# EFFECT OF DEXMEDETOMIDINE AND ESMOLOL IN ATTENUATING THE HAEMODYNAMIC RESPONSES DURING LARYNGOSCOPY AND ENDOTRACHEAL INTUBATION IN ELECTIVE UPPER ABDOMINAL SURGERIES

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#### ABSTRACT

#### BACKGROUND

Laryngoscopy and endotracheal intubation produces distinct but transient increase in cardiac workload. In this study, a comparison is made between dexmedetomidine, esmolol and control in their effect in attenuation of pressure response during laryngoscopy and endotracheal intubation.

#### MATERIALS AND METHODS

With written consent, we studied hundred and twenty-five (125) adult patients of American Society of Anesthesiologists physical status I and II, aged between 30 to 60 years, of either sex, undergoing elective upper abdominal surgeries. The patients were randomly allocated into three groups: 1) 42 patients in group (D) received Dexmedetomidine (1  $\mu$ g/kg), 2) 42 patients in group (C) received normal saline 15 ml, and 3) 41 patients in group (E) received Esmolol (2 mg/kg).

All patients received the drugs intravenously over 10 minutes and 3 min before induction of general anaesthesia. Premedication, induction and intubation were similar. Heart rate (HR), systemic arterial pressures were recorded at baseline, after study drug infusion, after induction, immediately and 3, 5, 7, 10 min after intubation.

Study Design- Prospective, randomized, double blind, controlled study.

Statistical Analysis- Analysis of variance and t-test as appropriate.

#### RESULTS

The heart rate, systolic arterial pressures and rate-pressure product immediately after intubation and thereafter were significantly lower in Group D (P<0.001) when compared to Group E and Group C. Group E had the same post-intubation parameters reliably (P<0.001) lower than Group C.

#### CONCLUSION

Dexmedetomidine and esmolol were both effective in attenuating the haemodynamic response to intubation, but dexmedetomidine was more effective than esmolol in lowering the haemodynamic response.

#### **KEYWORDS**

Dexmedetomidine, esmolol, normal saline, laryngoscopy, intubation, haemodynamic response.

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#### BACKGROUND

Laryngoscopy and endotracheal intubation is a noxious stimulus, which can provoke untoward response in cardiovascular, respiratory and other physiological systems.<sup>1</sup> The effect is transient, occurring immediately after intubation and lasting for 5-10 minutes. Intubation response may be well tolerated by normal, fit, ASA (American Society of Anaesthesiologist) physical status-I patients;<sup>2,3</sup> but can be

Financial or Other, Competing Interest: None. Submission 09-06-2018, Peer Review 11-06-2018, Acceptance 25-06-2018, Published 29-06-2018. Corresponding Author: Dr. Amalendu Bikas Chatterjee, Subhankar Sarami, Pratap Bagan East, Bankura -722101, West Bengal. E-mail: chatterjee\_amalendu@yahoo.com DOI: 10.18410/jebmh/2018/429 COOSO deleterious in patients with poor cardiovascular reserve.<sup>4</sup> In patients with coronary artery disease (CAD), hypertension, raised intra-cranial pressure it may be associated with myocardial infarction, arrhythmias, cardiac failure or cerebral haemorrhage.<sup>5,6</sup>

Methods to attenuate these responses, both pharmacological and physiological, have been extensively studied.<sup>7,8</sup> Treatment modalities include topical lignocaine spray, deeper planes of anaesthesia by inhalation/intravenous agents or narcotics, calcium channel blockers, a-2 agonists,  $\beta$ -2 blockers, vasodilators such as sodium-nitroprusside, nitroglycerine etc.<sup>9</sup> Although there are several methods, research is still ongoing.<sup>10</sup>

Esmolol is an ultra-short acting,  $\beta$  adrenergic receptor antagonist, with proven efficacy to provide haemodynamic stability during laryngoscopy and tracheal intubation.<sup>11</sup> The peak effect of esmolol is 2 minutes from the time of

intravenous administration and is metabolised by esterases resulting in short duration of action (10-15 minutes). These characteristics make esmolol a useful drug to blunt the response.

Dexmedetomidine,<sup>12</sup> a highly selective alpha-2 receptor agonist, decreases systemic noradrenalin release results in attenuation of sympathoadrenal responses and maintain haemodynamic stability during laryngoscopy and tracheal intubation.

