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## RESEARCH ARTICLE

### A COMPARISON OF ETOMIDATE AND THIOPENTONE AS INDUCTION AGENTS IN LAPAROSCOPIC SURGERIES

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#### ABSTRACT

This study was designed to compare the hemodynamic effects of etomidate and thiopentone during induction in laparoscopic surgeries. Stimulation of laryngeal and tracheal tissues during intubation causes catecholamine discharge with an increase in sympathico-adrenergic activity and also an increase in systemic blood pressure and heart rate. These increases begin with laryngoscopy. Various induction agents are used for laparoscopy. Etomidate is characterized by hemodynamic stability, minimum respiratory depression and cerebral protective effects. Its lack of effect on sympathetic nervous system and its effect on increased coronary perfusion makes it an induction agent of choice. Thiopentone is a barbiturate derivative having rapid onset of action and rapid awakening with single induction dose due to rapid uptake.

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## INTRODUCTION

Laparoscopic surgery is minimal invasive surgery that is associated with several advantages like smaller scar, reduced bleeding, reduced postoperative pain and infection which contributes to shorter hospital stay. CO<sub>2</sub> insufflation will cause increased abdominal pressure and hypercapnia which may contribute to undesirable hemodynamic changes. Laparoscopy leads to ventilatory changes as the pneumoperitoneum created decreases thoraco-pulmonary compliance by 30%-50% in healthy and obese patients. Reduction in functional residual capacity and development of atelectasis due to elevation of the diaphragm and changes in the distribution of pulmonary ventilation and perfusion from increased airway pressure can be expected. Various induction agents are used for laparoscopy. Etomidate, carboxylated imidazole is characterised by hemodynamic stability, minimal respiratory depression and cerebral protective effects. Its lack of effect on sympathetic nervous system and its effect of increased coronary perfusion makes it an induction agent of choice. Thiopentone is a barbiturate derivative having rapid onset of action and rapid awakening with single induction dose due to rapid uptake.

**Aims and objectives:** The aim of this study is to compare and evaluate the intraoperative hemodynamic changes after induction of anesthesia with Etomidate and Thiopentone for patients undergoing laparoscopic surgeries. Objective is to compare hemodynamic parameters –Pulse, Systolic BP, Diastolic BP, Mean arterial pressure.

## MATERIALS AND METHODS

Following the approval by hospital ethics committee, after obtaining written, informed consent from patient's relatives, study was done. Sixty patients, aged 20-70 years, either sex, ASA grade I-II-III, scheduled to undergo laparoscopic surgery under general anaesthesia were included in the study. Preoperative assessment was done and investigations were noted. In operation theatre intravenous cannula of proper size was inserted into the largest vein on the forearm and an infusion of DNS was started. All the patients were pre-medicated with intravenous glycopyrrolate 0.004mg/kg, ondansetron 0.15mg/kg, fentanyl 2µg/kg. ECG, NIBP, SpO<sub>2</sub> was monitored. Base line hemodynamic parameters were recorded. All the patients were preoxygenated with 100% O<sub>2</sub> for 3mins. Now patients were randomly divided to one of the following two groups 30 patients each as per study drug injected.

Group 1: Patient receiving etomidate 0.5mg/kg.  
Group 2: - Patient receiving thiopentone 5mg/kg.

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Patient was induced with either of the above induction agents, intubation was done with inj. succinylcholine 2mg/kg. Anaesthesia was maintained with oxygen and inhalational agent Sevoflurane. Titration of inhalational agents was targeted to maintain MAP within 20% of baseline values. Muscle relaxation was maintained by intermittent doses of Vecuronium bromide. Intraoperative ECG NIBP spo2 EtCo2 was monitored .

On completion of surgery , patient was reversed using Inj. Glycopyrrolate 0.008mg/kg IV and Inj. Neostigmine 0.05mg/kg IV. All observation were recorded and results were analysed statistically. Data was expressed as mean  $\pm$  SD. For comparing data between 2 groups, Student-t test (analysis of variance) was used and p value < 0.05 was interpreted as clinically significant.

## OBSERVATION AND RESULTS

In our study total 60 patients were selected. 30 patients received Thiopentone and rest 30 patients received Etomidate as intravenous induction agent. All the patients completed the study.

Group E (n=30) received Etomidate  
Group T (N=30) received Thiopentone  
Observations were compiled and result were analysed statistically.

