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A comparative study to assess the efficacy of addition of dexmedetomidine to levobupivacaine in brachial plexus block

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ABSTRACT

Background: Brachial plexus block is one of the best surgical technique for upper limb surgeries. It is a good anesthetic and analgesic agent. Though levobupivacaine has a duration of action of three to eight hours it has an acceptable analgesic property and few other practical constraints. Addition of vasoconstrictors like α -adrenergic agonists, hyaluronidase, neostigmine, opioids have been found to be effective. Dexmedetomidine, an $\alpha 2$ receptor agonist has found to be a better adjuvant to levobupivacaine by few literatures.

Objectives: To assess the effect of levobupivacaine 0.5% alone and with dexmedetomidine $100~\mu g$ as an adjuvant to levobupivacaine 0.5% on the onset and duration of sensory and motor block, the duration of perioperative analgesia, complications and sedation score.

Materials and Methods: This study was conducted after ethical committee clearance in a tertiary care medical college hospital in northern Kerala, India. 60 patients between the age of 18 and 60 years planned for upper limb surgery under brachial plexus block, who fulfilled the inclusion and exclusion criteria were included in the study. A pre anesthetic checkup was done to assess the patient before including in the study. The patients who were included in the study were randomly assigned into two groups – one group received 1 ml ($100\mu g$) dexmedetomidine with 39 ml of 0.5% Levobupivacaine(LD group) and the second group received 1 ml of 0.9% normal saline and 39 ml of 0.5% Levobupivacaine(LS group) as anesthetic agent through sealed envelope technique. The clinical history, vitals and hemodynamic parameters were monitored and noted prior to and during the surgery. The data collected was analyzed and tabulated.

Results: There was no statistically significant difference between both groups with respect to age, Gender and weight signifying proper random allocation of study subjects. None in LD group required post operative analgesia. The mean duration of onset of sensory and motor block was earlier and duration of motor and sensory analgesia was prolonged with the addition of Dexmedetomidine to Levobupivacaine. The hemodynamic stability was better with Dexmedetomidine. Bradycardia was reported in 20% of study participants who were administered dexmedetomidine

Conclusion: Dexmedetomidine added to levobupivacaine provides better anesthetic and analgesic care in upper limb surgeries done using brachial plexus block.

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1. Introduction

The effectiveness of Brachial plexus block had been proved as the best for a long period for the surgeries in upper limb. ¹ It is found to be one of the efficient methods for emergency

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and routine upper limb surgeries. It provides very good intraoperative anesthesia as well as postoperative analgesia without any systemic side effects. The complications like side effects of anesthetic agents, stress of tracheal intubation and laryngoscopy are avoided in this type of block and therefore commonly used in regional nerve block of upper extremity. Post operatively, patients are free from edema,

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pain, cerebral depression, nausea, vomiting and vasospasm

Levobupivacaine has the duration of action ranging from three to eight hours and hence it is the most commonly used local anesthetic drug.² This anesthetic drug has established itself as cheaper and safer for many years.³ But patchy or incomplete analgesia and delayed onset are few of the practical constraints. Few drugs are added to levobupivacaine in trial to decrease these limitations, to upsurge the quality, prolong the duration of action and analgesia.

To prolong the duration of brachial plexus block and improve the quality, vasoconstrictors like α -adrenergic agonists, hyaluronidase, neostigmine, opioids, can be utilized. But these vasoconstrictors have been found to cause certain side effects. There arise the need to identify the ideal additive and many researchers tried the novel alpha 2 adrenergic agonists.

Alpha 2 adrenergic agonists not only decrease the requirements of intraoperative anesthetic agents, but also they have cardiovascular stabilizing properties, sympatholytic analgesic and sedative property. In order to reduce the time of onset of nerve block, to prolong the duration of block and to improve the quality of blockade, these can be given in peripheral nerve blocks, intrathecal, epidural either alone or with local anesthetic agents. ^{2,4}

An $\alpha 2$ receptor agonist, dexmedetomidine, is eight times more sensitive than clonidine. ^{5,6} Researches have found that dexmedetomidine, when used in many animals and humans had improved the onset and duration of motor / sensory blockade. When used as an adjuvant to local anesthetic agents in peripheral nerve blocks, it has increased the duration of analgesia. ⁷⁻¹⁴

The present study was designed to assess the effect of levobupivacaine 0.5% alone and with dexmedetomidine $100\mu g$ as an adjuvant to levobupivacaine 0.5% on the onset and duration of sensory and motor block, the duration of perioperative analgesia, complications and sedation score.

