

## A COMPARISON OF PROPOFOL WITH SEVOFLURANE AND PROPOFOL ALONE FOR INDUCTION AND INTUBATION

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**ABSTRACT: STUDY OBJECTIVE:** We aimed to study propofol with sevoflurane and propofol alone in evaluating intubating conditions, hemodynamic response during induction and intubation and induction side effects in adult patients undergoing various elective surgical procedures without muscle relaxants.

**DESIGN:** Prospective randomized study.

**SETTING:** Operation theatre of a teaching institute.

**PATIENTS:** The study population consists of 60 ASA I & II, non-obese, adult patients aged between 20-40yrs coming for elective surgical procedures under General Anaesthesia and had Mallampatti class I airway anatomy, 30 of these patients receive propofol alone-“GROUP A” and 30 of who receive propofol with sevoflurane-“GROUP B”.

**MEASUREMENTS AND RESULTS:** The heart rate, systolic blood pressure, diastolic blood pressure and mean arterial pressure before and after induction and post-intubation at 1, 3 and 5 minutes were recorded. Time to induction in seconds (Start of anaesthetic until loss of eye lash reflex), induction side effects like breath holding, cough, excitatory movements, laryngospasm and others (Bradycardia, hypoxia, hyperthermia, hypothermia and injection site pain) were noted.

Intubating conditions were better in Group-B than in Group-A, Group-B patients had significantly had more clinically acceptable intubating conditions than Group-A. There was no significant difference in heart rate after induction and intubation between the two groups, except 3min after intubation in Group-A there is significantly low heart rate. There was significant in reduction in systolic blood pressure after induction and intubation in Group-A, however there was no significant difference in diastolic blood pressure and mean arterial pressure between two groups. Induction time is significantly less in Group-A patients when compared to Group-B patients and there was no significant difference in induction side effects between two groups.

**CONCLUSION:** Combination of inhalational 4% sevoflurane with IV propofol 1.5mg/kg is superior to IV propofol 3mg/kg with respect to quality of intubation and less significance with respect to hemodynamic response during induction and intubation in adult patients undergoing various elective surgical procedures without muscle relaxants.

**KEYWORDS:** Induction, Intubation, Sevoflurane, Propofol, Hemodynamics.

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**INTRODUCTION:** Endotracheal intubation is the most important and crucial step during administration of general anaesthesia.<sup>1</sup> The ease with which endotracheal intubation is achieved depends on technical proficiency, depth of anaesthesia and degree of muscle relaxation. Intubation in anaesthesia using short-acting hypnotic drug is frequently facilitated by the simultaneous administration of a depolarizing muscle relaxant such as succinylcholine. However, succinylcholine administration may be associated, at times, with side effects such as postoperative myalgia, malignant hyperthermia, masseter spasm, histamine release, anaphylaxis, hyperkalemia, cardiac arrhythmias and increase in intracranial or intraocular pressure. These side effects have spurred research into the development of a non-depolarizing drug with short onset of action.

Even the use of non-depolarizing relaxants may be associated with undesirable effects such as prolonged neuromuscular blockage, the need to reverse neuromuscular blockade, or the inability to reverse the paralysis quickly if airway management via mask or tracheal intubation is not possible. For these reasons, a method of providing good intubating condition rapidly without muscle relaxants has been sought.<sup>2-13</sup> Propofol in combination with short-acting opioids such as fentanyl, alfentanil and remifentanil may provide adequate conditions for laryngoscopy and tracheal intubation without using muscle relaxants.<sup>13,14</sup>

Such a technique is of value in particular situations in which muscle relaxants have to be avoided (Myopathies, known allergic reactions to muscle relaxants) or in cases where succinylcholine is contraindicated (Hyperkalemia, burns, plasma cholinesterase deficiency, penetrating eye injury). Propofol is a short-acting intravenous anaesthetic with high lipid solubility and short elimination half-life.<sup>15</sup> However, propofol has been associated with several adverse effects, including hypotension, apnea, pain on injection, and excitatory patient movements.<sup>16</sup> Pain on injection can be avoided if propofol is administered after inhalation induction of anaesthesia. Potent inhalation agents can be used as an alternative to facilitate tracheal intubation.

