5 (2.5)	27.09 (1.8)
Mean CCI (SD)	3 (1.15)	3.17 (1.27)
ASA classification, n (%)		
ASA 1	11 (16.2)	13 (19.7)
ASA 2	57 (83.8)	53 (80.3)
Mean duration of surgery, mins (SD)	70 (8.5)	68 (10.2)
Mean preoperative VAS (SD)	6.29 (0.648)	6.0 (0.765)
Mean preoperative knee flexion, ° (SD)	100.5 (15.4)	105 (9.8)

ASA, American Society of Anesthesiologists; CCI, Charlson Comorbidity Index; SD, standard deviation; VAS, visual analogue scale.

Methods

This study was a placebo-controlled double-blinded RCT conducted at a single centre. The trial was approved by the Institutional Ethics Committee (SS/2018/IEC 296) and prospectively registered with the Clinical Trials Registry of India (CTRI/2018/08/015386). The trial was to compare the postoperative analgesic efficacy of single-shot ACB alone (control group) versus single-shot ACB combined with intra-articular ropivacaine infusion via IAC (study group), in patients undergoing primary total knee arthroplasty. The trial was conducted after obtained written informed consent of trial participants and according to the Declaration of Helsinki guidelines.¹¹

Patients undergoing unilateral TKA for primary osteoarthritis (OA) of the knee were eligible for participation. Inclusion criteria comprised patients of either sex, aged 20 years and older, with primary OA of the knee joint, who consented to undergo unilateral primary TKA. Exclusion criteria were patients undergoing bilateral simultaneous or staggered TKA, inflammatory arthropathy, post-traumatic arthritis, history of previous surgeries on the knee, gross deformities (varus or valgus more than 20° or fixed-flexion deformity more than 15°), and patients who could not comprehend the visual analogue pain scale (VAS).

All patients participating in this study were operated under spinal anaesthesia, and received 2.5 ml of 0.5% hyperbaric bupivacaine intrathecal. All surgeries were performed by a single surgeon (AVGR), using a standard medial-parapatellar approach and a posterior-stabilized prosthesis design (P.F.C. Sigma Posterior-Stabilized Knee, DePuy Synthes, USA). In all cases, an IAC was used. The participants of the study group were randomized to receive ultrasound-guided ACB in combination with a local anaesthetic infusion (20 ml, 0.2% ropivacaine, administered every six hours for the first 24 hours after

surgery) via IAC. The control group received ACB in combination with equal-volume placebo (20 ml normal saline, administered every six hours for the first 24 hours after surgery) via IAC. ACB was administered in both groups immediately after the surgery in the operating room, with 20 ml of 0.2% ropivacaine. A bolus of 20 ml of 0.2% ropivacaine equates to 40 mg of the local anaesthetic. Administration of intra-articular boluses every six hours adds up to a cumulative ropivacaine dose of 160 mg of ropivacaine, which is less than the daily maximum dose of the local anaesthetic (225 mg) in adults, as reported by Rosenberg.¹²

Patients in both study groups received oral celecoxib (200 mg) and gabapentin (300 mg) as pre-emptive analgesia 12 hours before surgery. A uniform postoperative analgesic protocol was implemented with intravenous (IV) paracetamol (1 g) every eight hours for the first 24 hours postoperatively, followed by a switch to oral celecoxib (200 mg) twice a day, oral paracetamol (1 g) three times a day and gabapentin (300 mg) once a day till discharge. Breakthrough pain in the recovery room (first 24 hours) was managed with IV fentanyl bolus given, based on body weight, by the anaesthesiologist. Breakthrough pain in the ward was managed with stat intravenous tramadol 50 mg. IACs were removed on postoperative day (POD) 2 in all cases and all patients were discharged on the second postoperative day.

Outcomes measured. The primary outcome (postoperative pain) was assessed using the VAS. The VAS score was scored out of 10 and was recorded with the patient at rest. VAS pain was recorded at six and 12 hours postoperatively on POD 0 (day of surgery), every eight hours on POD 1, and once on POD 2 before the patient is discharged. All patients were started on physical therapy with knee ROM and quadriceps exercises on POD 1.

The secondary outcomes (active knee ROM, additional analgesic requirements for breakthrough pain, and immediate postoperative complications) were recorded on POD 2. Side effects of ropivacaine toxicity (including blurred vision, chest pain, dizziness, syncope, peri-oral paraesthesias, acute onset delirium, and cardiac arrhythmias) were monitored and documented in the immediate postoperative period.

