COMPARISON OF LOW-DOSE BUPIVACAINE WITH FENTANYL AND BUPIVACAINE ALONE FOR SPINAL ANAESTHESIA FOR LOWER LIMB SURGERIES

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ABSTRACT

BACKGROUND

Bupivacaine is the widely used anaesthetic agent for spinal anaesthesia. Though, it has some advantages of producing good surgical anaesthesia and a longer half-life when compared to other local anaesthetics, the incidence of adverse effects on haemodynamic stability like hypotension was found to be more common. Adjuvants like opioids have been used in combination with bupivacaine to lower the dose of each agent and maintain the analgesic efficacy and thereby reducing the incidence and severity of adverse effects. Fentanyl, a lipophilic opioid, has rapid onset and offset of action.

The aim of the study is to compare the efficacy and the incidence of adverse effects between bupivacaine alone and lowdose bupivacaine with fentanyl as spinal anaesthesia among the patients undergoing lower limb surgeries.

MATERIALS AND METHODS

A prospective longitudinal study was conducted for a period of one year in the Anaesthesiology Department at Vinayaka Mission Kirupananda Variyar Medical College Hospital. A total of 80 patients were included for the study. They were divided into two groups of 40 each, group H (bupivacaine 75 mg, n=40) and group L (bupivacaine 5 mg with 25 mg fentanyl, n=40). The patients were positioned in left lateral position and under sterile precautions 23G Quincke spinal needle was inserted between the L3 and L4 interspace, and depending on the patients allotted group, the anaesthetic agent was administered. Blood pressure, pulse rate, respiratory rate and saturation was recorded at 2 minute intervals for the first 10 minutes and then subsequently at 5 minutes interval. Adverse events such as nausea, vomiting, shivering, pruritus, respiratory depression and transient neurological symptoms if occurred were noted.

RESULTS

The maximum sensory level attainment was T9 in both the groups. The Bromage motor score was significantly higher in the fentanyl with low-dose bupivacaine group. The mean reduction of BP was higher among the patients who received high dose of bupivacaine and the difference was found to be statistically significant. No adverse events were reported in both the groups.

CONCLUSION

The combination of low-dose bupivacaine with fentanyl can be a preferred alternative for elective lower limb surgeries than a high dose of bupivacaine alone.

KEYWORDS

Bupivacaine, Fentanyl, Spinal Anaesthesia, Sensory and Motor Block.

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BACKGROUND

Spinal anaesthesia is the most commonly used technique for lower abdominal and lower extremity surgeries, because of its distinct advantages over general anaesthesia.

Bupivacaine is the widely used anaesthetic agent for spinal anaesthesia. Though, it has some advantages of

Financial or Other, Competing Interest: None. Submission 20-06-2017, Peer Review 25-06-2017, Acceptance 29-06-2017, Published 03-07-2017. Corresponding Author: Dr. Karthi Vellakalpatti Mani, Assistant Professor, Department of Anaesthesiology, IRT-PMCH, Perundurai, Erode. E-mail: dr.karthivm@gmail.com DOI: 10.18410/jebmh/2017/643 Tero Se producing good surgical anaesthesia and a longer half-life when compared to other local anaesthetics, but the incidence of adverse effects on haemodynamic stability like hypotension was found to be more common.¹ Perioperative hypotension may affect postoperative recovery and also the high incidence of coronary disease, increases risk of ischaemia secondary to hypotension.

In patients with history of hypertension, diabetes and coronary artery disease, it is essential to limit the extent to haemodynamic adverse effects. So, in these situations by using small dose of local anaesthetic agents, we can limit the extent of block, but bupivacaine solely in low dose will not be able to provide adequate surgical anaesthesia.²⁻⁵ Addition of opioids to local anaesthetic for spinal anaesthesia was first introduced in 1979 with intrathecal morphine. They

act on opioid receptors present in the substantia gelatinosa of dorsal horn of spinal cord to produce the anaesthetic effect.

Fentanyl, a lipophilic opioid, has rapid onset and offset of action. Adjuvants like opioids have been used in combination with bupivacaine to lower the dose of each agent and maintain the analgesic efficacy and thereby reducing the incidence and severity of adverse effects.^{6,7} They have synergistic antinociceptive effects and the opioids have been shown to reduce the incidence of hypotension. Studies had shown that the addition of fentanyl to hyperbaric bupivacaine improves the quality of intraoperative and early postoperative subarachnoid block.8 The present study was designed to evaluate whether a low dose of bupivacaine produces adequate surgical anaesthesia with lesser haemodynamic side effects when combined with fentanyl.

