

Quality of anesthesia induced by dexmedetomidine and acetaminophen in regional anesthesia comparative study

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Abstract

Introduction: Intravenous regional anaesthesia was first described in 1908 by August kar Gustav Bier, a German surgeon and pioneer of spinal anaesthesia, for anaesthesia of forearm and hand. He described a new method of producing analgesia of limb which he named 'vein anaesthesia'. August bier first presented his new method of intravenous regional anaesthesia at the 37th congress of the german surgical society on 22nd April 1908. Dexmedetomidine, an imidazole compound, is the pharmacologically active dextroisomer of medetomidine. It is a potent alpha-2 adrenoceptor agonist with eight times higher affinity for the alpha-2 adrenoceptor than clonidine, Alpha-2 agonists produce sedation, analgesia, hypnosis, anxiolysis and sympatholysis. **Aims and Objectives:** To evaluate the anaesthetic and analgesic effectiveness of Dexmedetomidine and acetaminophen when administered as adjuncts to lidocaine in intravenous regional anaesthesia to find out sensory and motor block onset times. **Methodology:** This was a clinical trial, in this patients were divided randomly into three group of 30 patients each. Patients were divided into **GROUP I:** Control Group, **GROUP II:** Dexmedetomidine. **GROUP III:** Paracetamol (Acetaminophen). The data was analyzed using computer software microsoft excel and SPSS version 10.0 for windows. The data was presented as mean and standard deviation and statistical significance was analyzed using one-way analysis of variance (ANOVA). Post-hoc intergroup significance was assessed using bonferroni, s t test. A 'p' value of <0.05 was considered statistically significant. Qualitative variable was analyzed using chi-square test. All analysis was conducted in accordance to intention to treat principle. **Result:** Onset of sensory block was significantly lower in Group II and Group III as compared to Group I (0.0001), Onset of Motor block was significantly lower in Group II and Group III as compared to Group I (0.0001). Excellent Quality of anesthesia was significantly higher in Group II as compared other groups (0.00001). There was significantly lower Visual Analogue Scale in Group II as compared to other groups, among intra-operative analgesia at 10, 20, and 30, 40, 50, 60. **Conclusion:** It is concluded that the addition of dexmedetomidine or acetaminophen to lidocaine in intravenous regional anaesthesia definitely improve the quality of anaesthesia and analgesia to a variable extent. However, dexmedetomidine is more potent, and provides better quality of anaesthesia and analgesia, and prolongs the duration of postoperative analgesia more than acetaminophen

Keywords: Quality of Anesthesia, Dexmedetomidine, Acetaminophen.

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INTRODUCTION

Intravenous regional anaesthesia (ivra) was first described in 1908 by August karl Gustav Bier, a German surgeon and pioneer of spinal anaesthesia, for anaesthesia of forearm and hand. He described a new method of producing analgesia of limb which he named 'vein anaesthesia'. August bier first presented his new method of intravenous regional anaesthesia at the 37th congress of the German surgical society on 22nd April 1908. His method, which now bears his name, consisted of occluding the circulation in a segment of the arm with two bandages and injecting dilute local anaesthesia through a venous cut down in this isolated segment,

