

RESEARCH**Open Access**

Postoperative outcomes of mepivacaine vs. bupivacaine in patients undergoing total joint arthroplasty with spinal anesthesia

Laura A. Stock, Kevin Dennis, James H. MacDonald, Andrew J. Goins, Justin J. Turcotte*  and Paul J. King

Abstract

Background: Spinal anesthesia (SA) has been previously associated with improved outcomes after total joint arthroplasty (TJA). The purpose of this study was to compare outcomes between various local anesthetics.

Methods: This was a retrospective study of 1,328 patients undergoing primary TJA with SA from September 2020–2021 at a single institution. Patients were grouped based on TKA or THA and further separated and analyzed in terms of anesthetic agents—mepivacaine (M), hyperbaric bupivacaine (HB), or isobaric bupivacaine (IB). Subgroup analysis of same-day-discharge (SDD) patients and low- (<11 mg) and high-dose bupivacaine was performed. Statistical significance was assessed at $P < 0.05$.

Results: Mepivacaine use was associated with younger age, lower ASAs, and lower Charlson Comorbidity Index (CCI) scores in both THAs and TKAs. Postoperatively, significant differences were found between HB, IB, and M in LOS, the first PT ambulation distance and rates of SDD, and home discharge in both THAs and TKAs. No significant differences in outcomes were observed between high- and low-dose bupivacaine in THAs or TKAs. In SDD patients, a significant difference was found only in the first 6-clicks mobility scores. After controlling for age, BMI, sex, ASA, and procedure type, mepivacaine was found to be associated with shorter LOS, increased likelihood of SDD, home discharge, POD-0 ambulation, and, further, the first ambulation distance. No significant differences were observed in 6-clicks mobility scores, urinary retention, 30-day ED returns or 30-day readmissions.

Conclusions: Both bupivacaine and mepivacaine are safe and effective local anesthetics for patients undergoing TJA as evidenced by low, similar rates of urinary retention and 30-day ED returns and readmissions. Mepivacaine does appear to facilitate early ambulation, shorter LOS and home discharge and should be considered as the local anesthetic of choice for patients undergoing rapid recovery TJA.

Keywords: Spinal anesthesia (SA), Total joint arthroplasty (TJA), Surgical outcomes

Background

As the population ages, exponential growth in total joint arthroplasties (TJA) are expected. Prior studies have cited an expected 284% increase in total hip arthroplasty (THA) and a 401% increase in total knee arthroplasty

(TKA) by the year 2030 [1]. In patients undergoing TJA, the use of different anesthetic techniques has also been shown to influence a variety of surgical outcomes, such as length of stay, time to ambulation, distance of ambulation, home discharge rate, and readmission or emergency department (ED) return rate [2, 3].

Pain management is a key component of postoperative care that influences the patient's early ambulation and ability to participate in physical therapy, both of which are important predictive measures for improved patient

*Correspondence: Jturcotte@aaahs.org

Anne Arundel Medical Center, 2000 Medical Parkway, Suite 503, Annapolis, MD 21401, USA



© The Author(s) 2022. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit <http://creativecommons.org/licenses/by/4.0/>.

outcomes. Early ambulation has been shown to result in shorter length of stay (LOS), lower pain levels, and lower incidence of deep venous thrombosis (DVT) and pulmonary infection [4]. Further, physical therapy intervention on postoperative day 0 has been associated with a shorter length of stay, longer ambulation distance, and greater home discharge rate [5, 6].

Anesthetic selection is a critical component of early postoperative pain management and early ambulation. Although general anesthesia is historically used in these procedures, multiple recent studies have demonstrated the benefits of regional anesthesia (RA) [7, 8]. Regional anesthesia methods include spinal blocks, epidural blocks, and peripheral nerve blocks [9]. One of the most widely used anesthesia methods for total joint procedures is spinal anesthesia. Recent studies have focused on evaluating the effectiveness of various local spinal anesthetics, primarily mepivacaine and bupivacaine, to compare their effect on postoperative outcomes following THA and TKA. Mepivacaine is a short-acting local anesthetic with an intermediate duration of action between 90 to 240 minutes, depending on dosage and administration [10, 11]. Bupivacaine is a long-acting local anesthetic with an early onset and duration of action of 2 to 5 hours [12]. However, the onset and duration may also be manipulated depending on the formulation of the bupivacaine, such as hyperbaric or isobaric. Hyperbaric bupivacaine has a density greater than cerebrospinal fluid whereas isobaric bupivacaine is equal to the density of cerebrospinal fluid [13]. This allows for a quicker onset for the hyperbaric bupivacaine and shorter duration of motor and sensory block compared to the slower onset and longer duration of action of the isobaric formulation [13]. One study found those treated with mepivacaine had earlier ambulation, earlier return to motor function, and were more likely to have the same-day discharge compared to those treated with hyperbaric or isobaric bupivacaine [2]. Another study used a double-blind, randomized clinical trial to investigate the effects of mepivacaine and bupivacaine on a patient's return to motor function following total hip and knee arthroplasty and found a mean time of 26 minutes sooner with mepivacaine [3]. Mepivacaine has not only been shown to shorten time to ambulation and length of stay, but also has a faster neurological recovery when compared to bupivacaine [11].

