To Determine the Effect of Clonidine as an Adjuvant for Ropivacaine in Lower Limb Orthopedic Surgery

Amrata Asthana¹, JP Tiwari², DK Tripathi³, Bhaskar Prasad²

¹JR-3, Dr. Ram Manohar Lohia Institute of Medical Sciences, Lucknow, ²Consultant Anaesthesiologist, Dr. Ram Manohar Lohia Institute of Medical Sciences, Lucknow, ³Senior Consultant Anaesthesiologist, Dr. Ram Manohar Lohia Institute of Medical Sciences, Lucknow

Abstract

Background: The present study determined the effect of Clonidine as an adjuvant for ropivacaine in lower limb orthopedic surgery. Subjects and Methods: The present study was conducted o 60 patient's undergoing lower limb surgery. Group I patients were given 30 mL 0.75% ropivacaine alone for sciatic femoral block with Group II patients in whom sciatic femoral block will be achieved using 30 mL 0.75% Ropivacaine plus 1 μg/kg clonidine. The time for the first rescue analgesia in the post-operative period, SBP, DBP sedation was assessed by a modified Ramsay sedation scale. Results: Group I comprised of 16 males and 14 females and group II had 18 males and 12 females. The difference was non- significant (P> 0.05). ASA grade I was seen 18 patients in group I and 24 in group II, ASA II was seen in 12 in group I and 6 in group II patients. The difference was non- significant (P> 0.05). Intra- operative variations in the mean arterial pressure (mmHg) between the cases of the two groups were significant at 5 min, 10 min,15 min & 60 min during the operation as P<0.05. Post-operative variations in the mean systolic blood pressure (mm of Hg) between the cases of the two groups was not significant at time intervals as P>0.05. The difference in the sedation score between the cases of the two groups was highly significant at time intervals of 1 hour, 2 hours, 3 hours, 8 hours and Immediately post-op as P<0.001 & it was significant at time interval of 6 hr as p<0.05. Conclusion: Authors found that clonidine added to ropivacaine 0.75% in an appropriate technique has low side effect profile with considerable therapeutic benefit and enhances the quality of combined sciatic femoral nerve block.

Keywords: Clonidine, Ropivacaine, Sedation.

Corresponding Author: Dr. JP Tiwari, B14, type 4. Dr. RMLIMS Hospital Block Campus, Vibhuti Khand Gomtinagar, Lucknow-226010

Received: April 2020 Accepted: April 2020

Introduction

Orthopaedics is the branch of surgery concerned with conditions involving the musculoskeletal System. Orthopedic surgeons use both surgical and nonsurgical means to treat musculoskeletal trauma, spine disease, sports injuries, degenerative diseases, infections, tumors, and congenital disorders. The femoral nerve is the largest branch of the lumbar plexus and arises from the ventral rami of the second, third and fourth lumbar nerves.^[1]

Femoral nerve block can be used to provide surgical anaesthesia, usually in combination with a sciatic nerve block. This may be advantageous in situations where it would be preferable to avoid general or neuraxial anaesthesia, knee joint, vessels and lies outside the femoral sheath. [2]

Contraindications for FNB are patient refusal, inflammation or infection over injection site, allergy to local anaesthetics, anticoagulation or bleeding disorders and pre-existing peripheral neuropathies etc.^[3] The sciatic nerve is so large, it can be blocked from several different locations along the lower extremity. Labat's sciatic nerve block is the classic approach, targeting the nerve in the gluteal region. Other sciatic nerve blocks include the anterior and lateral approaches, which allow the patient to remain in the supine

position, as well as the parasacral and prone approaches. Raj's subgluteal approach is performed in the supine position with the hip flexed. Sciatic nerve blocks require adequate set-up because this large nerve resists local anesthetic penetration, leading to longer block onset times. [4]

Ropivacaine is a long acting local anaesthetic agent belonging to amino amide group. It is structurally related to Bupivacaine, but it is a pure Senantiomer of 1-propyl-2c,6c-pipecoloxylidide, developed for the purpose of reducing cardio toxicity and improving motor and sensory blockade and even more motor-sparing than bupivacaine. [5]

Ropivacaine because of its less lipophylic and stereo selective properties has higher threshold for cardio toxicity and CNS toxicity. [5] It inhibits platelet aggregation and it inhibits the growth of staphylococcus aureus, E. Coli and pseudomonas aeruginosa. Clonidine is an imidazoline derivative having complex actions. It is a partial agonist with high affinity to alpha 2 receptors especially alpha 2A subtype. [6] The present study determined the effect of Clonidine as an adjuvant for ropivacaine in lower limb orthopedic surgery.

