

# **Original Research Paper**

## **Anaesthesiology**

# ADDITION OF INTRATHECAL DEXAMETHASONE TO BUPIVACAINE FOR SPINAL ANESTHESIA IN ORTHOPEDIC SURGERY

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ABSTRACT
Objectives: Spinal anesthesia provides profound nerve block that can be produced in a large part of the body by simple injection of a small amount of local anesthetic. But Intrathecal local anesthetics have limited duration. Different drugs have been used as additive to prolong spinal anesthesia. corticosteroids in prolonging the analgesic effects of local anesthetics in peripheral nerves is well documented. The purpose of this investigation was to determine whether the addition of dexamethasone to intrathecal bupivacaine would prolong the duration of sensory analgesia or not.

**Methods:** We conducted a randomized, prospective, double-blind, case-control, clinical trial. A total of 50 patients were scheduled for orthopedic surgery under spinal anesthesia. The patients were randomly allocated to receive 15 mg hyperbaric bupivacaine 0.5% with 2 cc normal saline (control group) or 15 mg hyperbaric bupivacaine 0.5% plus 8 mg dexamethasone (case group) intrathecally. The patients were evaluated for quality, quantity, and duration of block; blood pressure, heart rate, nausea, and vomiting or other complications.

#### Results:

There were no signification differences in sensory level, and onset time of the sensory block between two groups. Sensory block duration in the case group was  $118\pm10.68$  minutes and in the control group was  $88.44\pm8.37$  minutes which was significantly higher in the case group (P<0.001). The duration of analgesia was  $408.92\pm72.44$  minutes in the case group; whereas it was  $223\pm43.67$  minutes in the control group (P<0.001). The frequency of complications was not different between two groups.

## **KEYWORDS**:

Spinal anesthesia is the common block for lower abdomen and orthopedic surgery And avoids the risks of general anesthesia such as aspiration and difficulty with airway management.[1] Bupivacaine (H) is appropriate for procedures lasting up to 90-120 minutes.[1–3] To prolong duration of block, various additives such as adrenaline, phenylephrine, clonidine, opioids, etc. were added to Bupivacaine.[2–5] The additions of adrenaline cause tachycardia, pallor, and hypertension, which can be risky in patients with cardiovascular disease.[1] Intrathecal opioid administration has central and respiratory depression effects. Recently, some studies reported the effects of corticosteroids in quality and quantity of the sensory block in the peripheral nerves.[3–5]

Dexamethasone reduces inflammation and transmission of nociceptive C-fibers and by suppressing ectopic neural discharge.[6] It has been shown that the duration of postoperative analgesia was prolonged when dexamethasone is given as an adjunct for peripheral nerve blocks.[7,8] . The purpose of this investigation was to evaluate the effect of conjugation of dexamethasone with bupivacaine on the duration and onset time of spinal anesthesia.

## METHODS

Fifty patients with class American Society of Anesthesiologist (ASA) I-II, between 18 and 50 years old, scheduled for orthopedic surgery under spinal anesthesia, were included in this prospective, randomized, double-blind clinical trial. The orthopedic procedures were on lower limbs with surgery duration around 30-60 minutes. After ethics committee approval, written informed consent was obtained from each patient preoperatively.

Patients with history of steroid therapy, allergy to the drugs, hypertension, neurologic or psychological disorders, spine surgery, low back pain, opium addict or using any drug that modifies pain perception were excluded from the study.

Patients were randomly allocated into two groups, intrathecal bupivacaine- dexamethasone as the case group and intrathecal bupivacaine-normal saline as the control group.

After IV line preparation, a 6-8cc/kg lactated ringers solution was infused to all patients. Patients received no premedication, and upon arrival of patients into the operating room, ECG, peripheral oxygen saturation (SPO2), and noninvasive arterial blood pressure (NIBP) were monitored and recorded at 3-minute intervals until the end of surgery and vital signs were recorded every 15 minutes in the Post Anesthesia Care Unit (PACU).