The purpose of this study was to investigate the effects of spinal anesthesia on the surgical outcomes of total knee and hip arthroplasty. We compared the effects of hyperbaric bupivacaine, isobaric bupivacaine, and mepivacaine as well as high and low doses of bupivacaine on postoperative outcomes, such as length of hospital stay, ambulation on day 0, failed same day discharge attempt,

first 6-clicks mobility score, the first PT ambulation distance, urinary retention, home discharge rate, and 30-day readmission or ED return rate.

Methods

Study population

This study was deemed institutional review board exempt by the institution's clinical research committee. A retrospective observational study was performed in 1,328 patients who underwent primary unilateral THA or TKA with spinal anesthesia from September 2020 through September 2021. Patients undergoing bilateral or revision surgery, and those receiving general anesthesia were excluded from the study. All procedures were performed by 10 board-certified orthopedic surgeons. Procedures were performed at either an acute care hospital or an affiliated ambulatory surgery center (ASC).

Perioperative protocol

All patients were cared for in a coordinated Joint Replacement Center and received written educational materials, a nurse taught preoperative course, preoperative medical evaluations, and preoperative strengthening programs, including home exercise or outpatient physical therapy. An established rapid recovery protocol was utilized for all patients and the protocol included a multimodal pain management regimen of celecoxib, acetaminophen, pregabalin, and short-acting opioids. Patient-controlled analgesia and nerve blocks were not used in this patient population. All patients received spinal anaesthetic agents administered into the intrathecal space via a lumbar puncture. At anesthesiologist discretion and on patient request, spinal anesthesia was paired with propofol sedation. Patients receiving spinal anesthesia were not intubated, mechanically ventilated, and did not receive inhaled anesthetic agents.

Study design and data analysis

The patients were first separated between THA and TKA. Patient presentation and outcomes were then compared between those receiving hyperbaric bupivacaine, isobaric bupivacaine or mepivacaine. Patient demographics (age, body mass index [BMI], and sex) and comorbidity burden as measured in terms of the American Society of Anesthesiologists (ASA) score and Charlson Comorbidity Index (CCI) and were compared across groups. For each group, details of local anesthetic dosing were evaluated. Postoperative outcomes were then compared across groups. Univariate analysis, including Chi-square tests and Fisher's Exact tests, were used to compare categorical variables. For continuous measures, two-sided independent samples *t*-tests and one-way analysis of variance (ANOVA) were conducted to compare presentation and

outcomes across anesthetic types. *Post-hoc* Bonferroni adjustment was performed to make multiple comparisons across continuous endpoints. Two subgroup analyses were then performed to compare outcomes between low dose (<11 mg) and high dose (≥11 mg) bupivacaine, and between same-day discharge patients receiving either bupivacaine or mepivacaine. Finally, multivariate linear and Logistic regression models were created to evaluate the compare outcomes between patients receiving bupivacaine and mepivacaine after controlling for age, BMI, sex, ASA score, and procedure type. All statistical analyses were performed by means of SPSS (version 27.0, IBM, Armonk, NY). Statistical significance was assessed at $P < 0.05$.

Outcome measures

The following outcomes were assessed: 0-day LOS, LOS days, postoperative day 0 (POD0) ambulation, failed same day discharge (SDD) attempt, first Activity Measure for Post-Acute Care (AM-PAC) 6-Clicks mobility score, the first physical therapy (PT) documented ambulation distance, urinary retention, home discharge, 30-day readmission and 30-day ED return. POD0 ambulation included any documented ambulation by nursing or PT. The 6-Clicks mobility score is a tool used to assess

a patient’s mobility and ability to complete tasks. It contains a series of questions graded on a 1 to 4 point scale, in which a physician or physical therapist completes in order to assess a patient’s progress [14]. Urinary retention was defined as any instance in which a straight or foley catheter was placed postoperatively. Readmissions and ED-returns were captured for patients returning to both the same hospital and outside hospitals.