which resulted in prompt analgesia (Bier, 1908)¹. The earliest agent injected was prilocaïne, the technique gained popularity when holmes used lidocaine and introduced several modifications, including either a second cuff or subcutaneous band of local anesthesia to control tourniquet pain (Holmes, 1963)² Lidocaine remains the standard local anaesthesia agent for surgical procedures in north America (Henderson *et al.*, 1997)³ and prilocaïne is used widely in Europe (Bader *et al.*, 1988)⁴ intravenous regional anesthesia is easy to administer, reliable and cost-effective, Major nerve blocks such as brachial block and femoral-sciatic block require technical expertise. Conversely, the administration of intravenous regional anesthesia requires only the skill necessary to perform a venipuncture (Brown *et al.*, 1989)⁵. The ideal intravenous regional anaesthesia solution should have the following features: rapid onset reduced tourniquet pain and prolonged post-deflation analgesia, local anaesthetics alone are not able to bestow all such attributes to the IVRA solutions; hence a multitude of adjuncts like opioids, tramadol, nonsteroidal anti-inflammatory drugs, clonidine, Dexmedetomidine, muscle relaxants, potassium magnesium, ketamine and alkalization with sodium bicarbonate have been used to improve the overall quality of anaesthesia and analgesia (Choyce and Peng 2002; Turan *et al.*, 2002)^{6,7}. Dexmedetomidine, an imidazole compound, is the pharmacologically active dextroisomer of medetomidine. It is a potent α -2 adrenoceptor agonist with eight times higher affinity for the α -2 adrenoceptor than clonidine (Bhanna N *et al.*, 2000)⁸ α -2 agonists produce sedation, analgesia, hypnosis, anxiolysis and sympatholysis (Miller 7th edition; p.751). Perioperative administration of Dexmedetomidine decreases the requirement for opioid or non-opioid analgesics both intra- and postoperatively (Jaakola ML *et al.*, 1993)⁹. Aho and colleagues concluded that intravenously administered Dexmedetomidine relieved pain and reduced opioid requirement after laparoscopic tubal ligation. However, patients who received Dexmedetomidine were more sedated than other patients in the study (Aho M *et al.*, 1991)¹⁰. Jaakola and colleagues demonstrated the analgesic efficacy of Dexmedetomidine in human tourniquet pain in their study (Jaakola ML *et al.*, 1991). The quality of intraoperative anaesthesia, shortens the onset of motor and sensory block, decreases pain and improves post-operative analgesia without any significant side-effects (Memis D *et al.*, 2004). However, Esmaoglu and associates observed that addition of 1 μ g/kg of Dexmedetomidine to lidocaine for intravenous regional anaesthesia improves the quality of anaesthesia and decreases analgesic requirements. But

has no effects on the sensory and motor block onset times (Esmaoglu A *et al.*, 2005)

AIMS AND OBJECTIVES

To evaluate the anaesthetic and analgesic effectiveness of Dexmedetomidine and acetaminophen when administered as adjuncts to lidocaine in intravenous regional anaesthesia to find out sensory and motor block onset times.

METHODOLOGY

After obtaining approval from hospital ethical committee, the study was conducted in Government Medical College, Jammu in the department of anaesthesiology and intensive care on ASA physical status I and II patients aged between 20-25 years, of either sex, scheduled for hand or forearm surgery, lasting less than 1 hour duration. History of drug allergy, Patients with sickle cell anemia, Patients with bleeding and coagulation disorders, Patients with liver disorders, Patients with Raynaud's disease, scleroderma, myasthenia gravis, renal insufficiency, history of convulsions, Pregnancy and lactation were excluded from the study. The patients were divided randomly into three groups of 30 patients each. Patients were divided into GROUP I: patients in this group received 10ml of preservative free lidocaine 2% diluted with saline to a total volume of 40ml. GROUP II: patients in this group received 10 ml of preservative free lidocaine 2 % and 0.5 μ g/kg of Dexmedetomidine [i.e. 0.5 ml for a 50 kg adult] mixed with saline to a total volume of 40 ml. GROUP III: patients in this group received 10 ml of preservative free lidocaine 2% mixed with 30 ml (300 mg) of paracetamol solution to make a total volume of 40 ml. The data was analysed using computer software Microsoft Excel and SPSS version 10.0 for Windows. The data was presented as mean and standard deviation and statistical significance was analyzed using one-way analysis of variance (ANOVA). Post-hoc intergroup significance was assessed using Bonferroni's *t* test. A 'p' value of <0.05 was considered statistically significant. Qualitative variable was analyzed using chi-square test. All analysis was conducted in accordance to intention to treat principle.

RESULTS

Table 1: Onset of sensory block (minutes)

	Onset of sensory block (minutes)		
	mean	\pm SD	Range
Group I	5.20	1.08	3-7
Group II	1.66	0.55	1-3
Group III	4.53	1.23	2-7

F= 105.1 p-value = 0.0001 (HS)

From Table 1: In group I, mean onset of sensory block was 5.20 ± 1.08 minutes. In group II, mean onset of

sensory block was 1.66 ± 0.55 minutes. In group III, means onset of sensory block was 4.53 ± 1.23 minutes. Using ANOVA, there was statistically significant among all the three groups ($p < 0.05$).