This background study guided us to perform this randomized, prospective double blind study.

#### MATERIALS AND METHODS

After written informed consent of each patient and clearance from Institutional ethical committee a study was conducted at B. S. Medical College, Bankura. One hundred and twenty five (125) normotensive, ASA physical status I and II patients of either sex, aged 30-60 years, who were scheduled for elective upper abdominal surgeries under general anaesthesia (GA) requiring endotracheal (ET) intubation, were included in this study. Pre-anaesthetic checkup were done for all patients. Patients were excluded as per our exclusion criteria. On the night before surgery, patients were given 0.5mg alprazolam orally.

Pre-operative baseline vital parameters of patients including pulse rate (PR), systolic blood pressure (SBP), diastolic blood pressure (DBP), mean arterial pressure (MBP) and oxygen saturation (SpO2) were recorded. In operation theatre, Ringer's lactate was infused (6 ml/kg). Routine standard monitors such as pulse oximetry, electrocardiography (ECG) and non-invasive blood pressure were applied, and monitoring started. Premedication done ondansetron with intravenous 0.08 mg/kg and glycopyrrolate 0.004 mg/kg, midazolam 0.02 mg/kg 10 min before induction. The study drugs were premixed to a volume of 15 ml and were presented as coded syringes. The anaesthesiologists who prepared and who administered the medications were different. The patients were blinded to the treatment group and all recordings were performed by an anaesthesiologist blinded to the group allocation.

Computer generated random number table was used for patients allocation. The patients in control Group C (n=42)

received 15 ml of normal saline, Group E (n=41) received esmolol 2.0 mg/kg and Group D (n=42) received dexmedetomidine 1.0  $\mu$ g/kg as slow IV infusion over a period of 10 min. After pre-oxygenation the patients were induced with Thiopentone sodium 5 mg/kg and intubated after succinylcholine 2 mg/kg. After confirming the position and fixing the endotracheal tube anaesthesia was maintained with, 66% N<sub>2</sub>O in 33% oxygen and 1% sevoflurane in 6 l of fresh gas flow and vecuronium 0.08 mg/kg bolus followed by intermittent dose of 0.02 mg/kg. At the end of the surgery all patients were reversed with appropriate dose of neostigmine and glycopyrrolate. Patients were extubated after adequate recovery and then shifted to anaesthesia recovery room.

Vital parameters such as PR, SBP, DBP and MBP were recorded at baseline, after study drug infusion, after induction, immediately and 3, 5, 7 and 10 min after intubation. No surgical intervention was allowed throughout the study period of 10 min.

### RESULTS

Sample size was calculated based on primary objective of difference of mean arterial blood pressure (MBP) of 20 mmHg (SD 15) immediately after intubation between dexmedetomidine and esmolol group. Considering the type I error and power of the study as 5% and 90% respectively a total of 41 patients in each of the three groups was optimum sample size, expecting a dropout of 5%. All raw data were entered into a predesigned excel spreadsheet and analysed using standard statistical software SPSS-16. Chi square test was used to compare the three groups in terms of gender and one -way Analysis of variance (ANOVA) was used to compare them based on their age and body weight. The baseline parameter and the change of that parameter before and after intubation at different time intervals were compared between the three study groups by Analysis of Variance (ANOVA). Also, the parameters at different time intervals between any two among the three study groups were analysed by unpaired student's t test. A P value of less than 0.05 was considered statistically significant.

Parameter		Group			
Faranielei	Dexmedetomidine	Esmolol	Control		
Age (mean ± SD) in years	45.71 ± 9.354	46.19 ± 7.649	45.95 ± 7.774		
Body weight (mean ± SD) in kilograms	57.93 ± 9.295	$58.66 \pm 8.880$	58.71 ± 4.836		
Gender ratio (M:F)	22:20	21:20	20:22		
Table 1. Mean Age, Body Weight and Gender Ratio of the Study					

The table 1 shows the comparison of age, body weight and gender ratio between the three groups. There was statistically no significant difference in these demographic parameters between these three groups ( $p \ge 0.05$ ). Age of the study subjects range from 30 to 60 years. Body weight of the subjects range from 40-75 kg.