**Table 1. Pulse rate**

	Group-E (N=30)	Group-T (N=30)	P Value
BASELINE	77.9 $\pm$ 6.01	75.8 $\pm$ 4.93	0.1444
PREMEDICATION	81.76 $\pm$ 6.98	80.56 $\pm$ 5.24	0.4545
INDUCTION	79.83 $\pm$ 5.26	80.3 $\pm$ 6.77	0.7650
INTUBATION	82.56 $\pm$ 5.72	90.63 $\pm$ 6.80	<0.0001
1 MINS	79.5 $\pm$ 6.38	94.66 $\pm$ 5.04	<0.0001
3 MINS	80.76 $\pm$ 5.96	93.1 $\pm$ 4.12	<0.0001
5 MINS	80.73 $\pm$ 4.84	91.66 $\pm$ 5.95	<0.0001
15 MINS	78.73 $\pm$ 5.67	87.93 $\pm$ 5.57	<0.0001
30 MINS	80.56 $\pm$ 5.63	82.26 $\pm$ 6.20	0.2708

**Table 2. Systolic Blood Pressure**

	Group-E (N=30)	Group-T (N=30)	P Value
BASELINE	118.9 $\pm$ 10.06	119.1 $\pm$ 8.26	0.6430
PREMEDICATION	127.6 $\pm$ 9.13	124.2 $\pm$ 9.21	0.1564
INDUCTION	120.2 $\pm$ 6.61	124.5 $\pm$ 9.92	0.0529
INTUBATION	120.9 $\pm$ 6.72	125.1 $\pm$ 7.79	0.0292
1 MINS	121.3 $\pm$ 7.57	128.5 $\pm$ 9.41	0.0018
3 MINS	120.1 $\pm$ 7.58	127.03 $\pm$ 7.60	0.0008
5 MINS	121.8 $\pm$ 6.9	126.43 $\pm$ 8.59	0.0308
15 MINS	122.1 $\pm$ 6.84	124.8 $\pm$ 8.50	0.1805
30 MINS	121.6 $\pm$ 8.47	123.2 $\pm$ 8.84	0.4770

**Table 3. Diastolic Blood Pressure**

	Group-E (N=30)	Group-T (N=30)	P Value
BASELINE	80.5 $\pm$ 7.71	81.7 $\pm$ 6.07	0.5056
PREMEDICATION	84.9 $\pm$ 7.61	84.1 $\pm$ 6.84	0.1564
INDUCTION	81.7 $\pm$ 6.75	82.6 $\pm$ 6.29	0.6701
INTUBATION	82.2 $\pm$ 6.43	81.8 $\pm$ 5.08	0.7901
1 MINS	82.6 $\pm$ 5.47	89.6 $\pm$ 5.61	<0.0001
3 MINS	80.6 $\pm$ 6.35	87.7 $\pm$ 7.96	<0.0001
5 MINS	81.1 $\pm$ 5.85	86.5 $\pm$ 5.13	0.0003
15 MINS	81.3 $\pm$ 5.16	85.1 $\pm$ 3.58	0.0016
30 MINS	81.7 $\pm$ 6.86	84.1 $\pm$ 5.95	0.1531

**Table 4. Mean Blood Pressure**

	Group-E (N=30)	Group-T (N=30)	P Value
BASELINE	93.4 $\pm$ 5.51	94.2 $\pm$ 4.50	0.5403
PREMEDICATION	99.2 $\pm$ 4.33	97.53 $\pm$ 5.66	0.2014
INDUCTION	94.6 $\pm$ 4.66	96.5 $\pm$ 5.73	0.1642
INTUBATION	95.2 $\pm$ 4.71	96.3 $\pm$ 4.11	0.3391
1 MINS	95.5 $\pm$ 3.55	102.5 $\pm$ 3.96	<0.0001
3 MINS	93.8 $\pm$ 4.43	100.8 $\pm$ 5.54	<0.0001
5 MINS	94.6 $\pm$ 4.31	99.8 $\pm$ 4.21	<0.0001
15 MINS	94.8 $\pm$ 3.55	98.4 $\pm$ 3.63	0.0003
30 MINS	94.8 $\pm$ 4.28	96.1 $\pm$ 4.30	0.2453

Table 1 shows the changes in heart rate in the two study groups and comparison among them to baseline value. In etomidate group heart rate did not significantly change during induction and during intubation and after 30 mins postintubation compared to pre-operative value. But in thiopentone group ,during intubation and after intubation till 15 mins heart rate increased significantly compared to pre-operative value.

Table 2 shows change in Systolic blood pressure in the two study groups. In etomidate group , induction and after 30 mins postintubation systolic blood pressure did not change significantly as compared to baseline. In thiopentone group, during and after 5mins postintubation there is significant increase in systolic blood pressure compared to baseline. Table 3 shows the changes in diastolic blood pressure in the two study groups and comparison of them with preinduction (baseline) value. In etomidate group, there was no significant changes in diastolic blood pressure during induction or after 30mins postintubation. But in thiopentone group there is significant increase in diastolic blood pressure after intubation till 15 mins. Table 5 shows changes in mean blood pressure in the two study groups. In etomidate group, there is no significant changes in mean blood pressure after induction or post intubation. But in thiopentone group there is significant increase in mean blood pressure following intubation.