Aim-

To compare the efficacy and the incidence of adverse effects between bupivacaine alone and low-dose bupivacaine with fentanyl as spinal anaesthesia among the patients undergoing lower limb surgeries.

MATERIALS AND METHODS

A prospective longitudinal study was conducted for a period of one year in the Anaesthesiology Department at Vinayaka Mission Kirupananda Variyar Medical College Hospital. The study was started after getting approval from the institutional ethical committee. All adult patients with ASA staging 1 and 2 and are undergoing elective lower limb surgery were included in the study. Emergency trauma patients and paediatric patients were excluded from the study. A total of 80 patients were included for the study. All 80 patients were randomly allocated into two groups of 40 each by sealed envelope technique to either the group H (bupivacaine 75 mg, n=40) or group L (bupivacaine 5 mg with 25 mg fentanyl, n=40). After a preanaesthetic evaluation, the procedure was explained to the patient and informed consent was obtained. All patients were premedicated with Tab. Diazepam 5 mg orally 2 hours prior to procedure. On arrival at the operation theatre, the patients were positioned in left lateral position and under sterile precautions 23G Quincke spinal needle was inserted between the L3 and L4 interspace and depending on the patients allotted group, group H patients received 75 mg bupivacaine and group L patients received 5 mg bupivacaine with 25 mg fentanyl. Henceforth, the data was collected by an investigator who was not aware of the drug administered. Onset of sensory level, peak sensory level and motor blockade was noted. Blood pressure, pulse rate, respiratory rate and saturation was recorded at 2 minute intervals for the first 10 minutes and then subsequently at 5 minutes interval. Adverse events such as nausea, vomiting, shivering, pruritus, respiratory depression and transient neurological symptoms if occurred were noted.

All data were entered and analysed by using SPSS version 21. Intergroup difference among the parametric

variables was analysed using Student's t-test and among the nonparametric variables Chi-square test was used to assess the statistical significance. P value <0.05 was considered as statistically significant.

RESULTS

Table 1 shows the demographic characteristics along with weight and height comparison between the two groups. It is seen from the table that all the parameters were almost similar in both the groups without showing any statistical significant difference between the two groups. Similarly, the haemodynamic parameters like pulse rate, blood pressure, respiratory rate and oxygen saturation, which were measured at the baseline (before spinal anaesthesia) were almost similar in both the groups without showing statistical difference (Table 2). After the spinal anaesthesia, among the various vital parameters, which were measured, the maximum sensory level attainment was T9 in both the groups, which proves that there is no change in attaining sensory level between the two groups (Figure 2), whereas the motor score (score 4) was higher among the group, which received low dose of bupivacaine along with fentanyl (group L) than the group, which received high dose of bupivacaine alone (group H), which was assessed through Bromage scoring system and the difference between them was found to be statistically significant (Figure 1). Though the motor score was higher among the patients who received low dose of bupivacaine with fentanyl, none of the patients required general anaesthesia and only required some additional dose of sedative like midazolam and none of the patients in either group required vasopressor or atropine. The haemodynamic response was better among the patients in group L, which was shown by mean reduction in the blood pressure. The mean reduction of BP was higher among the patients who received high dose of bupivacaine and the difference was found to be statistically significant (Figure 3), whereas the heart rate (Figure 4) and respiratory rate did not show any statistical significant difference between the two groups. No adverse events like nausea, vomiting and pruritus were reported in any of the group (Table 3).

Parameters	Group H Group L (n=40) (n=40)		P value			
Age in years (mean ± SD)	55 ± 10.4	52.2 ± 10.9	0.241*			
Gender M:F ratio	31:9	28:12	0.510**			
Weight in kg (mean ± SD)	60.7 ± 8.8	59.4 ± 8.1	0.504*			
Height in cm (mean ± SD)	161.9 ± 1.1	161.1 ±1.2	0.661*			
Table 1. Demographic Characteristics Among the Study Population						

* p value derived by using Student's T-test.