Figure 1. Gender Distribution in the Three Groups (Pie Diagram)

#### Comparison of Pulse Rate (PR)



Figure 2. Pulse Rate Comparison with Confidence Interval (95%)

The observations were recorded at baseline (PRB), after study drug infusion (PRI), after induction (PR2), immediately (PR3) and 3, 5, 7 and 10 minutes after intubation (PR4-7 respectively). The median values (ordinate) of the pulse rate are compared for each treatment group using linear charts for eight different time points (abscissa) – (PRb-PR7). The error bars represent 95% confidence internal to each median data point.

Parameters	Group E (N=41) (Mean ± SD)	Group D (N=42) (Mean ± SD)	t-value	P-value	Significance of difference*
PRb	83.73 ± 8.59	83.45 ± 6.55	-0.127	0.868	Not Significant
PR1	71.46 ± 7.90	74.43 ± 7.13	1.796	0.076	Not Significant
PR2	70.02 ± 7.52	71.79 ± 6.71	1.126	0.263	Not Significant
PR3	85.00 ± 9.19	72.88 ± 6.77	-6.852	<0.001*	Significant
PR4	88.29 ± 9.49	73.93 ± 6.69	-8.100	<0.001*	Significant
PR5	85.56 ± 7.92	72.33 ± 6.61	-8.272	<0.001*	Significant
PR6	80.63 ± 5.89	72.90 ± 6.35	-5.696	<0.001*	Significant
PR7	78.51 ± 6.05	73.88 ± 5.98	-3.484	0.001*	Significant
Table 2. Comparison of Pulse Rate between Group E & Group D					

Table 2. Comparison of means of the pulse rate at different stages between the patients receiving Esmolol (Group E) and the patients receiving Dexmedetomidine (Group D). (\* Statistical significance of difference is considered at a confidence interval of 95%).



Figure 3. Comparison of Pulse Rate

### **Pulse Rate**

For both Group D & E the mean pulse rate decreased below baseline after study drug infusion. Both the groups were also comparable after study drug infusion and after induction. They developed a significant difference among themselves only on/after intubation. The mean pulse rate of Group D never rose above baseline during the entire period of observation. In group E the highest rise was seen 3 minutes post intubation (5.8% of baseline) and ended below baseline after 7 minutes of intubation.

Esmolol was significantly effective than control in minimizing the pulse rate response though not as effective as dexmedetomidine, in the aforesaid dosage.

#### Comparison of Systolic Blood Pressure (SBP)



Figure 4. SBP Comparison with Confidence Interval (95%)

Figure 4- The median values (ordinate) of the SBP are compared for each treatment group using linear charts for eight different time points (abscissa) - (PRb-PR7). The error bars represent 95% confidence internal to each median data point.

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Parameters	Group E (N=41) (Mean ± SD)	Group D (N=42) (Mean ± SD)	t-value	P-value	Significance of difference*
SBPb	135.34 ± 11.79	132.83 ± 9.61	-1.063	.291	Not Significant
SBP1	121.95 ± 10.97	121.45 ± 7.00	248	.805	Not Significant
SBP2	113.17 ± 11.48	113.69 ± 8.05	.239	.811	Not Significant
SBP3	$128.46 \pm 11.66$	114.48 ± 7.82	-6.428	<0.001*	Significant
SBP4	135.46 ± 12.86	116.05 ± 8.26	-8.084	<0.001*	Significant
SBP5	120.59 ± 11.16	115.90 ± 7.88	2.269	.026*	Significant
SBP6	118.80 ± 9.48	113.60 ± 7.24	-2.934	.004*	Significant
SBP7	$118.02 \pm 6.40$	111.38 ± 7.24	-4.502	<0.001*	Significant
Table 3. Comparison of SBP between Group E & Group D					

Table 3 shows the comparison of means of the SBP at different stages between the patients receiving Esmolol (Group E) and the patients receiving Dexmedetomidine (Group D). (\* Statistical significance of difference is considered at a confidence interval of 95%).



### Systolic Blood Pressure (SBP)



Figure 5. SBP Comparison of Systolic Blood Pressure (SBP)

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### Systolic Blood Pressure (SBP)

For both Group D & E the mean SBP decreased below baseline after study drug infusion and further after induction. Both the groups were comparable after study drug infusion and after induction other than at baseline. They developed a significant difference among themselves only on/after intubation. The mean SBP of Group D never rose above baseline during the entire period of observation. The mean SBP of Group E was highest 3 minutes post intubation, and decreased significantly thereafter. The mean SBP in both Group E & D were significantly lower than the SBP values of Group C in all observations other than baseline.

