

A Comparison of Intravenous Infused Esmolol and Dexmedetomidine on Hemodynamic Response to Laryngoscopy and Endotracheal Intubation in Elective Laparoscopic Gynecological Surgery

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Abstract

Background: Rigid laryngoscopy and tracheal intubation remain the gold standard in airway management despite the emergence of new airway devices. These procedures ensure airway patency, ventilation control, and effective delivery of inhalation agents. However, they can stimulate sympathetic and sympatho-adrenal activities, causing temporary effects such as increased heart rate, blood pressure, intra-cranial pressure, and intra-ocular pressure. Complications like cardiac arrhythmia, myocardial ischemia, myocardial infarction, and cerebral hemorrhage may arise due to elevated catecholamine levels. Arterial hypertension during laryngoscopy and intubation primarily stems from increased cardiac output rather than systemic vascular resistance.

Methods: Several drugs and techniques have been used to manage the hemodynamic response to laryngoscopy and intubation. These include deepening anesthesia, omitting cholinergic medications, pre-treatment with Nitro-glycerine and Isosorbide dinitrate, Propranolol, Droperidol, Lidocaine administration, Beta-blocker Esmolol, Clonidine, Dexmedetomidine, Calcium channel blockers, Gabapentin, and opioids like Fentanyl and Remifentanyl. The choice of technique or drug depends on surgery necessity, duration, anesthetic technique, route of administration, and patient's medical condition.

Results: Opiates reduce sympathetic response via vasodilation, vasomotor depression, and vagal center stimulation. Beta blockers effectively prevent heart rate elevation compared to blood pressure response. Esmolol, a short-acting β -1 adrenergic antagonist, ensures hemodynamic

stability during laryngoscopy and intubation. Dexmedetomidine, a selective α_2 -adrenergic receptor agonist, provides analgesic and sedative effects when administered intrathecally, epidurally, or intravenously. It reduces cardiovascular response to tracheal intubation and lessens the need for thiopentone induction and perioperative fentanyl.

Conclusion: This study aimed to compare the efficacy of Esmolol and Dexmedetomidine, belonging to different drug classes, in attenuating pressure response during laryngoscopy and endotracheal intubation for elective laparoscopic gynecological surgeries under general anesthesia.

Keywords: Airway management, laryngoscopy, tracheal intubation, sympathetic response, hemodynamic response, anesthesia, Esmolol, Dexmedetomidine, laparoscopic gynecological surgeries, pressure response.

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Introduction

Despite the emergence of new airway devices in recent year, rigid laryngoscopy and tracheal intubation still remain gold standard in airway management. It maintains the patency of airway, it controls the ventilation, and helps in delivering inhalation agent to the patient.

Direct laryngoscopy and tracheal intubation causes increase in sympathetic and sympathoadrenal activities to the mechanical stimulation of larynx and trachea. Increase in heart rate and blood pressure are the most common effects seen following laryngoscopy and intubation it also increase intra cranial pressure and intra ocular pressure because of significant increase in circulating catecholamine's. These effects are temporary but it may lead to cardiac arrhythmia, myocardial ischemia, myocardial infarction, and cerebral haemorrhage.^[1-3] The arterial hypertension is due to increase in cardiac output rather than increase in SVR and associate with the transient rise in CVP (central venous pressure)[4]

Various drugs and techniques have been described to control the hemodynamic response to the laryngoscopy and intubation, such as deepening of anaesthesia, omitting cholinergic medications, pre-treatment with Nitro-glycerine and Isosorbide dinitrate (Elkayam et al)[5] Propranolol (McCammon

et al)[6], Droperidol (Curren J et al)[7], administration of Lidocaine (Stoelting et al)[8], Beta blocker-Esmolol[9], Clonidine, Dexmedetomidine (Scheinin B et al)[10], Calcium channel blockers, Gabapentin, Opioids (Dahlgaren et al)[11] like Fentanyl and Remifentanyl can be used for its prevention. The technique or drug of choice depends upon the necessity and duration of surgery, choice of anaesthetic technique, route of administration, and medical condition of patient.

