Research Article

Evaluation of two different doses of Dexmedetomidine in attenuation of pressor response during laryngoscopy and intubation

Shilpa Sale^{1*}, Shama Shikalgar², Prashant Lomate³, Ganesh Kumbhar⁴, Vikas Kumar⁵

^{1,5}PG Student, ²Professor, ³Assistant Professor, ⁴Lecturer, Department of Anaesthesia, BVDU Medical College and Hospital, Sangli, Maharashtra, INDIA.

Email: shilpasale@gmail.com

Abstract

Introduction: Pressor response to laryngoscopy and intubation have been associated with sympathetic and parasympathetic responses like Hypertension, Tachycardia due to increased plasma catecholamine concentrations. Premedication with Dexmedetomidine attenuates pressor response to laryngoscopy and intubation. Aims and Objective: to evaluate and compare effects of 2 different doses of Dexmedetomidine 1mcg/kg and 0.6mcg/kg in attenuating pressor response to laryngoscopy and intubation. Methods: 60 patients aged 18-50 years of either sex of ASA grade I and II scheduled for elective surgery were randomly divided into 2 groups. All patients were uniformly premedicated, 10 mins before induction. Group A received Inj Dexmedetomidine 1mcg/kg and Group B received Inj Dexmedetomidine 0.6mcg/kg iv. After induction with Thiopentone and succinylcholine, laryngoscopy and intubation was done. HR, SBP, DBP were recorded for both groups before premedication, 10min after premedication and 1min, 2min, 3min, 5min, 7min,10min after intubation. Data was analyzed and compared with paired t test. Results: HR and BP levels were lower at 10min after premedication and 5min after intubation than baseline levels in both groups. But in Dexmedetomidine 1mcg/kg group amount of increase in HR, SBP, DBP were lower compared to 0.6 mcg/kg. Conclusion: Dexmedetomidine 1mcg/kg is more effective than 0.6 mcg/kg in attenuating pressor response to laryngoscopy and intubation and provides more stable hemodynamic profile.

Keywords: Dexmedetomidine.

*Address for Correspondence:

Dr. Shilpa Sale, PG Student, Department of Anaesthesia, Bharati Vidyapeeth Medical College and hospital, Wanlesswadi, Sangli-Miraj Road, Sangli 416414 Maharashtra, INDIA.

Email: shilpasale@gmail.com

Received Date: 01/08/2015 Revised Date: 10/08/2015 Accepted Date: 13/08/2015



INTRODUCTION

The hemodynamic pressor responses to laryngeal and tracheal stimulation following laryngoscopy and tracheal intubation were described by Reid and Brace in 1940¹ and King *et al* in 1951. To date exact mechanism of hemodynamic responses to laryngoscopy and intubation have not been clarified but have been associated with both sympathetic ^{2,3,4} and parasympathetic response⁵.

These may include hypertension, Tachycardia due to increased plasma catecholamine concentrations⁴. It is usually well tolerated by healthy individuals, but even short lasting stimulation may be fatal in patient with hypertension, recent MI, preeclampsia, cerebrovascular diseases like tumor's, aneurysms or intracranial hypertension^{2,5,6}. Various attenuating agents for this response have been used including opioids, barbiturates, benzodiazepines, beta blockers, calcium channel blockers, vasodilators etc. 7,8,9,10,11 Dexmedetomidine is the newer highly selective, specific and potent alpha 2 agonist having 8 times more affinity for alpha2 adrenoreceptors, shorter duration of action than clonidine and has an inhibitory effect on catecholamine release level^{12,13,14,13} thereby decreaseing their plasma Dexmedetomidine has additional effects such as anxiolysis, sedation, analgesia improved hemodynamic stability. Various studies have also documented that different doses of Dexmedetomidine can decrease the hemodynamic response to laryngoscopy and intubation ^{15,16,17,18,19,20,21,22}. So that we decided to compare the two different doses of Dexmedetomidine - 1mcg/kg iv and 0.6mcg/kg iv to control hemodynamic responses due to laryngoscopy and intubation.