Results

Demographics and comorbidities

Comparison of demographics and comorbidities between hyperbaric bupivacaine, isobaric bupivacaine, and mepivacaine found significant differences in age, ASA, and Charlson Comorbidities Index (CCI) score across groups. Both TJA cohorts showed that mepivacaine use was associated with younger patients (THA 63.2 ± 9.6 , $P=0.001$; TKA 66.1 ± 8.5 , $P<0.001$), ASA score of less than 3 (THA $P<0.001$; TKA $P=0.001$), and lower CCI scores (THA $P<0.001$; TKA $P=0.004$) (Table 1). The medication dosing for each of the three groups are detailed in Table 2.

Postoperative outcomes

In evaluating postoperative outcomes, patients receiving mepivacaine had the highest rate of 0-day length of stay

Table 1 Patient demographics and comorbidities

| Variable – n (%) or Avg. ± SD | Total Hip Arthroplasty | | | P-Value | Total Knee Arthroplasty | | | P-Value |
|-------------------------------|--------------------------------|-----------------------------|---------------------|------------------|--------------------------------|-----------------------------|---------------------|------------------|
| | Hyperbaric Bupivacaine (n=401) | Isobaric Bupivacaine (n=34) | Mepivacaine (n=232) | | Hyperbaric Bupivacaine (n=502) | Isobaric Bupivacaine (n=28) | Mepivacaine (n=131) | |
| Age | 66.7 ± 11.4 | 65.3 ± 12.0 | 63.2 ± 9.6 | 0.001 | 68.9 ± 8.3 | 68.9 ± 8.2 | 66.1 ± 8.5 | <0.001 |
| BMI | 29.4 ± 5.4 | 30.0 ± 6.0 | 28.5 ± 5.2 | 0.072 | 31.7 ± 5.3 | 31.3 ± 5.5 | 30.7 ± 4.9 | 0.061 |
| Female | 224 (55.9) | 14 (41.2) | 130 (56.0) | 0.242 | 314 (62.5) | 20 (71.4) | 131 (50.0) | 0.001 |
| ASA ≥ 3 | 154 (38.4) | 18 (52.9) | 50 (21.6) | <0.001 | 225 (44.8) | 10 (35.7) | 82 (31.3) | 0.001 |
| CCI | 3.15 ± 1.78 | 3.09 ± 1.73 | 2.48 ± 1.56 | <0.001 | 3.35 ± 1.76 | 3.29 ± 1.27 | 2.91 ± 1.70 | 0.004 |

P-Values <0.05 in bold

BMI body mass index, ASA American Society of Anesthesiologists Score, CCI Charlson Comorbidity Index

Table 2 Spinal anesthetic dosing

| Dose - mg | Total Hip Arthroplasty | | | Total Knee Arthroplasty | | |
|-----------------|--------------------------------|-----------------------------|---------------------|--------------------------------|-----------------------------|---------------------|
| | Hyperbaric Bupivacaine (n=401) | Isobaric Bupivacaine (n=34) | Mepivacaine (n=232) | Hyperbaric Bupivacaine (n=502) | Isobaric Bupivacaine (n=28) | Mepivacaine (n=131) |
| Mean | 10.9 | 12.7 | 63.4 | 10.7 | 11.0 | 65.2 |
| SD | 1.7 | 2.1 | 75.3 | 1.6 | 2.4 | 54.5 |
| Median | 10.5 | 12.5 | 60.0 | 10.5 | 11.1 | 60.0 |
| 25th Percentile | 10.5 | 11.5 | 56.0 | 9.8 | 10.0 | 60.0 |
| 75th Percentile | 12.0 | 15.0 | 60.0 | 11.3 | 12.0 | 60.0 |

(THA 60.8%; TKA 52.7%, both $P < 0.05$) and the shortest overall length of stay (THA 0.43 ± 0.59 ; TKA 0.51 ± 0.59 , both $P < 0.05$). No statistically significant differences in rates of failed SDD attempts were observed across anesthetic groups. Mepivacaine patients were also more likely to ambulate on postoperative day 0 compared to bupivacaine for both THA and TKA ($P < 0.05$). Mepivacaine and hyperbaric bupivacaine patients each had higher first 6-Clicks mobility scores compared to isobaric bupivacaine in THA but not in TKA (between groups $P = 0.003$, *post hoc* $P < 0.05$). In both the THA and TKA groups, significant differences in the first PT ambulation distance were observed across anesthetic types (both $P < 0.001$), but a significant improvement in ambulation distance between the mepivacaine and other groups was observed only in patients undergoing TKA. There was no significant difference in rates of urinary retention amongst either the THA or TKA groups. Significant differences in the rate of home discharge were observed in the THA group, with mepivacaine patients being discharged home more frequently (100%) than either bupivacaine group ($P < 0.05$). No significant difference in the rate of home discharge was observed in the TKA group. There was a significant difference in 30-day readmission rates for the TKA group, with patients receiving

hyperbaric bupivacaine experiencing higher rates than those receiving mepivacaine ($P < 0.05$). No significant differences in readmission rates were observed across anesthetic groups for THA patients, and no significant differences in 30-day ED returns were observed for either THA or TKA (Table 3).