Subjects and Methods

This randomized controlled study was conducted in the

Asthana et al: Clonidine as an Adjuvant for Ropivacaine

department of Anaesthesiology & department of Orthopaedic Surgery, Ram Manohar Lohia Combined Hospital, Lucknow on 60 patients undergoing elective lower limb orthopedic surgery in age ranged 18-50 years.

Patients' data such as name, age, gender etc. was recorded. Patients were divided into 2 groups. Group I patients were given 30 mL 0.75% ropivacaine alone forsciatic femoral block with Group II patients in whom sciatic femoral block will be achieved using 30 mL 0.75% Ropivacaine plus 1 μ g/kg clonidine. The time for the first rescue analgesia in the post-operative period, SBP, DBP sedation was assessed by a modified Ramsay sedation scale. Postoperative pain was recorded for 24 hrs using visual analogue scale.

Modified Ramsay Sedation Scale

| Indication | Score |
|--|-------|
| Anxious, agitated, restless | 1 |
| Awake, cooperative, oriented, tranquil | 2 |
| Semiasleep but responds to commands | 3 |
| Asleep but responds briskly to glabellar tap or loud auditory stimulus | 4 |
| Asleep with sluggish or decreased response to | 5 |
| glabellar tap or loud auditory stimulus | |
| No response can be elicited | 6 |

Results were tabulated and subjected to statistical analysis. P value less than 0.05 was considered significant.

Results

Table 1: Distribution of patients

| Table 1. Distribution of patients | | | | | | | |
|-----------------------------------|---------|---------|---------|--|--|--|--|
| Variables | Group A | Group B | P Value | | | | |
| Male | 16 | 18 | 0.421 | | | | |
| Female | 14 | 12 | | | | | |
| Total | 30 | 30 | | | | | |

[Table 1] shows that group I comprised of 16 males and 14 females and group II had 18 males and 12 females. The difference was non-significant (P>0.05).

Table 2: ASA status

| Variables | Group A | Group B | P Value |
|-----------|---------|---------|---------|
| ASA I | 18 | 24 | 0.201 |
| ASA II | 12 | 6 | |
| Total | 30 | 30 | |

[Table 2] shows that ASA grade I was seen 18 patients in group I and 24 in group II, ASA II was seen in 12 in group I and 6 in group II patients. The difference was non-significant (P> 0.05).

Table 3: Assessment of intra- operative mean arterial pressure

| | operative mean arterial pressure | | | | | |
|-----------|----------------------------------|---------------|------|------|-------|---------|
| MAP | Group | oup A Group B | | В | T | P Value |
| (mm/Hg) | Mean | SD | Mean | SD | Value | |
| Base Line | 94.7 | 5.24 | 91.4 | 5.39 | 0.010 | 0.314 |
| 5 Min | 90.3 | 5.35 | 86.6 | 4.24 | 1.564 | 0.004 |
| 10 Min | 86.2 | 5.46 | 84.8 | 4.37 | 1.624 | 0.003 |
| 15 Min | 88.3 | 5.45 | 85.4 | 4.31 | 1.589 | 0.040 |
| 30 Min | 87.3 | 5.36 | 85.3 | 4.54 | 1.632 | 0.108 |
| 45 Min | 88.5 | 5.20 | 86.2 | 4.55 | 1.625 | 0.073 |
| 60 Min | 85.5 | 5.50 | 84.9 | 4.53 | 1.712 | 0.001 |
| 90 Min | 86.7 | 5.35 | 83.2 | 5.52 | 1.742 | 0.243 |
| 120 Min | 86.2 | 5.40 | 84.0 | 4.03 | 0.754 | 0.950 |

[Table 3] shows that MAP after 60 minutes of start of operation was 85.5 ± 5.35 & 83.2 ± 5.35 (mmHg) whereas the readings after 120 minutes fluctuated to 86.2 ± 5.40 & 84.0 ± 4.03 (mmHg) for the cases of groups A & B respectively. Intra- operative variations in the mean arterial pressure (mmHg) between the cases of the two groups were significant at 5 min, 10 min,15 min & 60 min during the operation as P<0.05.