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Sevoflurane with its relatively pleasant smell, low airway irritability and low blood-gas solubility allowing smooth and more rapid induction and recovery, Sevoflurane as compared with propofol, has the advantage of providing better hemodynamic stability and a smoother transition to the maintenance phase without a period of apnea.<sup>17</sup> Combination of lesser percentage of halothane with propofol has been studied and concluded that combination of inhalational agent and propofol is ideal for intubation in children.<sup>18</sup> Sevoflurane 8% can be used as an alternative to facilitate tracheal intubation.<sup>19</sup> but it is not cost effective. Combination of Sevoflurane 8% and propofol 1.5mg/kg has been tried for Laryngeal Mask Airway insertion.<sup>17</sup>

Induction of anaesthesia with a combination of lesser dose of propofol and lesser percentage of sevoflurane with opioid pre-medication may optimize the inserting conditions of endotracheal tube and decrease the side effects that may follow with propofol alone.<sup>17</sup> Hence an attempt was made with a combination of lesser percentage of Sevoflurane with reduced dosage of Propofol for intubation with endotracheal tube to evaluate intubation conditions, hemodynamic response during induction and intubation and induction side effects without muscle relaxants in adult patients of age group 20-40yrs undergoing various elective surgical procedures.

**MATERIALS AND METHODS:** This study was undertaken after approval by our institution Research and Ethics committee and obtaining patient's written informed consent, patients were randomized into two groups of 30 each, i.e. Group A and Group B. This is a prospective randomized study. The study population consists of 60 ASA I & II, non-obese, adult patients aged between 20-40yrs coming for elective surgical procedures under General Anaesthesia and had Mallampatti class I airway anatomy. Patients were excluded who were unwilling, had history or evidence of difficult airway, malignant hyperthermia, allergy to volatile anaesthetics or propofol, patients who on MAO-inhibitors and patients with body mass index more than 1.5 times normal.

A thorough pre-anaesthetic evaluation was conducted on the day before surgery. Detailed history and cardio-respiratory examination was carried out in all patients. All relevant investigations were done. Nil per oral status for a minimum of 6 hrs was advised. On the day of surgery, after arrival of patient to the operation theatre pulse-oxymeter, ECG, and non-invasive blood pressure monitors were connected. The baseline heart rate, systolic blood pressure, diastolic blood pressure and mean arterial pressure were recorded.

After doing a thorough cockpit drill of continuous flow anaesthesia machine and availability of emergency drugs with ETCO<sub>2</sub> monitor, an intravenous line with Ringer's Lactate was secured using either 18G or 20G intravenous cannula. All patients were pre-medicated with IV fentanyl 2µg/kg, IV midazolam 1mg & IV Glycopyrrolate 0.2mg 5min before induction. All patients were pre-oxygenated with 100% O<sub>2</sub> for 3 min. Anaesthesia was then induced in Group-A patients by 67% N<sub>2</sub>O in O<sub>2</sub> and IV propofol 3mg/kg injected over 30s. Group-B patients were induced by mask with sevoflurane starting at 0.5% and incrementally increased to

4% inhaled concentration with 67% nitrous oxide in oxygen at a total gas flow of 8 liters/min and IV propofol 1.5mg/kg injected over 15s and tracheal intubation was attempted at 240s after the start of induction in both groups.

Lignocaine 0.2mg/kg added to propofol to prevent pain on injection. The heart rate, systolic blood pressure, diastolic blood pressure and mean arterial pressure before and after induction and post-intubation at 1, 3 and 5 minutes were recorded. Time to induction in seconds (Start of anaesthetic until loss of eye lash reflex), induction side effects like breath holding, cough, excitatory movements, laryngospasm and others (Bradycardia, hypoxia, hyperthermia, hypothermia and injection site pain) were noted.<sup>19</sup>

Tracheal intubation was performed using appropriately sized endo-tracheal tube. Intubating conditions were assessed by anaesthesiologist who performed intubation using Copenhagen Consensus Conference (CCC) score.<sup>19</sup> which graded the quality of tracheal intubation according to ease of laryngoscopy, position of the vocal cords, cough and movement of the limbs. Supplementation of endotracheal intubation with IV succinylcholine noted.

**RESULTS:** Statistical analysis of age, sex and weight distribution was done by using student's unpaired-t test. A p-value of less than 0.05 was regarded as significant. Both groups were found to be statistically similar with respect to age, sex and weight distribution. ASA grade is statistically similar between two groups with P=0.688.

Time to Induction (Sec)	Group A (n=30)	Group B (n=30)
1-100	30 (100.0%)	0
101-200	0	29 (96.7%)
>200	0	1 (3.3%)
<b>Total</b>	<b>30 (100.0%)</b>	<b>30(100.0%)</b>
<b>Mean ± SD</b>	<b>39.80±8.10</b>	<b>156.07±21.58</b>

**Table 1: Time to Induction (Seconds)**

Induction time is significantly less in Group A patients (39.80±8.10) when compared with Group B patients (156.07±21.58), (p<0.001).