In the subgroup analysis comparing low dose (< 11 mg) and high dose (≥ 11 mg) bupivacaine, no significant differences in any outcome measure were observed for either the THA or TKA groups (Table 4). In the second subgroup analysis comparing bupivacaine and mepivacaine in same-day discharge patients, significant difference was observed only in a higher first 6-Clicks mobility score in patients receiving bupivacaine. This was observed in both the THA and TKA groups ($P \leq 0.001$) (Table 5).

Risk-adjusted outcomes

The multivariate results presented in Table 6 display the effect of mepivacaine in comparison to bupivacaine, after controlling for age, BMI, sex, ASA, and procedure type (THA or TKA). After controlling for these factors, patients receiving mepivacaine were more likely to have 0-day LOS (OR: 5.767, $P < 0.001$) and to have shorter overall LOS ($\beta = -0.421$ days, $P < 0.001$). Further,

Table 3 Unadjusted postoperative outcomes: all spinal anesthesia groups

| Outcome – n (%) or Avg. \pm SD | Total Hip Arthroplasty | | | | Total Knee Arthroplasty | | | |
|------------------------------------------------|--------------------------------|-----------------------------|---------------------|------------------|--------------------------------|-----------------------------|----------------------|------------------|
| | Hyperbaric Bupivacaine (n=401) | Isobaric Bupivacaine (n=34) | Mepivacaine (n=232) | P-Value | Hyperbaric Bupivacaine (n=502) | Isobaric Bupivacaine (n=28) | Mepivacaine (n=131) | P-Value |
| 0-Day LOS | 96 (23.9)a | 8 (23.5)a | 141 (60.8)b | <0.001 | 55 (11.0)a | 4 (14.3)a | 138 (52.7)b | <0.001 |
| LOS Days | $0.92 \pm 0.79a$ | $1.12 \pm 1.00a$ | $0.43 \pm 0.59b$ | <0.001 | $1.09 \pm 0.91a$ | $1.00 \pm 0.61a$ | $0.51 \pm 0.59b$ | <0.001 |
| Failed SDD Attempt | 22 (5.5)a | 4 (11.8)a | 14 (6.0)a | 0.334 | 17 (3.4) | 0 (0.0) | 6 (2.3) | 0.449 |
| Ambulated POD0 | 282 (70.3)a | 19 (55.9)a | 196 (84.5)b | <0.001 | 328 (65.3)a | 17 (60.7)a | 225 (85.9)b | <0.001 |
| First 6-Clicks Mobility Score ^a | $20.48 \pm 2.75a$ | $18.82 \pm 2.92b$ | $20.43 \pm 2.51a$ | 0.003 | $20.13 \pm 2.75a$ | $20.5 \pm 2.5a$ | $19.98 \pm 2.50a$ | 0.607 |
| First PT Ambulation Distance (ft) ^b | $151.52 \pm 97.51a$ | $142.55 \pm 114.21a,b$ | $184.30 \pm 84.84b$ | <0.001 | $134.57 \pm 98.89a$ | $111.67 \pm 77.44a$ | $165.24 \pm 101.79b$ | <0.001 |
| Urinary Retention | 5 (1.2)a | 2 (5.9)b | 3 (1.3)a,b | 0.097 | 6 (1.2)a | 0 (0.0)a | 2 (0.8)a | 0.734 |
| Home Discharge | 389 (97.0)a | 31 (91.2)a | 232 (100.0)b | 0.001 | 486 (96.8)a | 27 (96.4)a,b | 260 (99.2)b | 0.106 |
| 30-Day Readmission | 8 (2.0)a | 0 (0.0)a | 4 (1.7)a | 0.699 | 30 (6.0)a | 0 (0.0)a,b | 4 (1.5)b | 0.008 |
| 30-Day ED Return | 15 (3.7)a | 0 (0.0)a | 5 (2.2)a | 0.305 | 15 (3.0)a | 0 (0.0)a | 9 (3.4)a | 0.599 |

P-Values < 0.05 in bold

Subscript letters denote *post hoc* groups that do not differ from each other at $P < 0.05$, continuous values are Bonferroni adjusted.