Opiates decrease sympathetic response to noxious stimulation by vasodilation, depression of vasomotor and stimulation of vagal centre.[12-14]

Beta blockers with negative chronotropic, antihypertensive, antiarrhythmic and anti-ischemic properties make these agents more effective in preventing rise in heart rate than pressure response and blunts the hemodynamic responses to tracheal intubation in treated hypertensive patients.

Esmolol is a potent ultra-short acting (half-life 9 min) cardio selective adrenoreceptor selective β_1 competitive antagonist. It also obtunds the cardiac response to exercise and other stimuli. Esmolol is short acting because of its esterase induced rapid metabolic inactivation. It can be an efficient agent to

provide hemodynamic stability during laryngoscopy and intubation.[15]

Dexmedetomidine is a selective Alpha₂-adrenoreceptor agonist with a α_2/α_1 selectivity (alpha-2:alpha-1=1600:1). It has analgesic and sedative properties, when administered intrathecally, epidural or intravenously as an adjuvant.

It activates central alpha-2 adrenergic receptor and inhibits the release of norepinephrine from the adrenergic neurons, CNS and from postganglionic fibres of cardiac nerves and an increase of parasympathetic tone. This results in decrease in blood pressure, heart rate, cardiac output and peripheral venous resistance.[16]

dexmedetomidine can reduce the cardiovascular response to tracheal intubation and need of thiopentone for induction and perioperative fentanyl requirement.[17]

With above background the present study was designed to compare the efficacy of two different class of drugs Esmolol versus Dexmedetomidine for attenuation of pressure response (mean heart rate, mean systolic blood pressure, mean diastolic blood pressure, mean arterial pressure, mean rate pressure product) to laryngoscopy and endotracheal intubation in patient undergoing elective laparoscopic gynaecological surgeries under general anaesthesia.

Material and Method

This Hospital based, Randomized, double-blind, Superiority type of interventional study was conducted at the tertiary care center which is one of the largest tertiary care center of Northern India.

This study included all the Patients Undergoing laparoscopic gynecological surgery, aged 20-50 year of either gender with weighing of 40-70kg. Patients with anticipated difficult airway, chronic disease

(like hypertension, diabetes, hepatic disease), patients on beta blockers, pregnant and lactating women also patients taking laryngoscopy time more than 20 seconds excluded from this study.

Sample size : A Sample of 30 cases in each group calculated at 95% confidence interval & 80% power to verify the expected difference of 14.6 ± 1.92 in mean heart rate in two groups after 10 minutes of receiving Esmolol and Dexmedetomidine in patients undergoing elective gynaecological laparoscopic surgery.

A total of 60 eligible subjects were recruited consecutively till sample size was achieved. Subjects were randomly allocated into one of the following two groups using block randomization method to ensure the equal number of subjects in both group.

Group A- Patients received intravenous infusion of esmolol 1.5 mg/kg diluted in 20 ml NS over 1 minutes. Group B-Patients received intravenous infusion of dexmedetomidine 1 μ g/kg diluted in 20 ml NS over 10 minutes.