Table 4: Post-operative systolic blood pressure

| Systolic | Group | Group A Group B | | В | T | P Value |
|-----------|-------|-----------------|-------|------|-------|---------|
| Blood | Mean | SD | Mean | SD | Value | |
| Pressure | | | | | | |
| (mm/hg) | | | | | | |
| Base Line | 125.5 | 7.14 | 123.0 | 7.76 | 1.457 | 0.133 |
| 1 hr | 116.9 | 7.50 | 111.6 | 7.77 | 1.921 | 0.076 |
| 2 hr | 114.3 | 7.47 | 111.3 | 7.10 | 1.502 | 0.064 |
| 3 hr | 118.0 | 7.92 | 116.8 | 6.56 | 1.510 | 0.059 |
| 4 hr | 117.2 | 7.64 | 114.1 | 7.99 | 1.168 | 0.274 |
| 6 hr | 118.2 | 7.91 | 119.1 | 8.03 | 0.500 | 0.644 |
| 8 hr | 118.8 | 8.20 | 119.8 | 8.01 | 0.524 | 0.610 |

[Table 4] shows that the mean SBP readings immediately post operatively were $125.5\pm7.14~\&~123.0\pm7.76$ (mm of Hg), after 4 hours $117.2~\pm7.64~\&~116.8\pm7.99$ (mm of Hg) and after 8 hours $118.8\pm8.20~\&~119.8\pm8.01$ respectively for the cases of the groups A & B. Post-operative variations in the mean systolic blood pressure (mm of Hg) between the cases of the two groups was not significant at time intervals as P>0.05.

Table 5: Post-operative diastolic blood pressure

| Diastolic | Group A | | Group B | | T | P |
|------------------------------|---------|------|---------|------|--------|-------|
| Blood Pressure (mm/hg) | Mean | SD | Mean | SD | Value | Value |
| Base Line | 75.2 | 5.59 | 75.8 | 4.91 | 0.821 | 0.664 |
| 1 hr | 71.4 | 5.63 | 72.4 | 5.52 | 2.854 | 0.172 |
| 2 hr | 73.4 | 5.52 | 72.8 | 5.04 | -0.040 | 0.664 |
| 3 hr | 81.3 | 5.52 | 72.1 | 5.08 | 1.410 | 0.141 |
| 4 hr | 71.5 | 5.77 | 74.1 | 5.03 | -0.574 | 0.413 |
| 6 hr | 71.6 | 5.52 | 73.2 | 4.41 | 2.011 | 0.093 |
| 8 hr | 71.4 | 5.59 | 74.1 | 5.03 | -0.511 | 0.931 |

[Table 5] shows that the base line DBP immediately post-operative was 77.8 ± 5.77 & 76.8 ± 4.91 (mmHg), after 4 hours post-operative was 72.4 ± 5.59 & 73.1 ± 5.03 (mmHg) and after 8 hours was 72.4 ± 5.59 & 73.1 ± 5.03 (mmHg) respectively in the cases of both groups A & B. Post-operative variations in the mean diastolic blood pressure (mmHg) between the cases of the two groups A & B was not significant at any time interval post operatively as P>0.05.