Spinal anesthesia was performed in the sitting position at L3 -L4 level through a midline approach using a 25-gauge Quincke spinal needle. Patients of the control group received 15 mg (3 ml) of 0.5% hyperbaric bupivacaine diluted in preservative free normal saline (2 ml) and patients of the case group received 15 mg (3 ml) of 0.5% hyperbaric bupivacaine and 8 mg preservative free dexamethasone with the Dexadic brand name (2 ml), overall 5 ml volume intrathecally. To facilitate the double-blinding method, the medication was prepared and injected by an anesthesiologist who was not involved in the study. After performance of the spinal anesthesia patients were kept in supine position and oxygen 4 L min -1 was given through a face mask. The sensory block level was assessed by a pin prick test by a short bevel needle along the midaxillary line bilaterally. The sensory block level was controlled every 30 seconds for 20 minutes; then it was evaluated every 5 minutes until a 4 sensory level regression from highest level or to the end of the surgery. Onset time was defined from the time of injection of drugs into the intrathecal space to the peak of sensory and motor block and the duration of sensory block was defined from peak of sensory block up to 4 sensory level regressions or when the patients feel pain in the field of surgery.

Hypotension, a 20% decrease in systolic blood pressure from base line or systolic blood pressure <100 mm Hg and bradycardia, HR<50 beats/min was treated by IV mephentramine 6-18 mg plus crystalloid fluids; and IV atropine 0.5 mg respectively. Nausea and vomiting were also evaluated and were treated with 0.15 mg /kg IV metoclopramide.

After 4 dermatome block regression, pain assessment

intraoperatively or in PACU was done using the visual analogue pain scale (VAS) between 0-10 (0 = no pain, 10 = the most severe pain) every 1 hour. If the postoperative VAS was higher than 6, it was treated by morphine 2 mg IV. Patients were observed at the time of discharge from hospital and 1 month later and asked about any neurologic deficit.

The demographic data of patients were studied for each of the two groups. Continues covariates such as age, weight, height, and BMI were compared using the analysis of variance T-test. Onset time, sensory block duration, and duration of analgesia were analyzed by a T-test as appropriate, with the P value reported at the 95% confidence interval. For categorical covariates (sex, nausea/vomiting, hypotension, bradycardia, use of mephentramine, the use of atropine), the comparison was studied using a chi-squared test or Fisher's exact test. Sensory level compared by Mann-Whitney test. The significance level was defined as a P value less than 0.05. To calculate the sample size, a power analysis of  $\alpha$ =0.05 and  $\beta$ =0.80 showed that 25 patients per study group were needed to detect the difference between two groups.

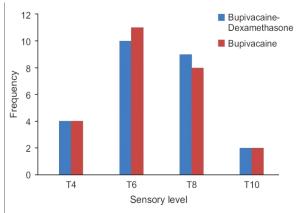
#### Go to: RESULTS

All patients (n=50) completed the study; there was no statistical difference in patients' demographics [Table 1]. The onset time of sensory block was 11.2±2.0 minutes for the case group and 10.9±1.8 minutes for the control Group (P=0.57). The maximum sensory level was between T4 and T10 in both groups and there was no significant difference (P=0.76) [Figure 1].

Table 1

Table 1					
	Bupivacaine		P value		
	Normal saline	Dexamethasone			
	n=25	n=25			
Age (years)	35.08±11.33	37.8±12.53	0.42		
Sex (F/M)	18.7	17.8	0.89		
Weight (kg)	76.84±8.42	76.28±10.01	0.73		
High (cm)	166.68±5.79	166.40±6.24	0.87		
BMI	27.71±3.38	27.48±2.84	0.8		





Sensory block level in the bupivacaine-dexamethasone versus bupivacaine-normal saline group The duration of the sensory block was 119.1±10.6 minutes in the case group and 89.4±8.3 minutes in the control group with a P value less than 0.001; also pain-free period in the case group was more than that in the control group(P<0.001). Receiving time to VAS >6 and the first analgesic dose prescription in the case group was significantly longer than that in the control group (P<0.001) [Table 2]. Hypotension was mild to moderate in both groups and was not different; except one patient in the control group who had a mean arterial pressure less than 60 mmHg and required 20 mg IV ephedrine to restore his blood pressure [Table 3].