Table 2: Onset of motor block (minutes)

	Onset of motor block (minutes)		
	Mean	± SD	Range
Group I	9.68	1.72	705-15
Group II	5.45	1.85	3.5-10
Group III	9.51	1.83	7.14

F=52.79, P= value = 0.0001(HS)

From Table 2: In group I, mean onset of motor block was 9.68 ± 1.72 minutes. In group II. Mean onset of motor block was 5.45 ± 1.85 minutes. In group III, mean onset

of motor block was 9.51 ± 2.83 minutes. Using ANOVA, there was statistically significant difference among all the three groups ($p < 0.05$).

Table 3: Quality of anesthesia (no. and % of patients)

	Quality of anesthesia (no. of patients)					
	Excellent		Good		Moderate	
	n	%	n	%	n	%
Group I	10	33.3%	15	50%	5	16.6%
Group II	28	93.3%	2	6.6%	0	0%
Group III	20	66.6%	10	33.3%	0	0%

$\chi^2=27.96$, p-value<0.0001 (highly significant) among all the three groups

Excellence Quality of anesthesia was significantly higher in Group II as compared other groups.

Table 4: Intra-Operative Analgesia by Visual Analogue Scale (0.10 Cm) At 10, 20, 30, 40, 50, 60 Minutes

	VAS at 10 minutes (Mean±SD)		Vas at 20 minutes(Mean± D)		VAS at 30 minutes(Mean±S)		Vas at 40 minutes(Mean±SD)		VAS at 50 minutes(Mean±SD)		VAS at 60 minutes(Mean±S D)	
Group I	0.56	0.50	0.93	1.11	1.56	1.40	1.23	0.72	1.53	0.57	1.90	0.48
Group II	0.13	0.34	0.20	0.40	0.30	0.46	0.56	0.50	0.73	0.52	0.00	0.52
Group III	0.33	0.47	0.26	0.46	0.40	0.49	0.66	0.47	0.10	0.75	0.30	0.65
p-value	F=7.01, p=0.0001(Hs)		F=9.22 p=0.0001(Hs)		F=18.26 p=2.83E-07 (Hs)		F= 16.35 p=0.0001(HS)		F= 12.29 p=0.0001(HS)		F= 24.47 p=3.62 (HS)	

From Table 4. There was significantly lower Visual Analogue Scale in Group II as compared to other groups, among intra-operative analgesia at 10, 20, and 30,40,50,60.

DISCUSSION

In our study, we used scoring scale given by palecha *et al*, to assess the sensory and motor block onset time. (palecha *s et al*. 2001) The mean time of onset of sensory and motor block in group I was 5.20 ± 1.08 minutes and 9.68 ± 1.72 minutes; in group II was 1.66 ± 0.55 minutes and 5.45 ± 1.85 minutes; and in group III was 4.53 ± 1.23 minutes and 9.51 ± 1.83 minutes, respectively. Statistically the difference was found to be significant amongst all the group using ANOVA ($p < 0.05$). in the intergroup comparison, between group I and II, it was observed that the sensory and motor onset time gets significantly shortened in bonferroni's t-test) in astudy by memis *et al* (2004), the mean time of onset of sensory and motor block was 5 ± 2 minutes and 10 ± 4 minutes respectively, when 0.5 ug/kg Dexmedetomidine was added to 40 ml of 0.5% lidocaine in IVRA, and they observed that the sensory and motor block onset time gets significantly shortened in Dexmedetomidine group as compared to control group($p < 0.05$).however, in study by esmaoglu *et al* (2005), statistically the onset time for sensory and motor block in Dexmedetomidine group