Therefore, although both Esmolol and Dexmedetomidine are effective in neutralizing the surge in SBP, the study here shows that Dexmedetomidine was significantly more effective than Esmolol in attenuating the reflex response to intubation, in the aforesaid dosage for SBP.

# Comparison of Rate Pressure Product (RPP)



Figure 6 shows the median values (ordinate) of the RPP are compared for each treatment group using linear charts for eight different time points (abscissa) –(RPPb-RPP7). The error bars represent 95% confidence internal to each median data point.

Parameters	Group E (N=41) (Mean ± SD)	Group D (N=42) (Mean ± SD)	t-value	P-value	Significance of difference*
RPPb	11257.32 ± 778.61	11101.67 ± 1339.23	156	.876	Not Significant
RPP1	8646.38 ± 532.37	9039.38 ± 997.53	2.23	.028	Significant
RPP2	7863.41 ± 597.78	8167.24 ± 993.11	1.684	.096	Not Significant
RPP3	10848.88 ± 864.83	8353.93 ± 1059.80	11.73	< 0.001	Significant
RPP4	11865.59 ± 789.51	8590.69 ± 1104.97	-15.313	< 0.001	Significant
RPP5	10257.17 ± 736.75	8394.48 ± 1063.63	-9.296	< 0.001	Significant
RPP6	9538.85 ± 523.16	8287.57 ± 958.39	-7.396	< 0.001	Significant
RPP7	9237.98 ± 460.44	8232.24 ± 891.97	-6.549	< 0.001	Significant
Table 4. Comparison of RPP between Group E & Group D					

Table 4 - Comparison of means of the RPP at different stages between the patients receiving Esmolol (Group E) and the patients receiving Dexmedetomidine (Group D). (\* Statistical significance of difference is considered at a confidence interval of 95%).



RPP3\* RPP4\* RPP5\* RPP6\*

GROUP D

GROUP E

RPP7

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RPPb

RPP1\* RPP2\*

GROUP C

5000

0



Figure 7. Rate Pressure Product (RPP)

### **Rate Pressure Product (RPP)**

For both Group D & E the RPP decreased below baseline after study drug infusion and further after induction. Both the groups were comparable after induction other than at baseline. They developed a significant difference among themselves thereafter.

There was a 37% after induction rise in RPP on attempt/during intubation in Group E. The RPP was highest in Group E; 3 minutes post intubation (rise by 5% of baseline). The RPP in Group E decreased thereafter and ended below baseline 5 minutes post intubation.

In contrast the mean RPP of Group D never rose above baseline during the entire period of observation and also the rise was only 2.2% during intubation from post induction values. The mean RPP in both Group E & D were significantly lower than the RPP values of Group C in all observations other than baseline.

Therefore, although both Esmolol and Dexmedetomidine were effective in neutralizing the surge in RPP, the study here shows that Dexmedetomidine was significantly more effective than Esmolol in attenuating the reflex response to intubation, in the aforesaid dosage.

#### DISCUSSION

The pressor response to laryngoscopy and endotracheal intubation though transient, may be potentially hazardous due to reflex sympathetic discharge caused by pharyngeal stimulation. Transient hypertension and tachycardia are probably of no consequence in healthy individuals but either or both may be hazardous to those with hypertension, myocardial insufficiency and cerebrovascular disease. These changes are the maximal at 1 minute after intubation and last for 5-10 minutes. Prophylaxis include topical lignocaine sprays, deeper planes of anaesthesia by inhalational agents; narcotics, calcium channel blockers, vasodilators such as sodium nitroprusside; nitroglycerin etc,<sup>1</sup> but they have got side effects such as sedation, respiratory depression, hypotension and bradycardia.

The analgesic, sedation, anxiolytic, sympatholytic and blunting of exaggerated haemodynamic responses by administration of dexmedetomidine are being extensively studied and are mainly mediated by the activation of a-2 receptors located in the postsynaptic terminals in the central nervous system (CNS), which causes decreased neuronal activity and augmentation of the vagal activity. Esmolol is a water soluble, rapid onset, ultra-shortacting, selective beta-adrenergic receptor antagonist with proven efficacy to provide haemodynamic stability during laryngoscopy and tracheal intubation. It has a half-life of nine minutes and without severe side effects. Esmolol seems to be an appropriate selection for attenuating the haemodynamic response to laryngoscopy and tracheal intubation, because of its cardioselectivity, rapid onset of action and short elimination half-life.