2. Materials and Methods

This cross sectional comparative study was carried out from November 2012 to November 2013 on sixty patients in a tertiary care medical college in northern part of Kerala, South India after getting the institutional human ethical committee concurrence. Patients posted electively for upper limb orthopedic and soft tissue lesion, between the age of 18 and 60 years of any gender, weighing above 60 kilograms with ASA Grade I and II and those who fulfilled the selection criteria and those who gave consent to participate in the study were included in the study. Patients with chronic kidney disease, hypertension, pregnant women, uncontrolled diabetes mellitus, cerebrovascular accident, COPD, coronary artery disease on anticoagulants, those with history of bleeding disorders, and those who were allergic to amide local anesthetics /alpha 2 agonist were

excluded from the study.

A pre-anesthetic checkup including complete history, general and systemic examination and fitness was assessed for all patients. Complete hemogram like RBC, WBC counts, platelets and other tests like blood urea, random blood sugar, bleeding time, clotting time, serum creatinine and ECG is mandatory if the subject is more than 45yrs. Patients were informed to be on nil per oral after 8 pm on the previous day of surgery. The participants were divided randomly into two groups of thirty each using sealed envelope technique.

The first group (LD group) was administered 1 ml $(100\mu g)$ dexmedetomidine with 39 ml of 0.5% Levobupivacaine. The second group was given 1 ml of 0.9% normal saline and 39 ml of 0.5% Levobupivacaine as anesthetic agent. Once the study participant entered the operation theatre, the chief consultant had used the pre assigned 30 envelope for each group in the pre shuffled order. Neither the investigator nor the patient knew which group they were assigned (double blinding).

The vital signs like respiratory rate, heart rate, systolic and diastolic blood pressure, pulse rate and oxygen saturation were monitored and noted immediately on entering the OT. Ringer lactate was started in the already secured intravenous line. Brachial plexus block was applied through the supraclavicular approach.

Nerve locator (Fisher and Paykel, New Zealand) was used for neural localization and it was achieved by connecting to a 22 G, 50-mm-long stimulating needle (Stimuplex, Braun, Germany). The location end point was a distal motor response with an output lower than 0.5 milliamperes in the median nerve region. Local anaesthetic solution in the labelled coded syringe was injected following negative aspiration.

Pin prick method was used to assess the sensory block. Sensory onset was considered when there was dull sensation to pin prick along the distribution of any two of the following nerves like musculocutaneous nerve, radial nerve, median nerve, ulnar nerve. When there was complete loss of sensation to pin prick, we can consider it as complete sensory block. Modified Bromage scale. Motor and sensory blocks were assessed for 30 minutes for every 3 minutes until after injection, and then every 30 minute until they have resolved. ¹⁵

Saturation of oxygen(SpO₂), heart rate, diastolic blood pressure, systolic blood pressure, were noted at 0, five, ten, fifteen, Thirty, sixty, ninety and one twenty minutes.

Side effects like heart rate less than fifty per min (bradycardia) and blood pressure less than 20% with respect to resting conditions (hypotension) were treated with appropriate measures. Then we noted the period of motor and sensory blocks once the surgery is started.

When the subject's visual analogue score >5, a rescue analgesia like intramuscular diclofenac sodium 75mg (1.5

mg/kg) was administered. Ramsay Sedation Scale (RSS) was used to assess the sedation before the block and 15 min then after.

All the data noted was entered in Microsoft excel sheet and was double checked. SPSS 16 software was used to analyze the collected data. The categorical variables were tabulated as frequency and percentages. Continuous variables were presented as Mean \pm Standard deviation. Independent sample T- test was used to measure the association between the vitals at different times. Chi-square test was applied to assess the relationship between the categorical variables. P<0.05 was considered as statistically significant.