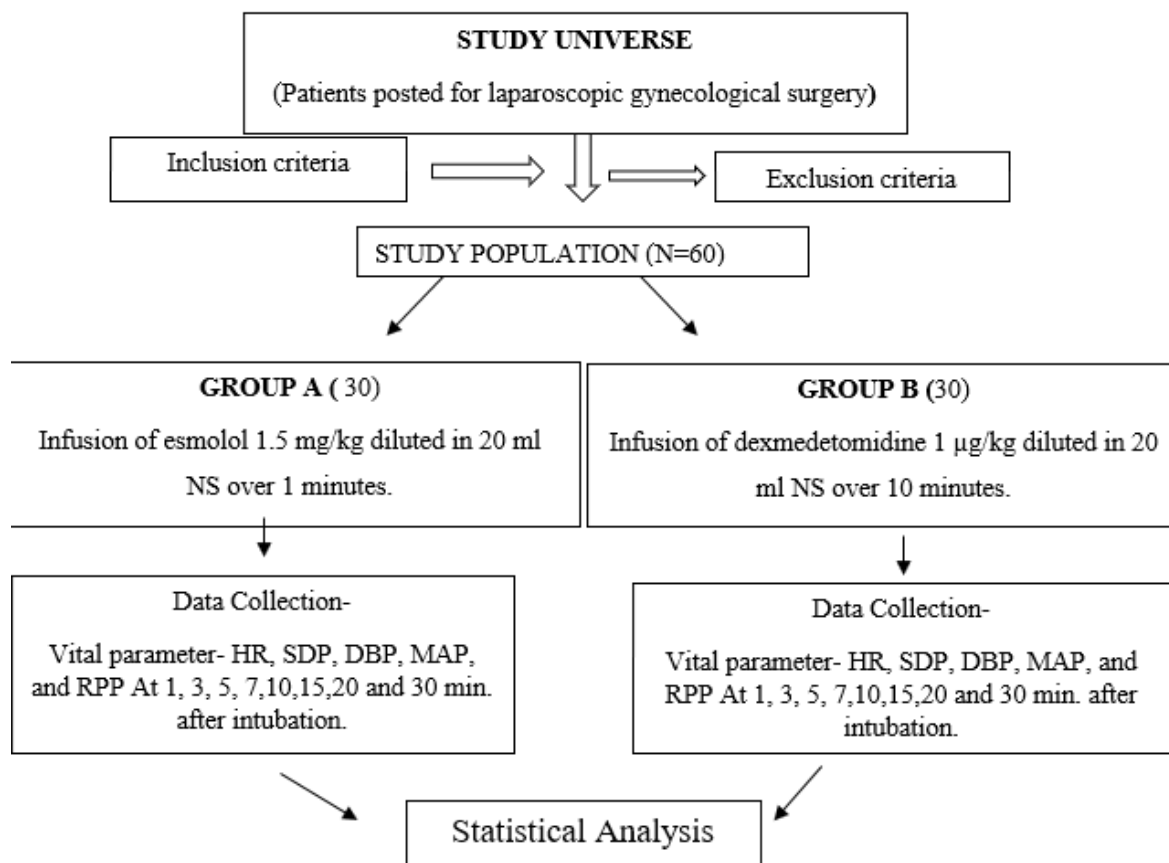
Allocation concealment was ensured using opaque sealed envelope method for group allocation. Allocation was done by a person not involved directly in the research to avoid selection bias. Neither the anesthetist nor the patient was aware of the groups and the drugs used (Double blind).

Patients heart rate, systolic blood pressure (SBP) and diastolic blood pressure (DBP), mean arterial pressure (MAP), respiratory rate (RR), arterial oxygen saturation (SPO₂), RPP, temperature were measured at 5 minute after starting infusion, after completion of infusion, after induction, , at 1 minute after intubation, at 3 minute after intubation, at 5 minute after intubation, at 10 minute after intubation, at 15 minute after intubation and at 30 minute after intubation.

All patients were subjected to standard Pre anesthetic checkup before the surgery

including detailed history, examination, vitals, routine investigations and markers.

Consort flow chart of the study



Statistical analysis: Quantitative data were summarized as mean and standard deviation and analyzed using Student “t-test”. Frequencies and percentage were analyzed using chi square test. A “p-value” <0.05 was taken as statistically significant. SPSS trial version 22 was used for statistical test.