## DISCUSSION

Early mobilization and shorter hospital stay are the main advantages of laparoscopic surgery. Being minimal invasive surgery, laparoscopic surgeries are very popular now a days. The main drawback is intra-operative hemodynamic instability due to pneumoperitoneum. Now a days many laparoscopic surgeries are conducted as a daycare surgery. Therefore, anesthesiologist must choose anesthetic agent which provide hemodynamic stability and rapid recovery. We compared 60 patients of ASA physical status I-II -III undergoing either laparoscopic surgery under general anesthesia for hemodynamic stability of etomidate and thiopentone. In the two groups heart rate and blood pressure were taken in stepwise manner, at baseline, premedication , induction, intubation and after intubation at 1,3,5,15 and 30 mins. The recorded values were tabulated for age, weight, sex, heart rate, systolic blood pressure, diastolic blood pressure and mean blood pressure. The result of the study was compared with the observation of other workers in the field of work taking into account for the differences as far as possible.

**The discussion of the study is as follows:**

### Haemodynamic Parameters

**Heart Rate:** As shown in table 1, baseline values of mean heart rate were comparable between both the groups with no statistically significant difference ( $P>0.05$ ). Changes in the heart rate during induction were also not statistically significant between both the Groups ( $P>0.05$ ). During intubation and after intubation there was significant increase in heart rate values in the group thiopentone compared to the group etomidate. During intubation and after intubation 1,3, 5, 15 and 30 mins heart rate values were  $82.56\pm 5.72, 79.5\pm 6.38, 80.76\pm 5.96, 80.73\pm 4.84, 78.73\pm 5.67, 80.56\pm 5.63$  in the group etomidate respectively. In group T (thiopentone) during intubation and after intubation 1,3,5,15,30 minutes mean heart rate values were  $90.63\pm 6.80, 94.66\pm 5.04, 93.1\pm 4.12, 91.66\pm 5.95, 87.93\pm 5.57, 82.26\pm 6.2$  respectively. Comparing the heart rate of two groups P value was calculated and was found to be significant as P value is  $<0.05$ . P value during and after intubation was found to be  $<0.0001$  till 15 minutes and at 30 minutes after was 0.2708. This showed that heart rate remained stable throughout 30 mins post intubation in etomidate group while a significant increase was seen in thiopentone group following intubation. Mousami das, Basant ku in their study showed post induction and after intubation, in etomidate group heart rate did not significantly change compared to pre induction. In thiopentone group, post induction and after intubation heart rate increased significantly compared to pre-induction.<sup>(35)</sup>

**Systolic Blood Pressure:** As shown in table 2, baseline values of systolic blood pressure were comparable between both the groups with no statistically significant difference ( $P>0.05$ ). Changes in the systolic blood pressure after giving study drug i.e during induction were also not statistically significant between both the groups ( $P>0.05$ ). Systolic blood pressure during induction in group E was ( $120.2 \pm 6.61$ ) and in group T was ( $124.5 \pm 9.92$ ) which showed a mild increase in thiopentone group but was insignificant as P value is 0.0529. During intubation and after intubation till 5 mins there was significant increase in systolic blood pressure values in the group thiopentone compared to the group etomidate. During intubation and after intubation 1,3, 5,15 and 30 mins systolic blood pressure values were  $120.9\pm 6.72, 121.3\pm 7.57, 120.1\pm 7.58, 121.8\pm 6.9, 122.1\pm 6.84, 121.6\pm 8.47$  in the group etomidate. In group T (thiopentone) during intubation and after intubation 1,3,5,15,30 minutes systolic blood pressure values were  $125.1\pm 7.79, 128.5\pm 9.41, 127.03\pm 7.60, 126.43\pm 8.59, 124.8\pm 8.5$  and  $123.2\pm 8.84$  respectively. Comparing the systolic blood pressure during and after intubation of two groups P value was calculated and was found to be significant as P value is  $<0.05$ . P value during and after intubation was found to be 0.0292, 0.0018, 0.0008, 0.0308 till 5 minutes and at 15 and 30 minutes after was 0.1805 and 0.4770 which suggested that while using etomidate systolic blood pressure remained stable throughout and with thiopentone there was significant increase till 5 minutes after intubation and settled to baseline within 15 minutes. Haris et al detected in their study that there were significant increases in SAP values in the group receiving thiopentone after intubation.<sup>(3)</sup> Mehmet Levent ugyur et al in their study showed remarkable increase in SAP

in 1<sup>st</sup> and 5<sup>th</sup> minute postinduction in the group receiving thiopentone.<sup>(37)</sup>