Miller et al<sup>13</sup> reported that 100 mg of single bolus dose of esmolol was effective for controlling the haemodynamic response to tracheal intubation. In another study, Liu et al. who used esmolol infusion to control haemodynamic responses associated with intubation, found significant decreases in PR and SBP, in the esmolol\_treated patients compared to the placebo group.<sup>14</sup> Oxorn et al.<sup>15</sup> concluded that esmolol in bolus doses of 100 mg and 200 mg affects solely the chronotropic response in a significant manner. Kindler et al. reported that esmolol administration before laryngoscopy was sufficient to control PR after intubation, but it did not affect sbp.<sup>16</sup> Similar to the above studies in our study Esmolol was significantly more effective than normal saline in obtunding the surge in PR and blood pressure.

al.17 et studied Scheinin that 0.6 µq/kq dexmedetomidine decreased, but not totally suppressed, the haemodynamic response to tracheal intubation in healthy individuals. Keniya et al. stated that the pre-treatment with dexmedetomidine 1.0 µg/kg attenuated, but not totally obtunded the cardiovascular response to tracheal intubation after induction of anaesthesia.18 In this study, the percentage rise in PR and other haemodynamic parameters between induction and post intubation were minimal in the dexmedetomidine group compared to Control and Esmolol.

Bradycardia and hypotension have been reported in some studies pertaining to the effect of dexmedetomidine administration.<sup>19,20</sup> But in our study, neither bradycardia nor hypotension was observed in the patients. Dexmedetomidine has been used I V in doses ranging from 0.1 to 10 µg/kg/h but higher doses have been associated with a significant increase in incidence of bradycardia and hypotension. Rapid administration of dexmedetomidine might produce tachycardia, bradycardia and hypertension followed administered by hypotension. We dexmedetomidine, 1.0 µg/kg slowly, over 10 mins.

Yallapragada SV et al.<sup>21</sup> studied on the efficacy of dexmedetomidine with that of esmolol in attenuating

laryngoscopic and intubation response after rapid sequence induction. In their study they concluded that dexmedetomidine is superior to esmolol in attenuating the haemodynamic response to laryngoscopy and tracheal intubation. Subsequently, Reddy SV et al.<sup>22</sup> again studied dexmedetomidine versus esmolol to attenuate the haemodynamic response. Similar to this study, they found that the suppression in cardiovascular responses was greater with dexmedetomidine 1.0  $\mu$ g/kg than that resulted from infusion of esmolol 2.0 mg/kg.

Monitoring of PR and ECG has shown no evidence of myocardial insult in any of the patient in any group in our study.

In our study infusion of dexmedetomidine 1.0 µg/kg prior to induction of anaesthesia suppressed the haemodynamic response to tracheal intubation in normotensive patients. This suppression in cardiovascular responses was found to be greater with dexmedetomidine infusion than with esmolol. In the present study the haemodynamic response to laryngoscopy and intubation were studied for a period of 10 minutes as this is the average period for which haemodynamic changes are believed to last. It was found that with this dose dexmedetomidine had better control over PR, SBP, DBP and MBP. On comparison between the two groups, the heart rate, blood pressure and rate pressure product was better controlled with dexmedetomidine than esmolol.

### CONCLUSION

In this study, infusion of dexmedetomidine  $1.0 \mu g/kg$  prior to induction of anaesthesia suppressed the haemodynamic response to tracheal intubation in normotensive patients. This suppression in cardiovascular responses was found to be greater with dexmedetomidine than that resulted from infusion of esmolol 2.0 mg/kg.