Result

Both the groups were comparable in relation to their baseline characteristics like age, gender, ASA grade and type of surgery (Table 1)

Table 1: Age, gender, ASA grade, type of surgery wise distribution of patients.

| | | Group D | Group T | p-value |
|------------------|-----------------|--------------|--------------|---------|
| Age (M±S.D.) | In Years | 31.226± 5.b | 30.66 ± 4.77 | 0.450 |
| Weight (M ±S.D.) | In kg | 56.83 ± 5.34 | 58.88 ± 4.42 | 0.111 |
| ASA | Grade I | 30 (100%) | 30 (100%) | <0.05 |
| | >Grade I | 0 | 0 | |
| Type of surgery | Infertility | 73.33% | 70% | |
| | Aub | 3.33% | 0 | |
| | Cervical Biopsy | 3.33% | 3.33% | |

| | | | |
|--|-----------------------|-------|-------|
| | Cervical Incompetance | 6.67% | 3.33% |
| | Endometrial Cyst | 3.33% | 3.33% |
| | Laparotomy | 3.33% | 0 |
| | Pid | 3.33% | 6.66% |
| | Polyp | 3.33% | 3.33% |
| | molar pregnancy | 0 | 3.33% |
| | LT Ovarian Cyst | 0 | 3.33% |
| | LT hydrosalpinx | 0 | 3.33% |

Table 2: Comparison of heart rate in Group A and Group B

| Heart Rate | Group A | | Group B | | Test of Significance | | |
|----------------------------|---------|-------|---------|-------|----------------------|---------|--------------|
| | Mean | SD | Mean | SD | T Test Value | P Value | Significance |
| Base Line | 90.87 | 6.998 | 91.03 | 6.935 | -0.163 | 0.871 | NS |
| 5 Min After Infusion Start | 90.37 | 6.82 | 89.27 | 6.41 | 1.229 | 0.236 | NS |
| After Induction | 86.43 | 4.202 | 82.20 | 4.55 | 3.75 | <0.001 | S |
| Just Before Intubation | 84.47 | 4.51 | 79.50 | 5.01 | 4.038 | <0.001 | S |
| 1 Min | 83.20 | 5.9 | 78.93 | 4.89 | 3.56 | <0.001 | S |
| 3 Min | 83.77 | 4.493 | 78.27 | 5.14 | 4.14 | <0.001 | S |
| 5 Min | 80.10 | 4.444 | 74.67 | 5.1 | 4.39 | <0.001 | S |
| 10 Min | 75.70 | 4.450 | 72.47 | 5.46 | 2.512 | 0.015 | S |
| 15 Min | 75.17 | 4.264 | 72.77 | 6.1 | 1.98 | 0.59 | S |
| 30 Min | 76.13 | 5.1 | 73.13 | 4.15 | 2.49 | 0.015 | S |

Table 3: Comparison of SBP in Group A and Group B

| SBP | Group A | | Group B | | Test of Significance | | |
|------------------------|---------|--------|---------|-------|----------------------|---------|--------------|
| | Mean | SD | Mean | SD | T Test Value | P Value | Significance |
| Base Line | 132.10 | 11.621 | 128.20 | 3.727 | 1.750 | 0.085 | NS |
| 5 Min After Infusion | 131.10 | 5.892 | 126.67 | 5.616 | 2.961 | 0.001 | S |
| After Induction | 127.73 | 7.277 | 110.27 | 6.2 | 10.05 | <0.01 | S |
| Just Before Intubation | 123.93 | 7.995 | 105.5 | 5.9 | 10.54 | <0.01 | S |
| 1 MIN | 115.50 | 6.580 | 102.90 | 4.496 | 8.700 | <0.01 | S |
| 3 MIN | 112.07 | 6.674 | 102.83 | 5.1 | 7.456 | 0.000 | S |
| 5 MIN | 118.03 | 6.950 | 101.07 | 4.112 | 12.45 | 0.000 | S |
| 10 MIN | 113.53 | 7.305 | 100.27 | 6.944 | 7.253 | 0.000 | S |
| 15 MIN | 111.60 | 8.548 | 99.67 | 5.061 | 6.652 | 0.000 | S |
| 30 MIN | 107.07 | 7.714 | 101.77 | 6.924 | 2.808 | 0.007 | S |