MATERIALS AND METHODS

After approval from ethical committee, the randomized prospective double blind study was carried out in tertiary referral medical college from December 2013.

Inclusion Criteria

- 60 patients belonging to
- Age group 18 to 50 years
- ASA grade I and II
- Mallampatti grade I
- Elective surgical procedures under GA

Exclusion Criteria

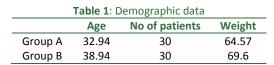
- Patient refusal
- ASA grade III and IV
- Severely Hypovolemic
- Difficult airway
- History of cardiac disease and neurological disease

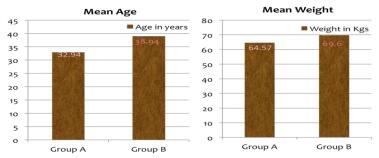
- Patients were subjected to routine investigations like CBC, BSL, Renal and Liver function tests.
 Written informed valid consent was taken from patient. Patients were premedicated with inj glycopylorate 0.004mg/kg i.m, ondansetron 0.15mg/kg iv, Inj midazolam 0.03 mg/kg iv half an hour before induction
- Patients were randomly divided in two groups

Group A: Dexmedetomidine 1mcg/kg **Group B:** Dexmedetomidine 0.6mcg/kg

Injection Dexmedetomidine1mcg/kg diluted up to 10ml with NS and Injection Dexmedetomidine 0.6mcg/kg diluted upto 10ml with NS in groups A and B respectively were administered intravenously 10 minutes before laryngoscopy and intubation. Heart rate, Systolic blood pressure, diastolic blood pressure were recorded before premedication, after premedication, 1 min, 2 min, 5min, 7min and 10 min after intubation. Induction was done with Inj. Thiopentone 4-5 mg/kg and Inj. Succinylcholine 2mg/kg followed by laryngoscopy and intubation. Patients were maintained on oxygen (50%). Nitrous oxide (50%), Sevoflurane, nondepolarising muscle relaxants. Analysis of data done.

RESULTS





Mean age and weight of both study groups are comparable to each other.

Figure 1: Mean Age and Weight

Table 2: Changes in heart rate

	HR (min)	Base line	10 min after pre med.	1 min after ETI	2 min after ETI	3 min after ETI	5 min after ETI	7 min after ETI	10 min after ETI
Group A	Mean	84.4	67.47*	80*	80.87*	79.8*	75.34*	73.6*	70.14*
	±SD	8.76	6.20*	10.45*	8.80*	7.31*	5.59*	6.07*	7.43*
Group B	Mean	81.47	78.8*	90.94*	94.6*	93.74*	90.54*	88*	85.34*
	±SD	7.14	7.49*	6.64*	6.72*	7.44*	7.97*	8.36*	7.34*

*Highly Significant

HR increased in both groups immediately after endotracheal intubation. But rise in HR 2min after endotracheal intubation is more in Group B compared to group A.

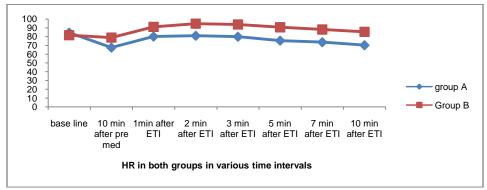


Figure 2: The increase was highly significant in Group B as compared to Group A during laryngoscopy and intubation

Table 3: Changes in systolic blood pressure

	SBP (mmgh)	Base line	10 min after pre med.	1 min after ETI	2 min after ETI	3 min after ETI	5 min after ETI	7 min after ETI	10 min after ETI
Group A	Mean	132.4	112.47*	126.8*	127.5*	124.53*	120*	113.2*	107.2*
	±SD	6.22	8.47*	8.97*	11.17*	12.40*	10.74*	10.8*	9.87*
Group B	Mean	132.27	126.47*	130.2*	145.14*	139.47*	132.2*	126.2*	119.5*
	±SD	6.84	7.34*	6.26*	5.26*	3.89*	4.29*	4.14*	5.59*

*Highly Significant

SBP increased in both groups immediately after endotracheal intubation. But rise in SBP 2min after endotracheal intubation is more in Group B than Group A.