**p value derived by using Chi-square test.

Parameters	Group H (n=40)	Group L (n=40)	P Value (Derived by Applying Student's t-Test)			
Systolic BP (mmHg)	18.9 ± 14.2	129.4 ± 12.4	0.887			
Diastolic BP (mmHg)	75.0 ± 11.8	76.5 ± 10.0	0.538			
Mean BP (mmHg)	89.1 ± 10.7	92.4 ± 10.3	0.167			
Pulse rate (beats/mins.)	84.8 ± 13.0	81.4 ± 15.0	0.295			
Respiratory rate (breaths/mins.)	17.5 ± 2.2	17.6 ± 1.8	0.812			
Saturation	99.78 ± 0.6	99.97 ± 0.1	0.625			
Table 2. Pre-Anaesthetic Haemodynamic Parameters between the Two Groups						

Parameters	Group H (n=40)	Group L (n=40)	Significance			
Maximum sensory level (median and range)	T9 (T8-T10)	T9 (T8-T11)	0.643*			
Modified Bromage score (median and range)	2 (2-4)	4 (4-6)	0.000*			
Maximum fall of systolic BP from baseline in mmHg, mean \pm SD	7.5 ± 2.3	2 ± 0.5	0.003**			
Maximum fall of diastolic BP from baseline in mmHg, mean ± SD	6.9 ± 1.8	1.8 ± 0.8	0.0028**			
Mephentermine used	Nil	Nil	-			
Atropine used	Nil	Nil	-			
Supplement with midazolam (number of patients)	3/40 (7.5%)	5/40 (12.5%)	0.091*			
Supplement with GA	Nil	Nil	-			
Maximum fall in respiratory rate (mean \pm SD)	1.7 ± 1.2	1.6 ± 1.3	0.673**			
Adverse events reported	Nil	Nil	-			
Table 3. Comparison of Vital Parameters Measured after Spinal Anaesthesia between the Two Groups						

*p value derived by Chi-square test.

**p value derived by Student's t-test.



Figure 1. Comparison of Modified Bromage Score between the Two Groups



Figure 2. Comparison of Sensory Level Block between the Two Groups



Figure 3. Comparison of Change in Mean BP among the Two Groups



Figure 4. Comparison of Change in Heart Rate among the Two Groups

DISCUSSION

This study had shown that addition of fentanyl to low dose of bupivacaine provided the same level of anaesthesia as that of higher dose of bupivacaine given alone. It is known that local anaesthetic agents cause fall in blood pressure due to the sympathetic blockade. Fentanyl has no effect on the sympathetic nerves, and by means of its synergism with local anaesthetic agents, it gives the same level of anaesthesia as equivalent to higher dose of bupivacaine.^{9,10}

The effect of decreasing the dose of bupivacaine usually causes a reduction in the density of blockade as proven by other studies.^{5,6} In the present study, it was shown that patients in the fentanyl group had a significantly lesser degree of motor blockade and especially one patient in the fentanyl group had the modified Bromage score of 6, which indicates absolutely no motor blockade, whereas all other sensory parameters were observed to be preserved. Few patients could perceive touch and pressure. Proprioception was also preserved in these cases. Some patients were

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trying to move their leg as they feel the limb to be numb and heavy and few patients were able to wriggle their toes as the patients were not able to perceive the movement. So, it resulted in giving additional dose of sedation intraoperatively. Anxious patients might be unsuitable for this drug combination considering the amount of stress the patient is put in when he feels the surgeon handling his limb. Patient cooperation is also required when the density of blockade is so low unlike when a conventional dose is used. Moreover, this dose combination might not be effective when muscle relaxation is required.

The baricity of the solution does not make any difference to the level of blockade or the density of blockade as shown by a study done by Roy G Soto et al.¹¹ So, the decrease in the density of blockade is not explained by the decrease in density of the drug administered. It is explained only by the lower dose of bupivacaine used in the group in which fentanyl was added to make equal volume.

The sensory level as assessed by pinprick was the same in both the groups. All patients had adequate analgesia and supplemental analgesics or conversion to general anaesthesia was not required in any of the cases.

Blood pressure was stable during the entire course of surgery in all patients. Though, the fall in blood pressure was significantly greater in the high dose of bupivacaine than the fentanyl combination group, it did not cause severe hypotension and the results of our study was almost in par with the previous studies, which had shown a significant decrease in blood pressure among the group, which received high dose of bupivacaine, but in those studies, the patients had developed hypotension.^{5,6,12,13} Blockade of two sympathetic segments might not cause hypotension, 14, 15, 16 which might be the cause for no hypotension reported in our study. Furthermore, all the patients selected for this study were classified as ASA I or II physical status. All of them had received an adequate preload before spinal anaesthesia was administered. Patients with a stable autonomic nervous system who have received adequate preloading and when only two sympathetic levels blocked can have a stable blood pressure.¹⁵ Moreover in the studies which had reported hypotension, the level of blockade was higher and the subjects chosen were elderly patients. This drug combination is proposed to be of use in patients who are haemodynamically unstable or have autonomic instability when a level of T10 itself might cause profound fall in blood pressure, it being dependent completely on the sympathetic nervous system in these patients.