Randomization. A total of 150 patients were enrolled in the study and subsequently 140 patients were randomized into two groups. Randomization was performed with the help of a computer-generated, blocked random-allocation sequence with a 1:1 ratio. Patients enrolled in the study and the operating surgeon were blinded to modality of analgesia prior to surgery. Postoperative pain and functional assessment by an arthroplasty fellowship trainee (VC), who was also blinded to the randomization scheme. A sealed envelope was attached to the file of the patient on admission, with details of random allocation. Patients were randomized into two groups: the study

Table II. Postoperative pain and range of motion assessment.

Outcome measured	Study group (n = 68)	Control group (n = 66)	Mean difference (95% CI)	p-value*
Mean VAS pain 6 hrs postop (SD)	4.88 (0.744)	4.98 (0.969)	0.1 (-0.39 to 0.19)	0.494
Mean VAS pain 12 hrs postop (SD)	5.5 (0.889)	6.62 (1.356)	1.12 (-1.46 to -0.67)	< 0.001
Mean VAS pain, 6 am, POD 1 (SD)	5.49 (0.906)	6.53 (1.427)	1.04 (-1.486 to -0.67)	< 0.001
Mean VAS pain, 12 pm, POD 1 (SD)	4.96 (0.905)	6 (1.129)	1.04 (-1.39 to -0.68)	< 0.001
Mean VAS pain, 6 pm, POD 1 (SD)	5.5 (0.68)	6.42 (0.929)	0.92 (-1.25 to -0.68)	< 0.001
Mean VAS pain, 6 am, POD 2 (SD)	5.53 (0.922)	6.52 (0.864)	0.99 (-1.30 to -0.69)	< 0.001
Mean active knee ROM on POD 2, ° (SD)	86.40 (7.220)	73.86 (7.884)	13 (9.97 to 15.10)	< 0.001

*Independent-samples *t*-test.

CI, confidence interval; POD, postoperative day; ROM, range of motion; SD, standard deviation; VAS, visual analogue scale.

group (combination of single-shot ACB and local anaesthetic infusion (bolus of 20 ml of 0.2% ropivacaine every six hours) via IAC), and the control group (combination of single-shot ACB and placebo (bolus 20 ml of normal saline every six hours) via IAC).

Statistical analysis. Assessment of whether the data were normally distributed was made using the Kolmogorov-Smirnov test. Continuous variables were analyzed using the independent-samples *t*-test. Parametric categorical variable data were analyzed using the chi-squared test and non-parametric data was analyzed with Fisher's exact test. The SPSS 19.0 software (IBM, USA) was used for the statistical analysis. The nature of hypothesis testing was two-tailed and a *p*-value < 0.005 was considered statistically significant. As VAS pain score assessment was a repeated outcome measure, analysis of variance (ANOVA) was performed to compare the mean scores at different time intervals to analyze significance.

To show a difference of 'one point' on the VAS scale (scored 0 to 10), pre-study power analysis estimated a sample size of 50 participants in each group with 95% confidence intervals (CIs) and power of 0.8. Significance was defined as a *p*-value < 0.05. Based on published studies,^{13,14} we considered a one-point difference on the VAS pain scale to be a clinically significant change. Estimating a loss to follow-up of 15% to 20%, we set a target of 75 participants in each group.

Results

The flow diagram of the trial, based on Consolidated Standards of Reporting Trials (CONSORT) guidelines,¹⁵ is illustrated in Figure 1. A total of 140 study participants were randomized into two groups: the study group (n = 70, ACB+ ropivacaine infusion via IAC) and the control group (n = 70, ACB+ normal saline placebo via IAC). Six patients were excluded (two from the study group and four from the control group) due to IAC blockade requiring premature removal of the IAC. Overall, 134 trial participants (68 study group, 66 control group) were included in the final data analysis.

Baseline demographic and clinical characteristics of trial participants are summarized in Table I.

Primary outcome assessment: postoperative pain and analgesic efficacy of intervention.