LOS length of stay, SDD same day discharge, POD postoperative day, PT physical therapy, ED emergency department

^a n = THA 367 HB, 33 IB, 188 M; TKA 502 HB, 28 IB, 262 M

^b n = 362 HB, 33 IB, 179 M; TKA 467 HB, 27 IB, 191 M

Table 4 Low dose (<11 mg) vs. High Dose (≥11 mg) bupivacaine

| Outcome – n (%) or Avg. ± SD | Total Hip Arthroplasty | | | Total Knee Arthroplasty | | |
|------------------------------------------------|-------------------------------|------------------------------|------------------|-------------------------------|------------------------------|------------------|
| | High Dose Bupivacaine (n=213) | Low Dose Bupivacaine (n=222) | P-Value | High Dose Bupivacaine (n=203) | Low Dose Bupivacaine (n=327) | P-Value |
| Avg. Dose | 12.41 ± 1.12 | 9.76 ± 1.23 | <0.001 | 12.19 ± 1.08 | 9.80 ± 1.15 | <0.001 |
| Isobaric Bupivacaine | 31 (14.6) | 3 (1.4) | <0.001 | 17 (8.4) | 11 (3.4) | 0.012 |
| 0-Day LOS | 48 (22.5) | 56 (25.2) | 0.511 | 18 (8.9) | 41 (12.5) | 0.191 |
| LOS Days | 0.95 ± 0.78 | 0.91 ± 0.84 | 0.620 | 1.07 ± 0.73 | 1.09 ± 0.98 | 0.823 |
| Failed SDD Attempt | 20 (6.0) | 6 (6.1) | 0.968 | 13 (3.4) | 4 (2.8) | 0.757 |
| Ambulated POD0 | 141 (66.2) | 160 (72.1) | 0.185 | 134 (66.0) | 211 (64.5) | 0.728 |
| First 6-Clicks Mobility Score ^a | 20.29 ± 2.95 | 20.39 ± 2.66 | 0.740 | 20.11 ± 2.71 | 20.17 ± 2.76 | 0.822 |
| First PT Ambulation Distance (ft) ^b | 160.42 ± 96.02 | 141.07 ± 100.99 | 0.052 | 132.54 ± 100.27 | 133.82 ± 96.53 | 0.887 |
| Urinary Retention | 5 (2.3) | 2 (0.9) | 0.276* | 2 (1.0) | 4 (1.2) | 1.000* |
| Home Discharge | 206 (96.7) | 214 (96.4) | 0.856 | 195 (96.1) | 318 (97.2) | 0.450 |
| 30-Day Readmission | 2 (0.9) | 6 (2.7) | 0.285* | 11 (5.4) | 19 (5.8) | 0.850 |
| 30-Day ED Return | 7 (3.3) | 8 (3.6) | 0.856 | 4 (2.0) | 11 (3.4) | 0.347 |

P-Values <0.05 in bold

LOS length of stay, SDD same day discharge, POD postoperative day, PT physical therapy, ED emergency department

*Denotes Fisher’s Exact Test

^a n= THA 198 High, 202 Low; TKA 195 High, 307 Low

^b n= THA 198 High, 197 Low; TKA 194 High, 307 Low

Table 5 Subgroup analysis of bupivacaine vs. mepivacaine in same day discharge patients

| Outcome – n (%) or Avg. ± SD | Total Hip Arthroplasty | | | Total Knee Arthroplasty | | |
|------------------------------------------------|------------------------|---------------------|--------------|-------------------------|---------------------|------------------|
| | Bupivacaine (n=104) | Mepivacaine (n=141) | P-Value | Bupivacaine (n=59) | Mepivacaine (n=138) | P-Value |
| Isobaric Bupivacaine | 8 (7.7) | 0 (0.0) | N/A | 4 (6.8) | 0 (0.0) | N/A |
| Ambulated POD0 | 104 (100.0) | 141 (100.0) | N/A | 59 (100.0) | 138 (100.0) | N/A |
| First 6-Clicks Mobility Score ^a | 21.26 ± 2.28 | 20.16 ± 2.38 | 0.001 | 22.08 ± 2.15 | 19.96 ± 2.54 | <0.001 |
| First PT Ambulation Distance (ft) ^b | 193.81 ± 73.84 | 208.93 ± 66.60 | 0.144 | 220.25 ± 90.13 | 215.53 ± 82.85 | 0.757 |
| Urinary Retention | 1 (1.0) | 2 (1.4) | 1.000* | 1 (1.7) | 0 (0.0) | 0.299* |
| Home Discharge | 104 (100.0) | 141 (100.0) | N/A | 59 (100.0) | 138 (100.0) | N/A |
| 30-Day Readmission | 0 (0.0) | 2 (1.4) | 0.509* | 2 (3.4) | 2 (1.4) | 0.585* |
| 30-Day ED Return | 0 (0.0) | 4 (2.8) | 0.138* | 2 (3.4) | 2 (3.6) | 1.000* |

P-Values <0.05 in bold

POD postoperative day, PT physical therapy, ED emergency department

*Denotes Fisher’s Exact Test

^a n= THA 91 B, 105 M; TKA 52 B, 103 M

^b n= THA 90 B, 96 M; TKA 52 B, 81 M

they were more likely to ambulate on the day of surgery (OR: 2.391, $P<0.001$), ambulated further ($\beta=21.785$ ft., $P<0.001$), and were more likely to be discharged home (OR: 6.537, $P=0.011$). In the risk-adjusted analysis, spinal anesthetic type had no significant effect on the same day discharge failure rates, the first 6-Clicks mobility

score, urinary retention, 30-day readmission, or 30-day ED returns.