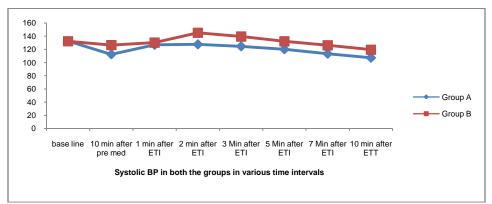


Figure 3: The increase was highly significant in Group B as compared to Group A during laryngoscopy and intubation

Table 4: Changes in dystolic blood pressure

	DBP (mmgh)	Base line	10 min after pre med.	1 min after ETI	2 min after ETI	3 min after ETI	5 min after ETI	7 min after ETI	10 min after ETI
Group A	Mean	81.87	69*	79.8	77.86	74.67*	71.27*	68.67*	66.2*
	±SD	6.19	5.38*	10.03	11.64	11.10*	9.38*	5.38*	5.48*
Group B	Mean	82.33	73.40*	76.07	81.07	79.47*	76.33*	73.73*	71.73*
	±SD	3.36	7.44	6.07	5.54	3.93*	3.14*	3.55*	3.31*

*Highly Significant

DBP increased in both groups immediately after endotracheal intubation. But rise in DBP 3 min after endotracheal intubation is more in Group B than Group A.

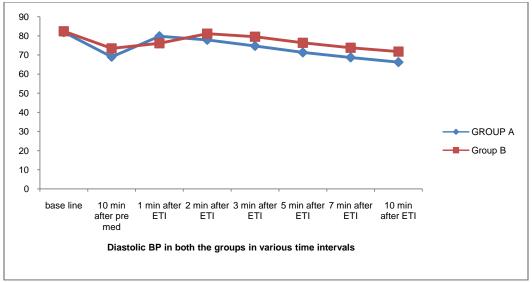


Figure 4: The increase was highly significant in Group B compared to Group A during laryngoscopy and intbation

DISCUSSION

Laryngoscopy and endotracheal intubation results in transient, but marked sympathetic and sympathoadrenal response. In patients with hypertension, Ischemic Heart Disease, cerebrovascular disease and intracranial aneurysms it can result in harmful consequences like left ventricular failure, pulmonary edema, myocardial infarction, ventricular dysarrhythmias and cerebral hemorrhage. It is for these reasons various studies had been carried out to search effective drug to attenuate this Hemodynamic response. Role of Dexmedetomidine as an analgesic, sedative, anxiolytic, sympatholytic and blunting agent for increased hemodynamic responses is being extensively studied. These responses are mediated by activation of alpha 2 receptors which are involved in regulating autonomic and cardiovascular systems. Alpha 2 receptors are located on blood vessels mediate vasoconstriction and on sympathetic terminals inhibit release of catecholamines. Alpha 2 receptors are also located within the postsynaptic terminals in central nervous system, their activation result in sedation, decreased neuronal activity and augmentation of vagal activity. 12,23,24,25,26,27,28 Our study was conducted to compare the efficacy of i.v. Dexmedetomidine 1mcg/kg with Dexmedetomidine 0.6mcg/kg to attenuate pressor response to laryngoscopy and intubation. Our study was carried out in 60 patients of ASA grade 1 and 2 who were divided in 2 groups of 30 each. The group A was given inj Dexmedetomidine 1mcg/kg and group B was given inj Dexmedetomidine 0.6 mcg/kg 10 minutes before laryngoscopy and intubation. Hemodynamic parameters like heart rate, systolic blood pressure, diastolic blood pressure were measured at different time intervals in each patient. We found that both groups of Dexmedetomidine