Table 4: Comparison of DBP in Group A and Group B

| DBP | Group A | | Group B | | Test of Significance | | |
|------------------------|---------|-------|---------|-------|----------------------|---------|--------------|
| | Mean | SD | Mean | SD | t-Value | p-value | Significance |
| Base Line | 90.77 | 9.457 | 86.03 | 4.047 | 2.520 | 0.014 | Ns |
| 5 Min After Infusion | 92.20 | 6.789 | 77.73 | 3.523 | 10.359 | <0.001 | S |
| After Infusion | 84.40 | 5.893 | 74.00 | 3.063 | 8.577 | <0.001 | S |
| After Induction | 80.43 | 4.651 | 72.00 | 3.040 | -8.313 | <0.001 | S |
| Just Before Intubation | 78.13 | 4.710 | 70.27 | 3.095 | 7.644 | <0.001 | S |
| 1 Min | 76.50 | 4.493 | 68.53 | 3.115 | 7.981 | <0.001 | S |
| 3 Min | 74.67 | 4.971 | 66.50 | 3.138 | 7.609 | <0.001 | S |
| 5 Min | 71.83 | 4.186 | 64.70 | 2.879 | 7.691 | <0.001 | S |
| 7 Min | 69.37 | 4.089 | 63.43 | 2.763 | 6.585 | <0.001 | S |
| 10 Min | 68.90 | 7.554 | 61.70 | 2.680 | 4.920 | <0.001 | S |
| 15 Min | 59.90 | 2.618 | 57.90 | 2.618 | 5.534 | <0.001 | S |
| 20 Min | 66.53 | 6.021 | 58.53 | 2.675 | 6.111 | <0.001 | S |
| 30 Min | 58.53 | 2.675 | 57.10 | 2.670 | 4.948 | <0.001 | S |

Table 5: Comparison of MAP in GROUP A and GROUP B

| MAP | Group A | | Group B | | Test of Significance | | |
|----------------------|---------|------|---------|-------|----------------------|---------|--------------|
| | Mean | SD | Mean | SD | t test value | P value | Significance |
| Base line | 105.54 | 5.4 | 104.09 | 5.46 | 1.123 | 0.266 | NS |
| 5 min after infusion | 104.83 | 5.9 | 99.10 | 5.48 | 3.89 | <0.001 | S |
| After infusion | 98.84 | 4.9 | 82.76 | 5.23 | 12.289 | <0.001 | S |
| After induction | 96.20 | 4.5 | 81.42 | 4.89 | 12.182 | <0.001 | S |
| before intubation | 93.40 | 6.1 | 79.77 | 5.01 | 9.458 | <0.001 | S |
| 1 Min | 91.83 | 6.2 | 78.66 | 6.944 | 7.776 | <0.001 | S |
| 3 Min | 89.47 | 5.46 | 76.94 | 5.06 | 9.219 | <0.001 | S |
| 5 Min | 87.23 | 5.48 | 76.82 | 4.64 | 7.941 | <0.001 | S |
| 7 Min | 84.66 | 5.23 | 75.63 | 6.92 | 5.702 | <0.001 | S |
| 10 Min | 83.78 | 4.89 | 74.22 | 6.94 | 6.168 | <0.001 | S |
| 15 Min | 81.56 | 5.01 | 73.16 | 5.01 | 6.494 | <0.001 | S |
| 20 Min | 79.00 | 6.10 | 72.79 | 4.66 | 4.431 | <0.001 | S |
| 30 Min | 77.36 | 5.65 | 71.99 | 6.94 | 3.287 | 0.007 | S |