Discussion

In alignment with previous studies, our risk-adjusted results demonstrated that mepivacaine was associated with shorter length of stay, increased likelihood of

Table 6 Risk-adjusted outcomes: bupivacaine vs. mepivacaine

| Endpoint | Mepivacaine β/OR | 95% CI | P-Value |
|-----------------------------------|---------------------|-----------------|------------------|
| 0-Day LOS | 5.767 | 4.357 – 7.634 | <0.001 |
| LOS Days (β) | -0.421 | -0.502 – -0.339 | <0.001 |
| Failed SDD | 0.801 | 0.458 – 1.400 | 0.436 |
| Ambulated POD-0 | 2.391 | 1.789 – 3.197 | <0.001 |
| First 6-Clicks Mobility Score (β) | -0.209 | -0.523 – 0.105 | 0.192 |
| First PT Ambulation Distance (β) | 21.785 | 10.459 – 33.111 | <0.001 |
| Urinary Retention | 0.661 | 0.225 – 1.942 | 0.452 |
| Home Discharge | 6.537 | 1.540 – 27.743 | 0.011 |
| 30-Day Readmission | 0.494 | 0.226 – 1.083 | 0.078 |
| 30-Day ED Return | 1.068 | 0.552 – 2.067 | 0.845 |

P-values <0.05 in bold

LOS length of stay, SDD same day discharge, POD postoperative day, PT physical therapy, ED emergency department

the same-day discharge, increased early ambulation rates and distance, and increased likelihood of home discharge after both THA and TKA. However, these findings might be skewed by patient selection and preferential use of mepivacaine in patients self-selecting for early discharge, as evidenced by the lack of significant differences in outcomes when comparing bupivacaine and mepivacaine in that population. When evaluating patients receiving bupivacaine, we observed no significant differences between high and low doses. Our findings suggest that both bupivacaine and mepivacaine are safe and effective local anesthetics for patients undergoing TJA as evidenced by low, similar rates of urinary retention and 30-day ED returns and re-admissions.

The results of our study are in alignment with prior studies, demonstrating that mepivacaine use is associated with earlier ambulation, increased same-day discharge, and decreased length of stay [2, 11]. Schwenk *et al.* carried out a randomized controlled trial comparing mepivacaine (52.5 mg), hyperbaric bupivacaine (11.25 mg), and isobaric bupivacaine (12.5 mg) amongst 154 patients undergoing THA [2]. The results of their study were consistent with our data in support of mepivacaine for earlier ambulation, increased discharge rate, and shorter length of stay, although higher levels of early postoperative pain and opioid consumption occurred with mepivacaine use [2]. Additionally, the study found no significant differences in urinary retention, transient neurologic symptoms, hypotension, muscle tension or dizziness, supporting our assertion that both bupivacaine and mepivacaine are safe for use in TJA with spinal anesthesia [2]. In another retrospective review of 156 patients undergoing TKA

at a single institution, mepivacaine was associated with shorter length of stay (28.1 ± 11.2 hours *vs.* 33.6 ± 14.4 hours, $P=0.002$) and less straight catheterization (3.8% *vs.* 16.5%, $P=0.021$), compared to bupivacaine. In alignment with the results of Schwenk *et al.*, patients receiving mepivacaine had slightly higher pain scores and morphine consumption in the post-anesthesia care unit, but showed no difference in pain scores or morphine consumption afterwards [15]. A systematic review by Siddiqi *et al.* investigated five studies comparing mepivacaine and bupivacaine in TJA and found that mepivacaine was associated with a faster return to motor function, shorter LOS, and decreased urinary retention [16]. Further, this study found no significant differences in pain scores or ambulation distance [16]. While our study did not evaluate pain scores specifically, our finding that mepivacaine patients were more likely to ambulate POD0 and ambulate further suggests that any increased levels of early postoperative pain may not be clinically significant enough to impact early recovery. In another small randomized controlled trial comparing 32 patients receiving either bupivacaine or mepivacaine, significant differences were found in return to sensory function ($P=0.015$) and return of motor function ($P=0.025$), both favoring mepivacaine [11]. Urinary retention occurrences and time to urination were also better with mepivacaine compared to bupivacaine ($P=0.039$) [11]. A study by Calkins *et al.* evaluated spinal anesthesia in THA at an ambulatory surgery center (ASC) and found mepivacaine, when compared with bupivacaine, to be associated with less time in the ASC ($P<0.001$), decreased time to controlled voiding ($P<0.001$), and decreased time to ambulation ($P<0.001$) [17]. Further, this study found no significant differences in pain scores, complications, ED returns, or re-admissions [17]. However, bupivacaine was associated with a greater number of patients experiencing zero pain compared to the mepivacaine group [17]. Based on these studies and the results of our study, significant evidence demonstrating that mepivacaine facilitates faster ambulation and decreased length of stay exists. Our study and the Schwenk's study deviated from the others presented in that we did not observe significant improvement in urinary retention in the mepivacaine group, although overall rates were low regardless of spinal anesthetics used. Based on prior evidence showing increased risk for postoperative urinary retention in TJA patients placed under spinal anesthesia who had a history of urinary retention and a larger volume of intraoperative fluid, we suggest restrictive intraoperative fluid management strategies be followed regardless of whether bupivacaine or mepivacaine is used [18]. Further, our results and those of