Induction Side Effects	Group A (n=30)	Group B (n=30)	P value
Breath holding	3 (10.0%)	0	0.237
Cough	6(20.0%)	2(6.7%)	0.254
Excitatory movements	3(10.0%)	1(3.3%)	0.612
Laryngospasm	0	0	-
Others	0	0	-

**Table 2: Induction Side Effects**

Both groups were found to be statistically similar with respect to breath holding, cough, excitatory movements, laryngospasm and other induction side-effects.

Number of Attempts	Group A (n=30)	Group B (n=30)
1	23 (76.7%)	29(96.7%)
2	5(16.7%)	1(3.3%)
3	2(6.6%)	0

**Table 3: Number of Attempts**

23.3% patients in group A required 2 or 3 attempts for intubation when compared with 3.3% in group B, which is highly significant ( $p < 0.001$ ).

Tracheal intubation supplemented with succinylcholine	Group A (n=30)	Group B (n=30)
No	26(86.7%)	30(100.0%)
Yes	4(13.3%)	0

**Table 4: Tracheal Intubation Supplemented with Succinylcholine**

None of the patients in Group B required succinylcholine supplementation to achieve intubation, when compared with 13.3% in Group A, which is not significant ( $p = 0.112$ ).

Heart Rate (bpm)	Group A (n=30)	Group B (n=30)	P value
Pre-Induction	90.97±9.86	89.13±13.63	t=0.597;p=0.553
Post-Induction	81.97±8.66	86.73±13.34	t=1.642;p=0.106
1 min after intubation	87.9±8.47	91.43±13.42	t=1.220;p=0.227
3 min after intubation	87.33±7.57	93.67±13.26	t=2.272;p=0.027*
5 min after intubation	87.67±8.1	89.23±13.33	t=0.550;p=0.584

**Table 5: Comparison of Heart Rate (bpm) between Two Groups**

There was no significant difference in heart rate after induction and post-intubation between the two groups except 3min after intubation which was significant ( $p = 0.027$ ).

MAP (mm Hg)	Group A (n=30)	Group B (n=30)	P value
Pre-Induction	96.1±8.05	93.17±8.12	t=1.405;p=0.165
Post-Induction	85.33±8.18	90.07±7.66	t=2.314;p=0.024*
1 min after intubation	91.77±7.79	94.50±7.74	t=1.363;p=0.178
3 min after intubation	94.00±8.38	95.10±7.65	t=0.531;p=0.597
5 min after intubation	92.20±7.96	93.63±7.77	t=0.705;p=0.483

**Table 6: Comparison of MAP (mm Hg) between Two Groups**

There was no significant difference in mean arterial pressure between the two groups following intubation, but there was a significant difference in mean arterial pressure following induction ( $p = 0.024$ ).

**DISCUSSION:** Laryngoscopy and tracheal intubation are essential skills associated with practice of anaesthesia. The drugs should be combined in such a way that it produces unconsciousness, analgesia and muscle relaxation without compromising hemodynamic stability, at the same time providing best intubating conditions.<sup>4</sup> usually a combination of hypnotic agent, opioid and a neuromuscular blocking agent is used. Over past few years, several factors have led researchers to ignore neuromuscular blocking agents for tracheal intubation. The driving force were introduction of

propofol, short acting opioids and sevoflurane in clinical practice. Propofol not only suppresses upper airway reflexes and pressor response to laryngoscopy and tracheal intubation but also provides faster recovery of consciousness, possess anti-emetic action and reduces incidence of airway complications.<sup>20</sup>

Although, succinylcholine is the gold standard to provide adequate relaxation because of its rapid onset within 30-60s and quick metabolism, routine use of this drug has been questioned following several reports of cardiac arrest in young children. In addition it has many other potential problems: myalgia, cardiac arrhythmias, elevated intraocular and intracranial pressure, hyperkalemia, malignant hyperthermia and prolonged apnea.<sup>20</sup> Non-depolarizing neuromuscular agents are alternative but are slower in onset and have a longer duration of action.

Sevoflurane a new inhalational agent with low blood-gas solubility and a relatively pleasant odour produces rapid induction and recovery. It causes less myocardial depression and cardiac arrhythmias than halothane.<sup>21</sup> In our study, we used fentanyl 2µg/kg intravenously, 5 min before induction, because in addition to analgesia, it also blunts pressor response against laryngoscopy and intubation. Fentanyl also has anti tussive action. Katochet al.<sup>22</sup> suggested that fentanyl blocks afferent nerve impulses arising from stimulation of the pharynx, larynx and lungs during intubation.

The peak effect of propofol from the time of administration of drug was around 90-100s, Mc Keating et al.<sup>3</sup> study showed that it is possible to perform laryngoscopy safely and smoothly at 120s after induction with propofol. Therefore we took 240s as a fixed time interval from the start of induction to intubation in Group A patients (IV propofol 3mg/kg). The use of fixed time interval tests an easily reproducible technique, independent of subjective assessments of depth of anaesthesia.