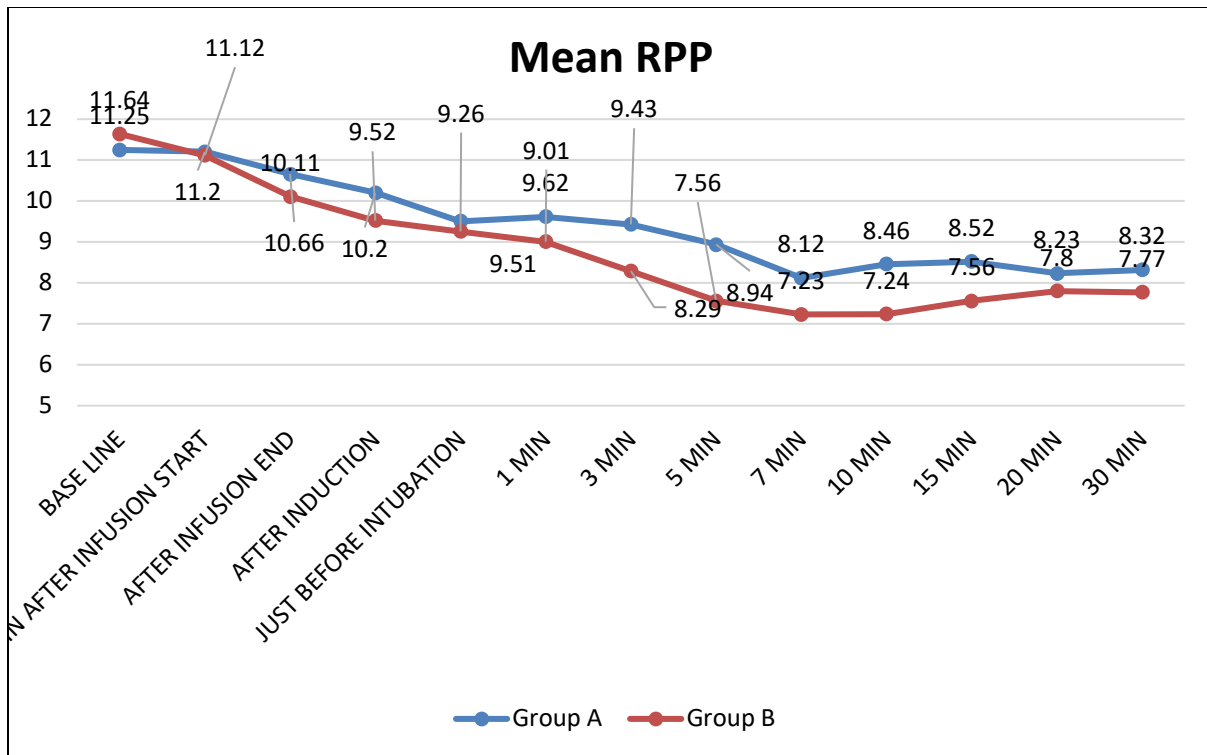


Figure 1: Comparison of RPP in Group A and Group B

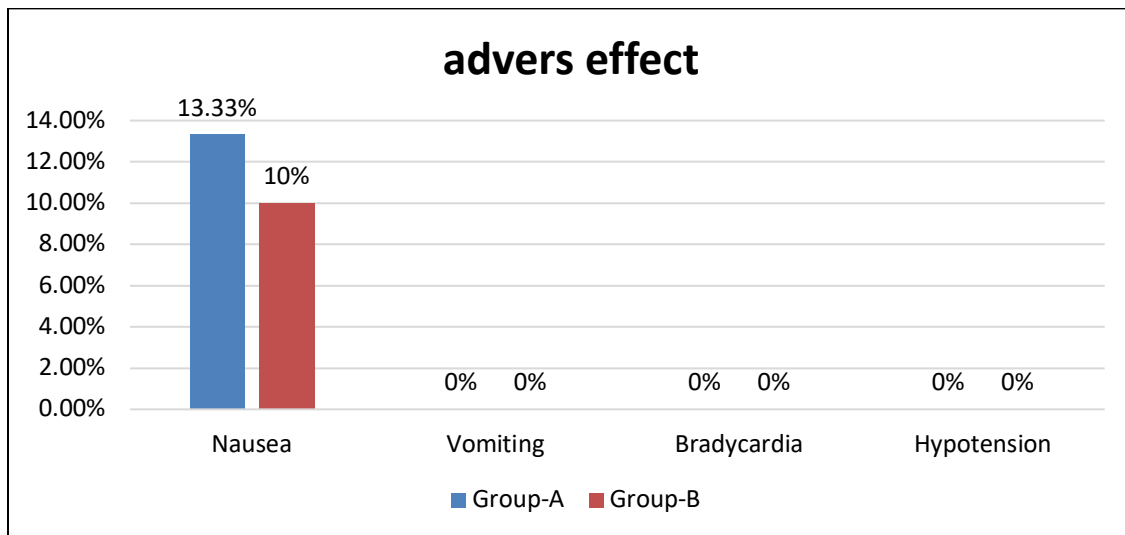


Figure 2: Adverse effects.

In reference to hemodynamic parameter Heart rate, SBP,DBP, MBP and RPP were seen significantly more downgrade (“p value<0.05) by Group B or in dexmedetomidine group as compare to Group A or Esmolol group at 5 minute after starting infusion, after completion of infusion, just before intubation, at 1 minute after intubation, at 3 minute after intubation, at 5 minute after intubation, at 10 minute after intubation, at 15 minute after intubation, at 30 minute after intubation.(table 2-5 and figure 1)

Only 13.3% patients in group A and 10% in group B reported nausea while none of the case shows side effect like vomiting, bradycardia, and hypotension. (Figure 2)

Discussion

Direct laryngoscopy and tracheal intubation causes increase in sympatho-adrenal activities due to the mechanical stimulation of larynx and trachea.

Various techniques and drugs have been described to control the hemodynamic response to the laryngoscopy and intubation in order to prevent any rise in myocardial work load and oxygen demand as well as to preserve the perfusion of vital organs.

The present study was undertaken to compare the efficacy and safety profile of these two different classes of drug.

In our study the demographic data and type of surgery were comparable in both the groups thus alleviating confounding factor.

In this study Heart rate, SBP, DBP, MBP and RPP were seen significantly lower (p value <0.05) in the Group B or in dexmedetomidine group as compare to Group A or Esmolol group at different time interval.

Reddy *et al* (2014)[18] studied consented 90 adult. Group E received 2.0 mg/kg of esmolol and Group D received 1.0 μ g/kg of dexmedetomidine, intravenously over 10 min and 3 min before induction of general anesthesia.