(4.8 ± 2.0 min. and 11.2 ± 4.6 min) was found to be similar as in control group ($p < 0.05$), although they used the dose of 1 ug/kg of Dexmedetomidine in 40ml of 0.05% lidocaine in ivra . Our finding are comparable to the study by Memis *et al*¹¹, but differs from the study by Esmaoglu *et al*¹². In the intergroup comparison, between II and III, we observed that there was statistically significant difference in the sensory and motor block onset time between Dexmedetomidine and acetaminophen group ($p < 0.05$ Using bonferroni's t-test). Thus, Dexmedetomidine provides earlier onset of sensory and motor block as compared to acetaminophen. This could be because of enhancement of local anaesthetic action of lidocaine. by Dexmedetomidine Yoshitomi *et al*. (2008)¹³, and not by acetaminophen. The change in the ph of injected solutions, due to adjuvants, added to lidocaine, can also affects the onset of block, which was however, not tested in our study. Quality of anaesthesia score was assessed on numeric scale as-excellent (4): no complaint from patient; good (3): minor complaint with no need of supplemental analgesics; moderate (2): complaint which required supplemental analgesics unsuccessful (1): patient given general anaesthesia (Esmaoglu *et al*, 2005) In group II, 28(93.3%) patients had excellent quality of anaesthesia score, 2(6.6%) had good quality of anaesthesia score. [range (3-4)] In group III, 20(66.6%) patients had excellent quality of anaesthesia score, 15(50%) had good

quality of anaesthesia score, and 5 (16.6%) had moderate quality of anaesthesia score [range (2-4)]. In group II and III, no patient had moderate score. In the intergroup comparison between group I and II, since more number of patients in Dexmedetomidine group had excellent score as compared to control group, Dexmedetomidine provides better quality of anaesthesia than control group. Memic *et al* (2004), also observed excellent quality of anaesthesia score [range (3-4)] in Dexmedetomidine group and good [range (2-3)] in control group and difference was statistically significant ($p < 0.05$). In the intergroup compare between group I and II, since more number of patients in acetaminophen group had excellent score as compared to control group. Acetaminophen provides better quality of anaesthesia than control group. Sen *et al* (2009)¹⁴, observed that anaesthesia quality was excellent [range (2-4)] in acetaminophen group and good ($p < 0.05$). Our findings are comparable to these studies. In the intergroup comparison between group II and III, more percentage of patient in Dexmedetomidine group had excellent quality of anaesthesia score than the percentage of patients in acetaminophen group, suggesting that Dexmedetomidine provides better quality of anaesthesia than acetaminophen. Intraoperative analgesia was assessed by visual analogue scale of 0-10 [0=no pain, 10=worst pain]. At 10 minutes, the difference in vas was statistically significant between group I and II with lower vas in group II ($P < 0.05$ using Bonferroni's t-test) there was statistically insignificant between group I and III, and group II and III ($p > 0.05$). At 20 minutes, the difference in vas was statistically significant between group I and II and group I and III with lower vas in group II and III. ($p < 0.001$ using Bonferroni's t-test). There was statistically insignificant difference between group II and III ($p > 0.05$). Similarly at 30, 40, 50 and 60 minutes, there was statistically significant difference in vas between group I and II, and group I and III ($p < 0.001$) and insignificant between II and III ($p > 0.05$). Statistically there was significantly lower vas in group II at 10, 20, 30, 40, 50 and 60 minutes when compared to control group ($p < 0.001$ using Bonferroni's t-test) in the study by Memis *et al* (2004), there was statistically significant difference in vas score at 5, 10, 15, 20 and 40 minutes after tourniquet inflation. There was statistically highly significant lower vas in Dexmedetomidine group (p -value < 0.001) as compared to control group. Esmaoglu *et al* (2005)¹², also observed significantly lower vas score in the Dexmedetomidine group with lesser requirement of intraoperative analgesics as compared to control group ($p < 0.05$). Our results are comparable to these studies. Sato *et al* (1991)¹⁵ reported that α_2 adrenergic receptors located at nerve ending have a role in the analgesia effects of the drug by preventing norepinephrine release.