3. Results

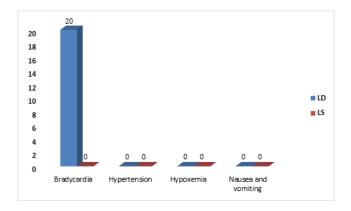


Fig. 1: Comparison of occurrence of side effects between both the groups

In the LD group males and females were 63.33% and 36.67% respectively. In LS group, it was 56.66% and 43.34% respectively. There was no statistical significance between the two groups. (Table 1)

The average age of study participants were 32.1 vs 30.3 in LD and LS group respectively. The mean weight of study participants in study and control group was 64.4 vs 63.2 respectively. Both the results were statistically significant (Table 2).

All the study participants in LD group did not require post-operative analgesia while all in LS group were given Postoperative analgesia. 26.67% in LD group were slightly drowsy compared to 100% in LS group. Both the results are statistically significant. (Table 1)

The mean duration of onset of sensory and motor block was nearly 4 min earlier in LD group compared to LS group.(5.32 vs 9.17 min) Onset of motor blockade was also 4 minutes earlier in LD group compared to LS (8.35 vs 12.28 min). The mean duration of sensory block(531 vs 921 min) and motor block(553 vs 943 min) were 390 minutes lesser respectively between both the groups. The duration post-operative analgesia was 402 min lesser between both

the groups (578 vs 980 min). All the results are statistically significant. (Table 2)

The diastolic blood pressure, systolic blood pressure, heart rate were comparatively maintained lesser than baseline for LD group from fifth min after induction of drug till two hours. (Tables 3, 4 and 5). Twenty percent of study population in LD group had bradycardia compared to none in LS group (chi-square value - 6.667; P value - 0.009). Bradycardia was noted in 20% if the study population. No other adverse effects was recorded in the study subjects.

4. Discussion

This cross-sectional study was done while applying supraclavicular brachial plexus block in upper limb surgeries to assess the impact of dexmedetomidine as an adjuvant drug to levobupivacaine. There is no statistical significance in age, sex and weight between both the groups signifying both the groups were similar before the start of the study and the sampling methods were followed appropriately.

Dexmedetomidine 100 μ g when used as an adjuvant to levobupivacaine 0.5% reduces the onset of sensory and motor blockade, prolongs the analgesic effect of motor and sensory block. The mechanism of action of $\alpha 2$ agonists causing sedation and analgesia is not completely known, but it seems to have multifactorial. Centrally, $\alpha 2$ agonists inhibits the substance P release at dorsal root neuron in the pain pathway and activates α^2 adrenoceptors in the locus coeruleus and thereby causing analgesia and sedation. Peripherally, $\alpha 2$ agonists decrease the release of noradrenaline, produce analgesia and produces $\alpha 2$ receptorindependent restraints on nerve action potentials. Activation of cation current by hyperpolarization causes the peripheral action of dexmedetomidine. For subsequent firing the nerve will not return from hyperpolarized state to resting membrane state. 16-19 Studies done by various other authors have also showed that addition of dexmedetomidine reduces the onset time of motor and sensory block and prolongs the duration of postoperative analgesia. 15,20-24

Addition of dexmedetomidine improves the hemodynamic stability of the patients. Similar findings were obtained in the several other researches conducted world wide. ^{25–27}

Bradycardia was noted in one fifth of the patients who were administered dexmedetomidine. Which coincides with the findings of Talke et al. ²⁸

5. Conclusion

The onset of motor and sensory blockade was reduced when dexmedetomidine was used as an adjuvant to 0.5% levobupivacaine and it also prolongs the duration of analgesia. It provides a good intraoperative sedation as well and helps to reduce the postoperative analgesic requirement.

 Table 1: Association between various parameters between groups

Parameter	Subclassification of	LD group		LS group		Chi square	P value	
	the parameter	No	%	No	%	value	P value	
Gender	Male	19	63.33	17	56.66	3.787	0.051	
	Female	11	36.67	13	43.34	3.767	0.031	
Post operative analgesia requirement Sedation Score	Not needed	30	100	0	0	60	< 0.001	
	Needed	0	0	30	100	00	<0.001	
	Sleeping but arousable	22	73.33	0	0	34.74	< 0.001	
	Slightly drowsy	8	26.67	30	100	34.74	<0.001	