showed attenuated pressor response to Laryngoscopy and endotracheal intubation. Different studies with different doses of Dexmedetomidine like 0.5 mcg/kg, 0.6mcg/kg, 0.75mcg/kg, 1 mcg/kg have been done to attenuate pressor response to intubation. So we decided to evaluate and compare efficacy of two different doses of Dexmedetomidine i.e. 1mcg/kg and 0.6mcg/kg. It was found in the study by Scheinin's et al that Dexmedetomidine 0.6mcg/kg attenuated hemodynamic response to intubation and required dose of thiopentone is significantly lower¹⁶. Jaakola *et al* concluded that, during the intubation blood pressure and heart rate is significantly reduced by 0.6 μg. kg⁻¹ dexmedetomidine ¹⁷. Yildiz et al found that a single dose of 1mcg/kg dexmedetomidine prevented cardiovascular hemodynamic response and decreased need for additional opioid during laryngoscopy and endotracheal intubation in elective minor surgery patients¹⁵. Ozkose et al. administered a single dose of 1mcg/kg Dexmedetomidine 10 min before induction. They reported that with control measurements, mean arterial pressures decreased up to 20% and heart rate is decreased up to 15% 1min and 3 min following intubation³¹. In the other study which was done by Tezer et al. it is concluded that sympathetic responses during laryngoscopy and intubation were effectively reduced by dexmedetomidine 1 µg.kg⁻¹h⁻¹ and esmolol 250 µg/kg/min³². Khan et al. demonstrated that heart rate, systolic and diastolic blood pressure were reduced by dexmedetomidine³³. In another study on the patients undergoing vascular surgery, it was observed that in the recovery period dexmedetomidine infusion led to supression on heart rate and plasma cathecholamine levels³⁴. Jaaokola et al¹⁷, R saraf et al³⁰, B. scheinin et al¹⁶, Tezer et al concluded in their study that

Dexmedetomidine 0.6 mcg/kg significantly reduced blood pressure and Heart rate during intubation. On the other hand, Yildiz et al¹⁵, Ozkose et al³¹, Aho et al¹⁸ Sukhminderjit singh Bajwa et al²⁹ in their studies concluded that sympathetic response during intubation effectively reduced by Dexmedetomidine 1mcg/kg which is comparable to our study. In this study in both groups, Heart rate, Systolic blood pressure, Diastolic blood pressure started decreasing up to 10min after premedication with study drug in respective group. These observations are comparable to those by Sukhminderjit singh Bajwa et al²⁹, B. scheinin et al¹⁶. We observed decrease of HR, SBP, DBP in Dexmedetomidine 1mcg/kg group is more than Dexmedetomidine 0.6 mcg/kg. In both groups HR, SBP, DBP increased after laryngoscopy and intubation. The peak increase in HR, SBP, DBP was seen at 1min after intubation. However increase in HR, SBP, DBP is less in group Dexmedetomidine 1mcg/kg as compared to Dexmedetomidine 0.6 mcg/kg. This increase was highly significant. No adverse cardiovascular effects from drug were seen in the present study. Bradycardia (HR<50/min), a possible consequence of administration of alpha 2 agonist, was counteracted by Atropine 0.6mg iv. Also no adverse effects like nausea, vomiting, respiratory depression, apnea, muscle rigidity were seen.

CONCLUSION

Both Dexmedetomidine 1mcg/kg and 0.6 mcg/kg attenuate pressor response to laryngoscopy and endotracheal intubation. But Dexmedetomidine at dose of 1mcg/kg is more effective than 0.6 mcg/kg in attenuating pressor response and provides more stable hemodynamic profile.