Table 2

	Bupivacaine		P value		
	Dexamethasone n=25	Normal saline n=25			
Onset time (minutes)	11.27±2.08	10.95±1.87	P=0.57		
Duration of sensory block (minutes)	119.12±10.69	89.44±8.37	P<0.001		
Duration of pain-free period	401.92±72.64	202.24±43.67	P<0.001		

Comparison of onset time, duration of sensory block and pain free period between two groups

Table 3

	Bupivacaine		P value
	Normal saline	Dexamethasone	
Nausea and vomiting	5 (20)	2 (8)	0.41
Hypotension	7 (28)	7 (28)	1.00
Bradycardia	4 (16)	6 (24)	0.48
Shivering	8 (32)	9 (38)	0.76

Figures in parenthesis are in percentage

Incidence of adverse events between two groups during the study period Two patients in the case group and three patients in the control group complained of postdural puncture headache which was treated by hydration and simple analgesia. Other complications such as bradycardia, nausea, and vomiting were not different between the two groups [Table 3] and no neurologic deficit was observed in any patients.

## Go to: DISCUSSION

Results in this study showed that the addition of 8 mg dexamethasone with bupivacine for spinal block significantly prolonged sensory block and postoperative analgesia compared with intrathecal bupivacaine, without any effects on the onset time of sensory block in orthopedic surgery.

Many studies demonstrated analgesic effects of steroids in neuroaxial and peripheral block.[9–12] Movafegh et al. found that the addition of dexamethasone (8 mg) to lignocine for spinal anesthesia provided significant prolongation of sensory and motor block compared to plain lidocaine and there is no difference between dexamethasone-lidocaine 5% and epinephrine (0.2 mg) - lidocaine 5% in sensory and motor block duration.The onset time of sensory and motor blockade were similar among these three groups.[13]

The vasoconstriction effects of topical steroids are mediated by occupancy of classical glucocorticoid receptors.[14,15] In our study, dexamethasone produced a significant prolonged sensory block which can be explained by vasoconstriction mechanism, in contrast with traditional theory of steroid action; steroids bind to intracellular receptors and modulate nuclear transcription.[16]

Mirzaie et al. reported that corticosteroids with bupivacaine can reduce the incidence of back pain after laminectomy in postoperative period. [17] Kotani et al. repoted that methyl prednisolone with bupivacaine intrathecally in patients with postherpetic neuralgia has excellent and long-lasting analgesia. [18]

Taguchi et al. reported that intrathecal injection of betamethasone successfully decreased the pain score in three patients with intractable cancer pain[19] Another study reported that epidural dexamethasone (5 mg) reduces postoperative pain score and morphine consumption following laparoscopic cholecystectomy with no apparent side effects[20] Atsuhrio reported that intrathecal or epidural methylprednisolone decreased continuous pain and in patients of postherpetic neuralgia. They repoted more analgesia in

the intrathecal group compared to the epidural group.[21]

Steroids have anti inflammatory as well as analgesic property but the mechanism of the analgesia induced by corticosteroid is not fully understood. [22,23] Epidural steroids were used for back pain treatment. Intrathecal dexamethasone may influence intraspinal prostaglandin production. Acute noxious stimulation of peripheral tissues leads to sensitization of dorsal horn neurons of the spinal cord by the release of substances such as glutamate and aspartate. These amino acids activate N-methyl-D-Aspartate receptors resulting in calcium ion influx which leads to activation of phospholipase A2, which converts membrane phospholipase to arachidonic acid. Corticosteroids are capable of reducing prostaglandin synthesis by inhibition of phospholipase A2 through the production of calcium-dependent phospholipid binding proteins called annexins and by the inhibition of cyclooxygenases during inflammation. [24]

Some authors also believe that analgesic properties of corticosteroids are the results of their systemic effects. [25] The block prolonging effect may be due to its local action on nerve fibers. [26]

### CONCLUSION

In our study,we used the combination of bupivacaine and 8 mg dexamethasone intrathecally. We found that the addition of dexamethasone significantly prolongs the duration of sensory block and decreases opioid requirements in postoperative management. Further studies are needed to evaluate the optimal dose of dexamethasone to be used in spinal anesthesia.

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