Both the drugs attenuated the pressure response. Of the two drugs administered, dexmedetomidine provided a consistent, reliable and effective attenuation of haemodynamic pressure when compared to esmolol (P value .001).

Shrivastav *et al* (2015) also compared dexmedetomidine (group D) 1 μ g/kg and esmolol (group E) 1.5 mg/kg and found that dexmedetomidine is more effective than esmolol for attenuating heart rate, SBP, DBP and MBP response to laryngoscopy and

intubation in elective neurosurgical patients. (P value 0.001). Selvaraj V *et al* (2016)[19] also observed the similar result with esmolol and dexmedetomidine on patients.

Similarly Gupta HB *et al* (2016)[20] and Sharma *et al* (2018) concluded that the SBP, DBP and MBP at 1 min, 3 min, 5min and 10 min after intubation was significantly lower in the dexmedetomidine group as compared to the esmolol group.

We observed, only 13.3% patients in group A and 10% in group B reported nausea while none of the case shows side effect like vomiting, bradycardia, and hypotension.

our study supported by Gupta HB *et al*[20] and T shering *et al* conclude that effect of dexmedetomidine on attenuation of pressor response to laryngoscopy and intubation is more pronounced than esmolol. The effectiveness of dexmedetomidine over esmolol makes it a useful drug for attenuation of hemodynamic response to laryngoscopy and intubation as a routine.

Conclusion

The dexmedetomidine infusion(1mcg/kg) was better than esmolol infusion (1.5mg/kg) in blunting the intubation response with better hemodynamic stability. There was no statistical significant difference of nausea in both the groups. None of the participants had experienced vomiting, hypotension& bradycardia. So the dexmedetomidine infusion was better than esmolol infusion.

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