**DIASTOLIC BLOOD PRESSURE:** As shown in table 3, baseline values of diastolic blood pressure were comparable between both the groups with no statistically significant difference ( $P>0.05$ ). Changes in the diastolic blood pressure after giving study drug i.e during induction were also not statistically significant between both the groups ( $P>0.05$ ). Diastolic blood pressure during induction in group E was ( $81.7\pm 6.75$ ) and in group T was ( $82.6\pm 6.29$ ) with insignificant P value 0.6701. After intubation from 1 min till 15 mins there was significant increase in diastolic blood pressure values in the group thiopentone compared to the group etomidate.

After intubation 1,3, 5,15 and 30 mins diastolic blood pressure were  $82.6\pm 5.47, 80.6\pm 6.35, 81.1\pm 5.85, 81.3\pm 5.16, 81.7\pm 6.86$  in the group etomidate respectively. In group T (thiopentone) after intubation 1,3,5,15,30 minutes mean diastolic blood pressure values were  $89.6\pm 5.61, 87.7\pm 7.96, 86.5\pm 5.13, 85.1\pm 3.58, 84.1\pm 5.95$  respectively. Comparing the diastolic blood pressure after intubation of two groups P value was calculated and was found to be  $<0.0001, <0.0001, 0.0003, 0.0016, 0.153$  which suggested that while using etomidate diastolic blood pressure remained stable throughout till 30 mins postintubation and with thiopentone there was significant increase till 15 minutes after intubation and settled to baseline within 30 minutes. Lindgren L et al. studied hemodynamic and catecholamine changes after induction of anesthesia with either thiopentone or propofol. With thiopentone, heart rate was greater than propofol before intubation. During induction systolic and diastolic blood pressure decreased more with propofol than with thiopentone.<sup>(25)</sup>

### MEAN BLOOD PRESSURE

As shown in table 4, baseline values of mean blood pressure were comparable between both the groups with no statistically significant difference ( $P>0.05$ ). Changes in the mean blood pressure after giving study drug i.e during induction was also not statistically significant between both the groups ( $P>0.05$ ). Mean blood pressure during induction in group E was ( $94.6\pm 4.66$ ) and in group T was ( $96.5\pm 5.73$ ) with insignificant P value 0.1642. During intubation there was no significant changes in both the groups. After intubation there was significant increase in mean blood pressure values in the group thiopentone compared to the group etomidate. After intubation 1,3, 5,15 and 30 mins were  $95.5\pm 3.55, 93.8\pm 4.43, 94.6\pm 4.31, 94.8\pm 3.55, 94.8\pm 4.28$  in the group etomidate respectively. In group T (thiopentone) after intubation 1,3,5,15,30 minutes mean blood pressure values were  $102.5\pm 3.96, 100.8\pm 5.54, 99.8\pm 4.21, 98.4\pm 3.63, 96.1\pm 4.3$  respectively. Comparing the mean blood pressure after intubation of two groups P value was calculated and was found to be significant as P value is  $<0.05$ . P value after intubation was found to be  $<0.001$  for 1,3,5 minutes and 0.0003 at 15 minutes till which suggested that while using etomidate mean blood pressure remained stable throughout till 30 minutes postintubation and with thiopentone there was significant increase till 15 minutes after intubation and settled to baseline within 30 minutes. Brossy MJ et al. In their study found that heart rate increased significantly above baseline after induction and intubation in both groups but there were no differences between groups. Arterial pressure

increased significantly at 1 min after intubation in both groups and at 2 min in the thiopentone group only.<sup>(36)</sup>

## Summary

This study was designed to study and compare the effectiveness of intravenous Etomidate and Thiopentone in attenuating hemodynamic response to direct laryngoscopy and endotracheal intubation in patients undergoing laparoscopic surgical procedures under General Anaesthesia.

## Conclusion

- The demographic profile of the patients in terms of age, body weight and sex were comparable in both the groups.
- There was no significant change in heart rate, systolic blood pressure, diastolic blood pressure and mean blood pressure in post induction and after intubation in Etomidate group.
- In Thiopentone group heart rate, systolic blood pressure, diastolic blood pressure, mean blood pressure increased significantly post intubation compared to preoperative values.
- From the above observations and discussion, it is concluded that Etomidate attains most properties of an ideal induction agent in comparison to Thiopentone as far as the haemodynamic responses during laryngoscopy and intubation is concerned in laparoscopic surgeries.

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