The pain difference, as assessed by VAS pain scores, was not significantly different in the first six hours after surgery (study group mean VAS pain score 4.88 (standard deviation (SD) 0.744) vs control group 4.98 (SD 0.969); mean difference = 0.1 with 95% CI -0.39 to 0.19; *p* = 0.494, ANOVA). By the end of POD 0, study group (ACB+ ropivacaine) with mean VAS pain score of 5.5 (SD 0.889) experienced significant reduction in pain compared to the control group (mean VAS 6.62 (SD 1.356), mean difference = 1.12, 95% CI -1.46 to -0.67; *p* < 0.001, ANOVA for repeated measures). Similarly, VAS scores at rest on POD 1 and POD 2 were lower in the study group compared to the control group (Table II) and the difference was found to be statistically significant (mean difference in VAS pain = 0.97 with 95% CI -1.254 to -0.686; *p* < 0.001, ANOVA for repeated measures).

Comparison of breakthrough pain and additional analgesic requirements.

None of the study group participants required additional analgesics for breakthrough pain. Adequate analgesia (VAS pain score < 5/10), was achieved in all study group participants with intra-articular ropivacaine infusion along with intravenous paracetamol 1 g, every eight hours. Three of 66 (4.5%) control group participants received additional analgesia for breakthrough pain. All three subjects received a single dose of IV tramadol (50 mg/IV) on POD 1, and two patients required another single dose of IV tramadol (50 mg) on POD 2.

Secondary outcomes. Active ROM (knee flexion) was assessed by goniometer on the second postoperative day. Patients in the study group had increased active knee ROM (mean flexion 86.4° (SD 7.22)), and was found to be significantly better than the control group (mean flexion 73.86 (SD 7.88), mean difference = 12.54 with 95% CI 9.97 to 15.1; *p* = 0.017, independent-samples *t*-test).

Perioperative complications are summarized in Table III. In total, 19 patients in the study group (27.9%) and 12 patients in the control group (18.2%) experienced increased wound soakage requiring daily compression dressing. Overall, 19 (27.9%) patients in the study group and 17 (25.75%) patients in the

Table III. Immediate postoperative complications in study population.

Complications, n (%)	Study group (n = 68)	Control group (n = 66)	p-value*
Nausea/vomiting	19 (27.94)	17 (25.75)	0.343
Breakthrough pain	0	3 (4.5)	0.451
Opiate analgesia for breakthrough pain	Nil	Tramadol cumulative dose 2 patients - 100 mg 1 patient - 50 mg	
Wound soakage	19 (27.94)	12 (18.18)	0.301

*Chi-squared test.

control group experienced nausea requiring medication, however this was not statistically significant. No patient had any other wound-related complication or infection. No incidence of catheter breakage or becoming stuck in the knee joint was encountered. No patients in either group had complications associated with toxicity of the local anaesthetic (including blurred vision, chest pain, dizziness, syncope, perioral paraesthesias, acute onset confusion or delirium, and no documentation of cardiac arrhythmias).

None of the trial participants, of either group, were documented to have surgical site infections (SSIs), peri-prosthetic joint infections (PJIs), complications related to treatment, or medical comorbidities in the first six months after surgery.

Discussion

This study demonstrated that combined single-shot ACB with intra-articular ropivacaine infusion was associated with a significant reduction in postoperative pain after primary TKA, in comparison to only single-shot ACB. Active knee ROM was significantly better in the study group.

Pain after TKA can cause dissatisfaction and negatively influence rehabilitation after surgery. Benefits of peripheral nerve blockade and its role in faster postoperative rehabilitation have been studied and widely reported. Multimodal analgesia is now the standard of care for analgesia after TKA. This includes pre-emptive analgesia, regional anaesthesia at the time of surgery followed by various options such as oral and IV analgesics, peripheral nerve blocks, and opiates.

Adductor canal blocks can be administered either as single-shot injections or by continuous adductor canal catheter infusions, and were introduced as 'pure sensory' nerve blocks without associated motor weakness. Single-shot adductor canal block has been established as an efficient source of pain control and improved rehabilitation.^{1,7} Compared to other available peripheral nerve blocks, such as the sciatic block or the femoral nerve block (FNB), ACB is associated with reduced motor deficits after surgery, thus aiding faster recovery and rehabilitation. Several reports support the use of ACB over

FNB to avoid quadriceps weakness and reduce the time required to attain an independent ambulatory status.¹⁶⁻²⁰ Toftdahl et al²¹ concluded that peri- and intra-articular analgesia following TKA has better analgesia and mobilization compared to continuous femoral nerve block.