Therefore, Dexmedetomidine, by preventing norepinephrine release from nerve terminals, produced analgesic effects and thus, significantly lower vas score was observed in Dexmedetomidine group as compared to control group. In our study, however, we observed lower vas score in acetaminophen group as compared to control group in the intraoperative period. Canbay *et al* (2008)¹⁶ reported that acetaminophen pretreatment appears to be effective in reducing the pain experienced during intravenous injection of propofol. This suggests the peripheral antinociceptive effects of acetaminophen. Deciga-c *et al*, (2004)¹⁷ reported that which are more resistant to lidocaine than A-delta fibres, and to opening of potassium channels located in primary afferent nerve endings. In the intergroup comparison between group II and III, there was statistically insignificant difference in the Dexmedetomidine and acetaminophen group ($p > 0.05$) suggesting that both the drugs significantly lower intraoperative vas scores are compared to control group and thus improve intraoperative analgesia.

CONCLUSION

It is concluded that the addition of dexmedetomidine or acetaminophen to lidocaine in intravenous regional anaesthesia definitely improves the quality of anaesthesia and analgesia to a variable extent. However, dexmedetomidine is more potent, and provides better quality of anaesthesia and analgesia, and prolongs the duration of postoperative analgesia more than acetaminophen.

REFERENCES

1. Bier A.A. A new method for local anaesthesia in the extremities. *Ann Surg* 1908; 48: 780-3
2. Holmes CMCK. Intravenous regional neural blockade. Neural blockade in clinical anaesthesia and management of pain, 3rd ed. Philadelphia: Lippincott-Raven, 1998; pp. 395-410.
3. Henderson CL, Warrier CB, McEwan JA, et al. A North American survey of intravenous regional anaesthesia. *Anesth Analg* 1997; 85: 858-63.
4. Bader AM, Conception M, Hurley RJ, et al. Comparison of lidocaine and prilocaine for intravenous regional anaesthesia. *Anaesthesiology* 1988; 69: 409-12.
5. Brown EM, McGriff JT, Malinowski RW. Intravenous regional anaesthesia (Bier's Block): review of 20 years experience. *Can J Anaesth* 1989; 36: 307-10.
6. Choyce A, Peng P. A systematic review of adjuncts for regional anaesthesia for surgical procedures. *Can J Anaesth* 2002; 49: 32-45.
7. Turan A, Karamanoglu B, Memis D, et al. Intravenous regional anaesthesia using prilocaine and neostigmine. *Anesth Analg* 2002; 95: 1419-22.
8. Bhana N, Goa KL, McClellan KJ. Dexmedetomidine. *Drugs* 2000; 59: 263-8.

9. Jaakola ML. Dexmedetomidine premedication before intravenous regional anaesthesia in minor outpatient hand surgery. *J Clin Anesth* 1994;6: 204-11
10. Aho Ms, Erkola OA, Scheinin H et al. Effect of intravenously administered Dexmedetomidine on pain after laproscopic tubal ligation. *Anaesth Analg* 1991; 73:112-8.
11. Memis D, Turan A, Karamanliglu B, et al. Adding Dexmedetomidine to lidocaine for intravenous regional anaesthesia. *Anesth Analg* 2004; 98: 835-40.
12. Esmaoglu A, Mizrak A, Akin A, et al. Addition of Dexmedetomidine to lidocaine for intravenous regional anaesthesia. *Eur J Anaesth* 2005; 22(6): 447-51.
13. Yoshitomi T, Kohijitani A, Maeda S, et al. Dexmedetomidine enhances the local anaesthetic action of lidocaine via an α -2A adrenoceptors. *Anesth Analg*. 2008; 107(1); 96-101.
14. Sen H, Kulahci Y, Qzkan S, et al. The analgesic effect of paracetamol when added to lidocaine in intravenous regional anaesthesia. *Anaesth Analg* 2009;109(4):1327-30
15. Sato J, Per IER. Adrenergic excitation of cutaneous pain receptors induced by peripheral nerve injury. *Science* 1991; 251: 1608-10.
16. Canbay O, Celebi N, Arun O, et al. Efficacy of intravenous acetaminophen and lidocaine on propofol injection pain. *Br J Anaesth* 2008; 100: 95-98.
17. Deciga-campos M, Lopez Munoz FJ. Participation of the L-arginine-nitric oxide/cyclic GMP-ATP-sensitive K_2 channel cascade in the antinociceptive effects of rofecoxib. *Eur J Pharmacol* 2004; 484:193-9.

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