Calkins *et al.* demonstrated no significant differences in 30-day ED returns or re-admissions, showing that early complications requiring intervention are similar across spinal anesthetic types. However, further study into rates of specific early and late complications is warranted to fully evaluate the effects of spinal anesthesia type on TJA recovery.

Our subgroup analysis found no significant differences between high dose and low dose bupivacaine, which is consistent with other published literature [19]. In a retrospective review of 761 TJAs, Herndon *et al.* found no significant differences in perioperative outcomes, including LOS and discharge disposition, when comparing 15 mg vs. <15 mg of isobaric bupivacaine [19]. While their study supports our data in finding no significant differences in surgical outcomes between high and low doses of bupivacaine, we suggest that the lowest possible therapeutic dose be used to mitigate risk of side effects.

A notable aspect of our study is the difference in unadjusted outcomes between patients undergoing THA and TKA. Specifically, 6-Clicks mobility scores and rates of home discharge showed statistically significant differences across anesthetic types in THA but not TKA patients, while 30-day re-admissions showed statistically significant differences in TKA but not THA patients. Based on the retrospective observational nature of this study, it is difficult to determine whether differences in patient characteristics and comorbidities, the surgical procedure, anesthesia type, or a combination of these factors accounts for these differences. However, our finding that mepivacaine use was not associated with differences in 6-Clicks mobility scores and re-admissions after adjusting for these confounding factors suggests that anesthesia type has less impact on these outcomes than procedure type and patient characteristics. Conversely, the finding that mepivacaine was associated with higher likelihood of home discharge after risk adjustment demonstrates that anesthesia type has a more direct effect on discharge disposition.

This study does contain multiple limitations. First, as an observational study it was exposed to selection bias, as the selection of anesthesia type was at the discretion of the provider. As previously described, our institution has informally adopted a recommendation that mepivacaine is the preferred local anesthetic for planned same-day discharge TJAs. This therefore makes it difficult to decipher whether the favorable outcomes observed in the mepivacaine group were due to patient selection or the anesthetic itself. Second, it is possible that additional confounding factors, including other specific comorbidities or psychosocial factors, influenced the observed outcomes. Third, our evaluation of isobaric and hyperbaric bupivacaine preparations was underpowered, thus

limiting any conclusions that can be drawn from any negative results.

Conclusion

The results of this study demonstrated that both bupivacaine and mepivacaine are safe and effective local anesthetics for patients undergoing TJA with spinal anesthesia. However, mepivacaine is associated with shorter length of stay, increased likelihood of the same-day discharge, increased early ambulation rates and distance, and increased likelihood of home discharge after both THA and TKA. We therefore suggest that mepivacaine should be considered the first-choice spinal anesthetic for patients undergoing rapid recovery TJA.

Abbreviations

SA: Spinal anesthesia; TJA: Total joint arthroplasty; M: Mepivacaine; HB: Hyperbaric bupivacaine; IB: Isobaric bupivacaine; SDD: Same-day-discharge; CCI: Charlson comorbidity index; THA: Total hip arthroplasty; TKA: Total knee arthroplasty; LOS: Length of stay; DVT: Deep venous thrombosis; RA: Regional anesthesia; ASC: Ambulatory surgery center; BMI: Body mass index; ASA: American Society of Anesthesiologists; POD0: Postoperative day 0; AM-PAC: Activity measure for post-acute care; PT: Physical therapy.

Acknowledgements

Not applicable.

Authors' contributions

LS, JT, and PK were major contributors in writing this manuscript. JT performed the statistical analysis regarding this project. KD contributed to data collection for this project. JT and PK provided oversight of this project. All authors read and approved the final manuscript.

Funding

This project did not receive any funding.