Direct delivery of local anaesthetic agents into the joint after TKA is another established analgesic modality. Gomez-Cardero et al⁹ reported that use of ropivacaine infusion pumps was associated with improved postoperative pain scores and reduced opioid use. It also improved immediate functionality and patient comfort, resulting in reduced LOS without added complications. An earlier randomized, double-blinded, placebo-controlled study for IAC (using ropivacaine) versus epidural plus single-shot FNB by Reinhardt et al²² concluded that IAC is a safe and effective analgesic modality.

In their study, Andersen et al²³ found that peri and intra-articular infusion of ropivacaine was associated with improved analgesia when compared to epidural infusions. Smith et al²⁴ reported lower pain scores for patients receiving intra-articular bupivacaine infusion when compared to periarticular liposomal bupivacaine injection. Ikeuchi et al²⁵ performed a double-blinded RCT to study the efficacy of local anaesthetic infusion (with steroid) after TKA, and reported significantly reduced pain in the group receiving local anaesthetic infusion via catheter. This study population received patient-controlled analgesia (PCA) as per institutional protocol, without use of peripheral nerve blockade after TKA.

In contrast, De-Weese et al²⁶ found they had more use of breakthrough analgesia in group receiving continuous intra-articular bupivacaine compared with another group on epidural analgesia. Reeves and Skinner²⁷ could not find any positive benefit of ropivacaine intra-articular infusion in their study.

A RCT by Beausang et al²⁸ compared the independent use of ACB versus intra-articular local anaesthetic infusions, and showed the superiority of ACB in reducing pain and opiate consumption 24 hours after TKA. Dannana et al¹⁰ evaluated pain relief after TKA with ACB with periarticular infiltration of the knee, versus ACB combined with local infiltration and intra-articular ropivacaine boluses. They concluded that the additional use of intra-articular ropivacaine was associated with significant reduction in pain after surgery. Gurava Reddy et al²⁹ performed a multicentre non-randomized study comparing continuous adductor-canal block by catheter versus local anaesthetic infusion by intra-articular catheter, with comparable pain relief with both analgesic methods.

Use of peripheral nerve blocks is not without complications. Feibel et al³⁰ reported increased falls after surgery in patients receiving peripheral nerve blocks, especially continuous FNB. ACB has also been reported to result in motor weakness due to proximal spread of the local anaesthetic.³⁰⁻³³ However, this is a relatively rare

complication. In this study, no patient experienced falls after surgery in the immediate postoperative period or during early follow-up.

The strengths of this study are its design (prospective, randomized, double-blinded, placebo-controlled), and the fact that the surgery performed by a single surgeon. The study was adequately powered to detect changes in the VAS pain scale, and there was a uniform analgesic protocol for management of postoperative pain.

The risk of introducing infection with IAC is minimal, and no study has reported any serious complications with IAC.⁸ The only disadvantage with IAC in our view is the additional cost and increased wound soakage.

A limitation of our study is that VAS pain scores were recorded with the patient at rest and not during active physical therapy. Further, there was a cumulative dose difference in local anaesthetic used between both the groups, however this was within the safe maximal-dose limits of ropivacaine. Further studies on optimal local anaesthetic drug dosage and duration of treatment are warranted.

In conclusion, combined single-shot ACB with intra-articular local anaesthetic infusion is effective in managing postoperative pain following TKA and promoting early rehabilitation of patients, without increasing the risk of complications. Further large volume trials are warranted to generate evidence on clinical significance on analgesia after TKA.



Take home message

- Breakthrough pain after total knee arthroplasty remains a impediment to recovery and rehabilitation. Adductor canal block (ACB) is effective in managing breakthrough pain after the effect of spinal anaesthesia wears off.

- Since the ACB analgesic effect is short-lived, we propose the use of intra-articular local anaesthetic infusion for breakthrough pain after surgery.

- In our study, patients benefited from local anaesthetic infusion with reduced pain and improved knee range of motion. Further large volume trials will help establish a clinical practice guideline for the use of intra-articular local anaesthetic infusions for postoperative pain management after primary total knee arthroplasty.

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Funding statement:

- No benefits in any form have been received or will be received from a commercial party related directly or indirectly to the subject of this article.

ICMJE COI statement:

- P. Mulpur reports an institutional grant from the *Journal of Bone & Joint Surgery* (Dr. Robert Bucholz Journal Club Grant 2020-2021), unrelated to this study.

Acknowledgements:

- We acknowledge the contribution of Dr. Khubchand Jhakotia (Consultant Anaesthesiologist).

Ethical review statement:

- This study was approved by the Institute Ethics Committee.

Open access funding

- The authors confirm that the open access fee for this study was self-funded.

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