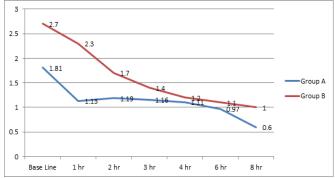


Figure 1: Sedation score

[Figure 1] shows that sedation score immediately post-operative was 1.81 ± 0.607 & 2.7 ± 0.36 , after a lapse of 1 hours score was 1.13 ± 0.438 & 2.3 ± 0.54 , after 4 hours score was 1.11 ± 0.303 & 1.2 ± 0.38 and after 8 hours post-operative score was 0.60 ± 0.490 & 1.0 ± 0.00 of the two groups A & B respectively. The difference in the sedation score between the cases of the two groups was highly significant at time intervals of 1 hour, 2 hours, 3 hours, 8 hours and Immediately post-op as P<0.001 & it was significant at time interval of 6 hr as p<0.05.

Discussion

Post-operative pain is the most common clinical problem in hospitals -among the surgical patients and is attributed as main reason for overnight hospital stay in 17-41% of surgical day care patients. Popivacaine with its efficacy, lower propensity for motor block and reduced potential for cardiotoxicity and central nervous system toxicity, appears to be an important option for regional anesthesia and management of postoperative pain. Popivacaine is a selective alpha 2 (α -2) adrenergic agonist with both analgesic and sedative properties. Plas use with ropivacaine in regional blocks has been associated with prolongation of the local anesthetic effect. The present study determine the effect of Clonidine as an adjuvant for ropivacaine in lower limb orthopedic surgery.

The patients were divided into two groups with only Ropivacine (Group A) & Ropivacine with Clonidine (Group B), each having 30 patients. Patients of group A received total 32 ml of ropivacaine 0.75% (12 ml for femoral block & 20 ml for sciatic block in each group). The distribution of patients was randomized in two groups using a computer generated program. Patients of group B received total 32 ml of ropivacaine 0.75% (12 ml for femoral block & 20 ml for sciatic block in each group) with Clonidine 1µgm/kg.

In present study, group I comprised of 16 males and 14 females and group II had 18 males and 12 females. ASA grade I was seen 18 patients in group I and 24 in group II, ASA II was seen in 12 in group I and 6 in group II patients. Anderson et al,[11] in which participants received bilateral ACBs containing 20 ml ropivacaine 0.5% + 1 ml clonidine 150µg/ml in one leg and 20 ml ropivacaine 0.5% + 1 ml saline in the other leg. There was no difference in duration of sensory block assessed with an alcohol swab: Mean duration in the leg receiving ropivacaine + clonidine was 19.4 hours compared to 19.3 hours in the leg receiving ropivacaine + placebo with a mean difference of 0.1 hour. No differences in block duration were detected when assessed by: Pinprick, mean difference 0.0 hour, maximum pain during tonic heat stimulation, mean difference -0.7 hour, warmth detection threshold, mean difference -0.1 hour or heat pain detection threshold.

We found that intra- operative variations in the mean arterial pressure (mmHg) between the cases of the two groups were significant at 5 min, 10 min,15 min & 60 min during the operation as P<0.05. Post-operative variations in the mean systolic blood pressure (mm of Hg) between the cases of the two groups was not significant at time intervals as P>0.05. Post-operative variations in the mean diastolic blood pressure (mmHg) between the cases of the two groups A & B was not

significant at any time interval post operatively as P>0.05. Pratap et al, [12] performed a bilateral trial on healthy volunteers resembling ours. They injected 0.5 ml of lidocaine 0.5% with 10 µg clonidine subcutaneously on one forearm and 0.5 ml of lidocaine 0.5% with saline subcutaneously on the opposite forearm. The duration of anesthesia was prolonged from 3.5 hours to at least 6 hours when clonidine was added. They assumed that 10 µg of clonidine would be without systemic effects and concluded that clonidine had a peripheral action in enhancing duration of anesthesia on superficial co-infiltration with lidocaine. However, $\alpha 2$ -receptors are more numerous at peripheral nerve endings compared to axons which may explain the difference in results.