REFERENCES

- Reid LC, Brace DE. Irritation of the respiratory tract and its reflex effect upon heart surg Gynec and Obst 1940; 70: 157-62
- Kayhan Z, Aldemir D, Mutlu H, Ög*űs, §E. Which is responsible for the haemodynamic response due to laryngoscopy and endotracheal intubation? Catecholamines, vasopressin or angiotensin? Eur J Anaesthesiol 2005; 22: 780-5.
- 3. Hung O. Understanding hemodynamic responses to tracheal intubation. Can J Anaesth 2001; 48: 723-6.
- 4. Kaymak Ç, Kocabas, Ş NA, Durmaz E, Öztuna D. Adrenoceptor (ADRB2) pharmacogenetics and cardiovascular phenotypes during laryngoscopy and tracheal intubation. Int J Toxicol 2006; 25: 443-9.
- Kovac AL. Controlling the hemodynamic response to laryngoscopy and endotracheal intubation. J Clin Anesth 1996; 8: 63-79.
- Morin AM, Geldner G, Schwarz U, et al. Factors influencing preoperative stress response in coronary artery bypass graft patients. BMC Anesthesiology 2004; 4: 7.

- Charuluxananan S, Kyokong O, Somboonviboon W, Balmongkon B, Chaisomboonpan S. Nicardipine versus lidocaine for attenuating the cardiovascular response to endotracheal intubation. J Anesth. 2000; 14:77–81. [PubMed]
- 8. Menda F, Koner O, Sayin M, Ture H, Imer P, Aykac B. Dexmedetomidine as an adjunct to anesthetic induction to attenuate hemodynamic response to endotracheal intubation in patients undergoing fast-track CABG. Ann Card Anaesth. 2010; 13:16–21. [PubMed]
- 9. Gunes Y, Gunduz M, Ozcengiz D, Ozbek H, Isik G. Dexmedetomidine-remifentanil or propofol-remifentanil anesthesia in patients undergoing intracranial surgery. Neurosurg Q. 2005; 15:122–6.
- Powroznyk A, Vuylsteke A, Naughton C, Misso S, Holloway J, Jolin-Mellgard A, et al. Comparison of clevidipine with sodium nitroprusside in the control of blood pressure after coronary artery surgery. Eur J Anaesth. 2003; 20:697–703. [PubMed]
- Abou-Arab MH, Heier T, Caldwell JE. Dose of alfentanil needed to obtain optimal intubation conditions during rapid-sequence induction of anaesthesia with thiopentone and rocuronium. Br J Anaesth. 2007; 98:604–10. [PubMed]
- 12. Hall JE, Uhrich TD, Ebert TJ. Sedative, analgesic and cognitive effects of clonidine infusions in humans.Br J Anaesth. 2001; 86:5–11. [PubMed]
- 13. Bajwa SJ, Bajwa SK, Kaur J, Singh G, Arora V, Gupta S, et al. Dexmedetomidine and clonidine in epidural anaesthesia: A comparative evaluation. Indian J Anaesth. 2011; 55:116–21. [PMC free article] [PubMed]
- Guler G, Akin Z, Tosun E, Eskitascoglu, Mizrak A, Boyaci A. Single-dose dexmedetomidine attenuates airway and circulatory reflexes during extubation. ActaAnaesthesiol Scand. 2005; 49:1088–91. [PubMed]
- 15. Yildiz M, Tavlan A, Tuncer S, Reisli R, Yosunkaya A, Otelcioglu S. Effect of dexmedetomidine on haemodynamic responses to laryngoscopy and intubation: Perioperative haemodynamics and anaesthetic requirements. Drugs R D. 2006; 7:43–52. [PubMed]
- Scheinin B, Lindgren L, Randell T, Scheinin H, Scheinin M. Dexmedetomidine attenuates sympathoadrenal responses to tracheal intubation and reduces the need for thiopentone and perioperative fentanyl. Br J Anaesth 1992; 68: 126-31.
- Jakola ML, Ali-Melkkila T, Kanto J, Kallio A, Scheinin H, Scheinin M. Dexmedetomidine reduces intraocular pressure, intubation response and anaesthetic requirements in patients undergoing ophthalmic surgery. Br J Anaesth 1992; 68: 570-5.
- Aho M, Lehtnen AM, Erkola O, Scheinin H, Lehtinen A, Kallio A, et al. The effect of intravenously administered dexmedetomidine on perioperative haemodynamics and isoflurane requirements in patients undergoing abdominal hysterectomy. Anaesthesiology 1991; 74: 997-1002.
- Mowfi HA, Aldossary N, Ismail SA, Alqutiani J. Effect of dexmedetomidine premedication on the intraocular pressure changes after succinylcholine and intubation. Br JAnaesth 2008; 100 (4); 485-9.
- 20. Basar H, Akpinar S, Doganci N, Buyukkocak U, Kaymak C, Sert O, et al. The effect of preanaesthetic, single dose dexmedetomidine on induction, haemodynamic and