The onset and duration of blockade was also similar between the groups. Patient related factors like height, weight and age seems to have a better correlation to the duration of blockade than addition of fentanyl to bupivacaine, although some studies have shown a difference in the duration of blockade age stratified comparison between the groups have not been done, which could have shown a different result.¹⁷⁻²⁰

There were no adverse effects observed with the addition of fentanyl in our study population. Pruritus found to be a frequent adverse effect of intrathecal opioids and

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previous studies had shown the incidence of pruritus to be between 20 to 70% in their study subjects.²¹⁻²² However, in our study, none of the patients getting fentanyl complained of pruritus. Respiratory depression was not observed in any of our patients receiving fentanyl. This is in agreement with previous studies.²³⁻²⁵ Because of its high lipid solubility, unlike morphine, it does not ascend cephalad to reach the respiratory centre to cause respiratory depression. Similarly, nausea, vomiting or urinary retention were not observed in any of our patients, whereas few other studies had reported these adverse effects.²⁶⁻²⁷ This study shows that fentanyl when added to bupivacaine provides the same level of anaesthesia as a higher dose of bupivacaine and this combination can be especially useful for patients having ischaemic heart disease, diabetics with end organ damage, renal failure without coagulopathy and in patients with autonomic neuropathy. It can be recommended only when the risk of using a higher dose of bupivacaine or giving general anaesthesia outweighs the minimal discomfort associated with this dose.

CONCLUSION

Thus, we conclude that adding fentanyl helps in reducing the dose of 0.5% hyperbaric bupivacaine for spinal anaesthesia in lower limb surgeries without showing any change in the sensory level block, but showing a minimal change in the motor level block. By its synergistic effect with 0.5% hyperbaric bupivacaine, it provides better intraoperative and postoperative analgesia, good haemodynamic stability with no incidence of complications like nausea, vomiting and shivering. Therefore, the combination of low-dose bupivacaine with fentanyl should be preferred alternative for elective lower limb surgeries than a high dose of bupivacaine alone.

REFERENCES

- Kararmaz A, Kaya S, Turhanoglu S, et al. Low-dose bupivacaine-fentanyl spinal anaesthesia for transurethral prostatectomy. Anaesthesia 2003;58(6):526-530.
- [2] Khanna MS, Singh IKJP. Comparative evaluation of bupivacaine plain versus bupivacaine with fentanyl in spinal anaesthesia in geriatric patients. Indian J Anaesth 2002;46(3):199-203.
- [3] Goel S, Bharadwaj N, Gower VK. Intrathecal fentanyl added to intrathecal bupivacaine for day case surgery: a randomized study. Eur J Anaesthesiol 2003;20(4):294-297.
- [4] Atallah MM, Shorrab AA, Mageed AYM, et al. Low-dose bupivacaine spinal anaesthesia for percutaneous nephrolithotomy: the suitability and impact of adding intrathecal fentanyl. Acta Anaesthsiol Scand 2006;50(7):798-803.
- [5] Korhonen AM, Valanne JV, Jokela RM, et al. Intrathecal hyperbaric bupivacaine 3 mg+ fentanyl 10 mg for outpatient knee arthroscopy with tourniquet. Acta Anaesthesiol Scand 2003;47:342-346.