Swadia VN et al.<sup>23</sup> and Bithal PK et al.<sup>24</sup> had found significantly greater time for tracheal intubation with sevoflurane i.e. (242.2±52.67s) and (325.93±44.02s) respectively. This difference was not only because of different clinical end points but also a different induction technique in which sevoflurane concentration was increased incrementally and ventilation was not assisted manually. In a study by Erhan E et al.<sup>5</sup> clinically acceptable intubating conditions were found in 93.3%, 66.7% and 40% in patients receiving propofol, thiopental or etomidate respectively. Patients receiving propofol found to have less severe coughing after intubation when compared to thiopental or etomidate.

In Thwaiteset al.<sup>25</sup> study, all children could successfully be intubated with 8% sevoflurane in nitrous oxide and oxygen at 150s. 91% children had excellent intubating conditions and 9% had good intubating conditions. They demonstrated that 8% sevoflurane with nitrous oxide in oxygen can provide acceptable intubating conditions at 150s.

In our study, tracheal intubation was accomplished in 100% of patients in Group B, 93.3% of those patients had acceptable intubating conditions when compared with 73.3% in Group A, which is highly significant ( $\chi^2 = 4.320$ ;  $p < 0.001$ ). 86.7% of patients had no cough in Group B, compared with 56.7% in group A.

Coughing was significantly associated more with Group A ( $p=0.037$ ). Limb movements were significantly more in Group A compared to Group B. None of the patients in Group B required succinylcholine supplementation to achieve intubation. 96.7% of patients were intubated at first attempt in Group B when compared with 76.7% in Group A. Number of attempts were significantly less in Group B ( $p<0.001$ ). In Group A 13.3% of patients required succinylcholine supplementation to achieve intubation because of vocal cords movement, coughing and excessive limb movements.

Only 76.7% of patients intubated at first attempt and remaining 23.3% required multiple attempts. During induction, 10% of patients in Group A had breath holding, 20% had cough and 10% had excitatory movements, which is not significant. Induction time in Group B patients were  $156.07\pm 21.58$ s, when compared with Group A ( $39.80\pm 8.10$ ). Induction time were more in Group B patients ( $t=27.629$ ;  $p<0.001$ ). In Swadia et al.<sup>23</sup> study anaesthesia was induced with 60% nitrous oxide in oxygen and incremental increase in concentration of sevoflurane from 1-7%.

Time interval from application of facemask to intubation was  $242\pm 52.67$ s. 80% of children had excellent intubating conditions. 16% had tachycardia, 8% had bradycardia and 80% had hypotension. Complications like laryngospasm, bronchospasm were not observed. In present study there was reduction in heart rate, systolic blood pressure, diastolic blood pressure and mean arterial pressure in Group-A patients after induction and intubation when compared with pre-induction values.

However, there was no significant difference among these parameters when compared with pre-induction values in Group B patients. Thus propofol decreased both heart rate and blood pressure, which indicates there was decrease in cardiac output. Similar results were found in other studies, Srivastava U et al.<sup>13</sup> found significant decrease in HR and arterial pressure from baseline in children given propofol and fentanyl. Steyn et al.<sup>26</sup> observed a no change in HR but found a significant fall in MAP after induction and following intubation with a dose combination of propofol 3mg/kg and alfentanil 15 $\mu$ g/kg in children.

In Swadia et al.<sup>23</sup> study sevoflurane group 16% patients, developed tachycardia, 8% had bradycardia and 80% had hypotension. In Bithal PK et al.<sup>24</sup> study HR was significantly high in the sevoflurane group, during post-induction and immediate post-intubation and 1min post-intubation. MAP also increased but slightly from baseline. In our study there was no significant difference in heart rate after induction and intubation between the two groups, except 3min after intubation, where heart rate is significantly low in Group A ( $87.33\pm 7.57$ ) when compared with Group B ( $93.67\pm 13.26$ ), ( $p=0.027$ ).

There was significant reduction in systolic blood pressure after induction and intubation in Group A patients when compared with Group B patients. However, there was no significant difference in diastolic blood pressure and mean arterial pressure between two groups, except mean arterial pressure being low in Group A following induction ( $p=0.024$ ).

**CONCLUSION:** We concluded that combination of inhalational 4% sevoflurane with IV propofol 1.5mg/kg is superior to IV

propofol 3mg/kg with respect to quality of intubation and less significance with respect to hemodynamic response during induction and intubation in adult patients undergoing various elective surgical procedures without muscle relaxants and also this combination is cost effective. This combination can also be attempted for anticipated difficult intubation.

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