We found that sedation score immediately post-operative was 1.81 ± 0.607 & 2.7 ± 0.36 , after a lapse of 1 hours score was 1.13 ± 0.438 & 2.3 ± 0.54 , after 4 hours score was 1.11 ± 0.303 & 1.2 ± 0.38 and after 8 hours post-operative score was 0.60 ± 0.490 & 1.0 ± 0.00 of the two groups A & B respectively. The difference in the sedation score between the cases of the two groups was highly significant at time intervals of 1 hour, 2 hours, 3 hours, 8 hours and Immediately post-op as P<0.001 & it was significant at time interval of 6 hr as p<0.05.

Conclusion

Authors found that clonidine added to ropivacaine 0.75% in an appropriate technique has low side effect profile with considerable therapeutic benefit and enhances the quality of combined sciatic femoral nerve block.

References

- Helayel PE, Kroth L, Boos GL, Jahns MT, Oliveira Filho GR. [Effects of intramuscular and perineural clonidine on sciatic nerve block with 0.5% ropivacaine.]. Revista brasileira de anestesiologia. 2005;55(5):483–90.
- Dalle C, Schneider M, Clergue F, Bretton C, Jirounek P. Inhibition of the I(h) current in isolated peripheral nerve: a novel mode of peripheral antinociception? Muscle & nerve. 2001;24(2):254–61.
- 3. Liu SS, Strodtbeck WM, Richman JM, Wu CL. A comparison of regional versus general anesthesia for ambulatory anesthesia: a meta-analysis of randomized controlled trials. Anesthesia and analgesia. 2005;101(6):1634–42.
- Capdevila X, Ponrouch M, Choquet O. Continuous peripheral nerve blocks in clinical practice. Current opinion in anaesthesiology. 2008;21(5):619–23.
- Ilfeld BM. Continuous peripheral nerve blocks: a review of the published evidence. Anesthesia and analgesia. 2011;113(4):904–25.
- Abdallah FW, Abrishami A, Brull R. The facilitatory effects of intravenous dexmedetomidine on the duration of spinal anesthesia: a systematic review and meta-analysis. Anesthesia and analgesia. 2013;117(1):271–8.
- Albrecht E, Kern C, Kirkham KR. A systematic review and meta-analysis of perineural dexamethasone for peripheral nerve blocks. Anaesthesia. 2015;70(1):71–83.
- Popping DM, Elia N, Marret E, Wenk M, Tramer MR. Clonidine as an adjuvant to local anesthetics for peripheral nerve and plexus blocks: a meta-analysis of randomized trials. Anesthesiology. 2009;111(2):406– 15.
- El Saied AH, Steyn MP, Ansermino JM. Clonidine prolongs the effect of ropivacaine for axillary brachial plexus blockade. Canadian journal of anaesthesia 2000;47(10):962–7.
- Casati A, Magistris L, Beccaria P, Cappelleri G, Aldegheri G, Fanelli G. Improving postoperative analgesia after axillary brachial plexus anesthesia with 0.75% ropivacaine. A double-blind evaluation of adding clonidine. Minerva anestesiologica. 2001;67(5):407–12.
- Andersen JH, Jaeger P, Sonne TL, Dahl JB, Mathiesen O, Grevstad U. Clonidine used as a perineural adjuvant to ropivacaine, does not prolong

Asthana et al: Clonidine as an Adjuvant for Ropivacaine

the duration of sensory block when controlling for systemic effects: A paired, blinded, randomized trial in healthy volunteers. PloS one. 2017;12(9).

12. Pratap JN, Shankar RK, Goroszeniuk T. Co-injection of clonidine prolongs the anesthetic effect of lidocaine skin infiltration by a peripheral action. Anesthesia and analgesia. 2007;104(4):982–3.

Copyright: © the author(s), 2020. It is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY 4.0), which permits authors to retain ownership of the copyright for their content, and allow anyone to download, reuse, reprint, modify, distribute and/or copy the content as long as the original authors and source are cited.

How to cite this article: Asthana A, Tiwari JP, Tripathi DK, Prasad B. To Determine the Effect of Clonidine as an Adjuvant for Ropivacaine in Lower Limb Orthopedic Surgery. Acad. Anesthesiol. Int. 2020;5(1):199-202.

DOI: dx.doi.org/10.21276/aan.2020.5.1.42

Source of Support: Nil, Conflict of Interest: None declared.