Table 2: Comparison of mean values of various parameters between both groups

Parameter	Group	Mean	SD	Mean diff	T value	P value
Age (years)	LS	30.3	11.1	1.80 0.595	0.505	0.554
Age (years)	LD	32.1	12.3		0.393	
Weight (Va)	LS	63.2	4.2	1.20	1.259	0.213
Weight (Kg)	LD	64.4	3.1	1.20		0.213
Onset of sensory block (min)	LS	9.17	0.889	3.820	15.464	< 0.001
Offset of sensory block (film)	LD	5.32	1.020	3.820		
Onset of motor block (min)	LS	12.28	1.06	3.93	9.66	< 0.001
Onset of motor block (mm)	LD	8.35	1.96	3.93	9.00	<0.001
Duration of sensory Block (min)	LS	921.72	37.328	390.72	-33.013	< 0.001
Duration of sensory Block (IIIII)	LD	531	53.0	390.72	-33.013	<0.001
Duration of Motor Block (min)	LS	943.14	34.68	389.31 -	-35.503	< 0.001
Duration of Motor Block (IIIII)	LD	553.83	49.037	369.31	-33.303	<0.001
Duration of Post Op Analgesia	LS	980.69	46.823	402.69	20.062	< 0.001
(min)	LD	578	53.989	402.09	-30.863	<0.001

Table 3: Comparison of heart rates between both groups at different time intervals

Time (Min)	LS Group		LD Group		M 1:66	T l	D1
	Mean	SD	Mean	SD	Mean diff	T value	P value
0	85.21	8.01	84.92	8.32	0.29	0.138	0.891
5	83.56	7.51	79.56	6.68	4	2.18	0.033
10	81.88	7.51	76.85	6.54	5.03	2.752	0.008
15	79.32	7.65	72.35	6.72	6.97	3.749	< 0.001
30	76.65	7.85	69.80	6.23	6.85	3.711	< 0.001
45	75.15	6.8	67.32	6.12	7.83	4.669	< 0.001
60	75.12	6.23	67.15	5.82	7.97	5.120	< 0.001
75	78.35	8.89	71.15	6.88	7.2	3.508	< 0.001
90	79.82	7.86	71.59	7.02	8.23	4.277	< 0.001
120	82.01	8.35	73.25	7.89	8.76	4.177	< 0.001

Table 4: Comparison of mean values of systolic BP between both groups

Time (Min)	LS Group		LD Group		M 1*66	m1	D
	Mean	SD	Mean	SD	Mean diff	T value	P value
0	131.12	9.32	134.68	9.67	3.56	-1.044	0.301
5	126.32	5.81	124.36	5.32	1.96	1.363	0.178
10	121.47	6.25	115.89	6.67	5.58	3.344	< 0.001
15	122.12	5.98	114.21	5.26	7.91	5.395	< 0.001
30	120.10	5.54	113.95	5.98	6.15	4.146	< 0.001
45	121.15	6.23	112.22	5.65	8.93	5.816	< 0.001
60	122.25	8.13	114.26	6.65	7.99	4.167	< 0.001
75	125.51	9.69	114.55	7.86	10.96	4.811	< 0.001
90	126.71	9.36	115.63	8.02	11.08	4.728	< 0.001
120	128.32	9.12	117.21	9.11	11.11	4.721	< 0.001

LS Group LD Group Time (Min) Mean diff T value P value Mean SD Mena SD 0 0.724 0.472 9.21 79.68 1.64 81.32 8.32 5 79.25 8.32 76.85 7.68 2.4 1.161 0.25 10 71.32 5.68 4.03 2.598 0.012 75.35 6.32 15 73.22 6.32 69.52 5.82 3.7 2.359 0.022 30 73.52 68.12 5.37 5.4 3.613 < 0.001 6.18 6.23 5.29 45 73.11 67.82 5.92 3.378 < 0.001 60 74.73 7.12 67.25 5.85 7.48 4.446 < 0.001 75 75.63 7.36 67.03 6.20 8.6 4.895 < 0.001 90 77.25 8.59 71.23 6.02 2.791 < 0.001 8.11 79.26 9.24 72.26 8.36 7 3.077 < 0.001 120

Table 5: Comparison of mean values of diastolic BP between both groups

However dexmedetomidine may cause bradycardia in limited cases, which is reversible with atropine. Hence we conclude from the study dexmedetomidine can be an ideal adjuvant for levobupivacaine for upper limb surgeries applying brachial plexus block.

6. Source of Funding

None.

7. Conflict of Interest

None.

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