- cardiovascular parameters. Journal of ClinAnaesth 2008; 20: 431-6.
- Kunisawa T, Nagata O, Nagashima M. Dexmedetomidine suppresses the decrease in blood pressure during anaesthetic induction and blunts the cardiovascular responses to tracheal intubation. Journal ofClinAnaesth 2009; 21:194-9.
- 22. Menda F, Koner O, Sayin M, Ture H, Imer P, Aykac B. Dexmedetomidine as an adjunct to anesthetic induction to attenuate haemodynamic response to endotracheal intubation in patients undergoing fast-track CABG. Ann Card Anaesth 2010; 13: 16-21.
- 23. Wijeysundera DN, Naik JS, Beattie WS. a-2 adrenergic agonists to prevent perioperative cardiovascular complications: A meta analysis. Am J Med. 2003; 114:742–52. [PubMed]
- Ebert T, Maze M. Dexmedetomidine: Another arrow for the clinician's quiver. Anesthesiology. 2004; 101:568–70. [PubMed]
- 25. Gerlach AT, Dasta JF. Dexmedetomidine: An updated review. Ann Pharmacother. 2007; 41:245–52. [PubMed]
- BekkerA, Sturaitis M. Dexmedetomidine for neurosurgical surgery. Operative Neurosurg. 2005; 57:1– 10.
- Tanskanen P, Kytta J, Randell T, Aantaa R. Dexmedetomidine as an anaesthetic adjuvant in patients undergoing intracranial tumor surgery: A double-blind, randomized and placebo-controlled study. Br J Anaesth. 2006; 97:658–65. [PubMed]

- 28. Sturaitis M, Kroin J, Swamidoss C, Moric M. Effects of intraoperative dexmedetomidine infusion on hemodynamic stability during brain tumor resection. Anesthesiology. 2002;98:A-
- SukhminderJit Singh Bajwa, JasbirKaur, Amarjit Singh-Attenuation of pressor response and dose sparing of opioids and anaesthetics with preoperative Dexmedetomidine, IJA, vol5 issue2,2012
- R. Saraf, M Jha, Sunil Kumar- Dexmedetomidine and pressor response, Paediatric Anesthesia and critical care Journal 2013;1(1):78-86
- Ozkose Z, Demir FS, Pampal k, et al Hemodynamic and anaesthetic advantages of Dexmedetomidine, an alpha2 agonist, for surgery in prone position. Tohoku J Exp Med.2006; 210:153-60.
- Tezer E, Sarıcaoğlu F, Çelebi N, Aypar Ü. Comparison of esmolol and dexmedetomidine in induction of anesthesia about hemodynamic and anaesthetic requirements. Turkish Journal of Anesthesia 2005; 13(4):247-252.
- Khan ZP, Munday IT, Jones RM, Thornton C, Mant TG, Amin D. Effects of dexmedetomidine on isoflurane requirements in healthy volunteers. 1:Pharmacodynamic and pharmacokinetic interactions. Br J Anaesth 1999; 83: 372-80.
- 34. Talke P, Chen R, Thomas B, et al. The hemodynamic and adrenergic effects of perioperative dexmedetomidine infusion after vascular surgery. Anesth Analg. 2000; 90(4): 834-9.

Source of Support: None Declared Conflict of Interest: None Declared