- [6] Singh H, Yang J, Thornton K, et al. Intrathecal fentanyl prolongs sensory bupivacaine spinal block. Can J Anaesth 1995;42(11):987-991.
- [7] Gupta A, Axelsson K, Thorn SE, et al. Low-dose bupivacaine plus fentanyl for spinal anaesthesia during ambulatory inguinal herniorrhaphy: a comparison between 6 mg and 7.5 mg of bupivacaine. Acta Anaesthsiol Scand 2003;47(1):13-19.
- [8] Al-Ghanem SM, Massad IM, Al-Mustafa MM, et al. Effect of adding dexmedetomidine versus fentanyl to intrathecal bupivacacine on spinal block characteristics in gynecological procedures: a double blind controlled study. American Journal of Applied Sciences 2009;6(5):882-887.
- [9] Tejwani GA, Rattan AK, McDonald JS. Role of spinal opioid receptors in the antinociceptive interactions between intrathecal morphine and bupivacaine. Anaesth Analg 1992;74(5):726-734.
- [10] Wang C, Chakrabarti MK, Whitwam JG. Specific enhancement by fentanyl of the effects of intrathecal bupivacaine on nociceptive afferent but not on sympathetic efferent pathways in dogs. Anaesthesiology 1993;79(4):766-773.
- [11] Soto RG, Paez JC, Smith RA. Impact of baricity of bupivacaine on intrathecal fentanyl-associated pruritus during combined spinal/epidural anesthesia for labor. The Internet Journal of Anaesthesiol 2009;20(1):1-6.
- [12] Kuusniemi KS, Pihlajamaki KK, Pitkanen MT, et al. The use of bupivacaine and fentanyl for spinal anaesthesia for urologic surgery. Anaesth Analg 2000;91(6):1452-1456.
- [13] Patra P, Kapoor MC, Nair TGM. Spinal anaesthesia with low dose bupivacaine and fentanyl for endoscopic urological surgeries. Journal of Anaesthetic and clinical pharmacology 2005;21(2):147-154.
- [14] Martyr JW, Clark MX. Hypotension in elderly patients undergoing spinal anaesthesia for repair of fractured neck of femur. A comparison of two different spinal solutions. Anaesth Intensive Care 2001;29(5):501-505.
- [15] Coe AJ, Revanas B. Is crystalloid preloading useful in spinal anaesthesia in the elderly? Anaesthesia 1990;45(3):241-243.
- [16] Choi DH, Ahn HJ, Kim MH. Bupivacaine-sparing effect of fentanyl in spinal anesthesia for cesarean delivery. Reg Anesth Pain Med 2000;25(3):240-245.
- [17] Patterson L, Avery N, Chan P, et al. The addition of fentanyl does not alter the extent of spread of intrathecal isobaric bupivacaine in clinical practice. Can J Anaesth 2001;48(8):768-772.
- [18] Casati A, Fanelli G, Aldegheri G, et al. Frequency of hypotension during conventional or asymmetric hyperbaric spinal block. Reg Anesth Pain Med 1999;24(3):214-219.
- [19] Infante KNE, Van Gessel E, Forster A, et al. Extent of hyperbaric spinal anaesthesia influences the duration of spinal block. Anaesthesiology 2000;92(5):1319-1323.
- [20] Hunt CO, Naulty JS, Bader AM, et al. Perioperative analgesia with subarachnoid fentanyl-bupivacaine for

cesarean delivery. Anaesthesiology 1989;71(4):535-540.

- [21] Roussel JR, Heindel L. Effects of intrathecal fentanyl on duration of bupivacaine spinal blockade for outpatient knee arthroscopy. AANA J 1999;67(4):337-343.
- [22] Mulroy MF, Larkin KL, Siddiqui A. Intrathecal fentanylinduced pruritus is more severe in combination with procaine than with lidocaine or bupivacaine. Reg Anesth Pain Med 2001;26(3):252-256.
- [23] Leighton BL, DeSimone CA, Norris MC, et al. Intrathecal narcotics for labor revisited: the combination of fentanyl and morphine intrathecally provides rapid onset of profound, prolonged analgesia. Anaesth Analg 1989;69(1):122-125.
- [24] Maves TJ, Gebhart GF. Antinociceptive synergy between intrathecal morphine and lidocaine during visceral and somatic nociception in the rat. Anesthesiology 1992;76(1):91-99.
- [25] Vercauteren MP, Coppejans HC, Hoffmann VL, et al. Small-dose hyperbaric versus plain bupivacaine during spinal anesthesia for cesarean section. Anesth Analg 1998;86(5):989-993.
- [26] Tuominen M, Taivainen T, Rosenberg PH. Spread of spinal anesthesia with plain 0.5% bupivacaine: influence of the vertebral interspace used for injection. British Journal of Anaesthesia 1989;62(4):358-361.
- [27] Fernandez-Galinski D, Rue M, Moral V, et al. Spinal anaesthesia with bupivacaine and fentanyl in geriatric patients. Anesth Analg 1996;83(3):537-541.