#### REFERENCES

- Tomori Z, Widdicombe J. Muscular, brochomotor and cardiovascular reflexes elicited by mechanical stimulation of respiratory tract. J Physiology 1969;200(1):25-49.
- [2] Forbes AM, Dally FC. Acute hypertension during induction of anaesthesia and endotracheal intubation in normotensive man. Br J Anaesth 1970;42(7):618-624.
- [3] Kihara S, Brimacombe J, Yaguchi Y, et al. Haemodynamic responses among three tracheal intubation devices in normotensive and hypertensive patients. Anesth Analg 2003;96(3):890-895.
- [4] Pyrs-Roberts, Greene LT, Meloche R, et al. Studies of anaesthesia in relation to hypertension II. Haemodynamic consequences of induction and endotracheal intubation. Br J Anaesth 1971;43(6):531-534.
- [5] Figueredo E, Garcia-Fuentes EM. Assessment of the efficacy of esmolol on the haemodynamic changes induced by laryngoscopy and tracheal intubation: a

meta-analysis. Acta Anaesthesiol Scand 2001;45(8):1011-1022.

- [6] Edwards ND, Alford AM, Dobson PMS, et al. Myocardial ischaemia during tracheal intubation and extubation. Br J Anaesth 1994;73(4):537-539.
- [7] Stoelting RK. Blood pressure and heart rate changes during short duration laryngoscopy for tracheal intubation: influence of viscous or intravenous lidocaine. Anesth Analg 1978;57(2):197-199.
- [8] Mikawa K, Obara H, Kusunoki M. Effect of nicardipine on the cardiovascular response to tracheal intubation. Br J Anaesth 1990;64:240-242.
- [9] Kovac AL. Controlling the haemodynamic response to laryngoscopy and endotracheal intubation. J Clin Anesth 1996;8(1):63-79.
- [10] da Silva Neto WV, Azevedo GS, Coelho FO, et al. Evaluation of haemodynamic variations during anesthetic induction in treated hypertensive patients. Rev Bras Anestesiol 2008;58(4):330-341.
- [11] Louizos AA, Hadzilia SJ, Davilis DI, et al. Administration of esmolol in microlaryngeal surgery for blunting the haemodynamic response during laryngoscopy and tracheal intubation in cigarette smokers. Ann Otol Rhinol Laryngol 2007;116(2):107-111.
- [12] Grewal A. Dexmedetomidine: new avenues. J Anaesthesiol Clin Pharmacol 2011;27(3):297-302.
- [13] Miller DR, Martineau RJ, Wynands JE, et al. Bolus administration of esmolol for controlling the haemodynamic response to tracheal intubation: the Canadian Multicentre Trial. Can J Anaesth 1991;38(7):849-858.
- [14] Liu PL, Gatt S, Gugino LD, et al. Esmolol for control of increases in heart rate and blood pressure during tracheal intubation after thiopentone and succinylcholine. Can Anaesth Soc J 1986;33(5):556-562.
- [15] Oxorn D, Knox JW, Hill J. Bolus doses of esmolol for the prevention of perioperative hypertension and tachycardia. Can J Anaesth 1990;37(2):206-209.
- [16] Kindler CH, Schumacher PG, Schneider MC, et al. Effects of intravenous lidocaine and/or esmolol on haemodynamic responses to laryngoscopy and intubation: a double-blind, controlled clinical trial. J Clin Anesth 1996;8(6):491-496.
- [17] Scheinin B, Lindgren L, Randell T, et al. Dexmedetomidine attenuates sympathoadrenal responses to tracheal intubation and reduces the need for thiopentone and peroperative fentanyl. Br J Anaesth 1992;68(2):126-131.
- [18] Keniya VM, Ladi S, Naphade R. Dexmedetomidine attenuates sympathoadrenal response to tracheal intubation and reduces perioperative anaesthetic requirement. Indian J Anaesth 2011;55(4):352-357.
- [19] Ben-Abraham R, Ogorek D, Weinbroum AA. Dexmedetomidine: a promising agent for anaesthesia and perioperative care. Isr Med Assoc J 2000;2(10):793-796.

- [20] Lawrence CJ, De Lange S. Effects of a single preoperative dexmedetomidine dose on isoflurane requirements and peri-operative haemodynamic stability. Anaesthesia 1997;52(8):736-744.
- [21] Yallapragada SV, Vidadala KS, Vemuri NN, et al. Comparison of the efficacy of dexmedetomidine with that of esmolol in attenuating laryngoscopic and

intubation response after rapid sequence induction. Anesth Essays Res 2014;8(3):383-387.

[22] Reddy SV, Balaji D, Ahmed SN. Dexmedetomidine versus esmolol to attenuate the haemodynamic response to laryngoscopy and tracheal intubation: a randomized double-blind clinical study. Int J Appl Basic Med Res 2014;4(2):95-100.