Availability of data and materials

The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

Declarations

Ethics approval and consent to participate

Institutional review board approval was obtained.

Consent for publication

All authors certify that they've read and approved this manuscript.

Competing interests

No relevant disclosures.

Received: 10 January 2022 Accepted: 14 June 2022

Published online: 13 July 2022

References

1. Singh JA, et al. Rates of total joint replacement in the united states: future projections to 2020-2040 using the national inpatient sample. *J Rheumatol.* 2019;46(9):1134–40.
2. Schwenk ES, et al. Mepivacaine versus bupivacaine spinal anesthesia for early postoperative ambulation. *Anesthesiology.* 2020;133(4):801–11.
3. Wyles CC, et al. More predictable return of motor function with mepivacaine versus bupivacaine spinal anesthetic in total hip and total knee

- arthroplasty: a double-blinded, randomized clinical trial. *J Bone Joint Surg Am.* 2020;102(18):1609–15.
4. Lei YT, et al. Benefits of early ambulation within 24 h after total knee arthroplasty: a multicenter retrospective cohort study in China. *Mil Med Res.* 2021;8(1):17.
 5. Chen AF, et al. Effect of immediate postoperative physical therapy on length of stay for total joint arthroplasty patients. *J Arthroplasty.* 2012;27(6):851–6.
 6. Sarpong NO, et al. Same-day physical therapy following total knee arthroplasty leads to improved inpatient physical therapy performance and decreased inpatient opioid consumption. *J Arthroplasty.* 2019;34(12):2931–6.
 7. Basques BA, et al. General compared with spinal anesthesia for total hip arthroplasty. *J Bone Joint Surg Am.* 2015;97(6):455–61.
 8. Paziuk TM, et al. General vs spinal anesthesia for total joint arthroplasty: a single-institution observational review. *J Arthroplasty.* 2020;35(4):955–9.
 9. JRH, F. Anesthesia for hip and knee surgery. 2018 cited 2021; Available from: <https://orthoinfo.aaos.org/en/treatment/anesthesia-for-hip-and-knee-surgery>.
 10. Salinas FV, Malik K, Benzon HT. Local anesthetics for regional anesthesia and pain management. *Raj's Pract Manage Pain.* 2008;811–38.
 11. Mahan MC, et al. Time of return of neurologic function after spinal anesthesia for total knee arthroplasty: mepivacaine vs bupivacaine in a randomized controlled trial. *Arthroplast Today.* 2019;5(2):226–33.
 12. Susan J. Dreyer, M.a.W.J.B., MD, Commonly used medications in procedures. *Pain procedures in clinical practice*, 2015: p. 5-12.
 13. Uppal V, et al. Hyperbaric versus isobaric bupivacaine for spinal anesthesia: systematic review and meta-analysis for adult patients undergoing noncesarean delivery surgery. *Anesth Analg.* 2017;125(5):1627–37.
 14. Clinic, C. 6 clicks functional measurement tool: why it's drawing crowds at conferences far and wide. 2017; Available from: <https://consultqd.clevelandclinic.org/6-clicks-functional-measurement-tool-why-its-drawing-crowds-at-conferences-far-and-wide/>.
 15. Mahan MC, et al. Mepivacaine spinal anesthesia facilitates rapid recovery in total knee arthroplasty compared to bupivacaine. *J Arthroplasty.* 2018;33(6):1699–704.
 16. Siddiqi A, Mahmoud Y, Secic M, Tozzi JM, Emara A, Piuizzi NS, et al. Mepivacaine Versus Bupivacaine Spinal Anesthesia for Primary Total Joint Arthroplasty: A Systematic Review and Meta-Analysis. *J Arthroplasty.* 2022;37(7):1396–1404.e5. <https://doi.org/10.1016/j.arth.2022.03.031>.
 17. Calkins TE, et al. Mepivacaine vs bupivacaine spinal anesthesia in total hip arthroplasty at an ambulatory surgery center. *J Arthroplasty.* 2021;36(11):3676–80.
 18. Lawrie CM, et al. Incidence and risk factors for postoperative urinary retention in total hip arthroplasty performed under spinal anesthesia. *J Arthroplasty.* 2017;32(12):3748–51.
 19. Herndon CL, et al. Lower dosing of bupivacaine spinal anesthesia is not associated with improved perioperative outcomes after total joint arthroplasty. *Arthroplast Today.* 2021;11:6–9.

Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

Ready to submit your research? Choose BMC and benefit from:

- fast, convenient online submission
- thorough peer review by experienced researchers in your field
- rapid publication on acceptance
- support for research data, including large and complex data types
- gold Open Access which fosters wider collaboration and increased citations
- maximum visibility for your research: over 100M website views per year

At BMC, research is always in progress.

Learn more biomedcentral.com/submissions

