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A COMPARISION BETWEEN SUBLINGUAL NITROGLYCERIN SPRAY AND I.V LIGNOCAINE IN ATTENUATION OF HEMODYNAMIC RESPONSES TO INTUBATION AND DIRECT LARYNGOSOPY

Soujan Chowta¹, Yashvanth R², Gauri S Kene³ Suchet Sharath⁴, Rachan Amin⁵, Akhil Rao U K⁶

 ^{1,3,6}Assistant Professor, Department of Anaesthesiology and Critical Care, Srinivas Medical College and Hospital, Mukka, Mangalore, Karnataka, India.
 ^{2,4,5}Senior Resident, Department of Anaesthesiology and Critical Care, Srinivas Medical College and Hospital, Mukka, Mangalore, Karnataka, India.

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Corresponding Author: Dr Yashvanth R, Senior Resident, Department of Anaesthesiology and Critical Care, Srinivas Medical College and Hospital, Mukka, Karnataka, India. **Email:** <u>yashgowda226@gmail.com</u>

Abstract

Background: Laryngoscopy and tracheal intubation is associated with a reflex sympathetic pressor response which results in increased heart rate and blood pressures. This may prove detrimental in high-risk patients. Objective of this study is to compare the effects of intravenous lignocaine and oral nitroglycerin spray in attenuation of this response. Methods: 90 ASA I and II status normotensive patients scheduled for elective surgical procedures were selected randomly and divided into three groups of 30 each. All patients received premedication, the first group was control group, in this group no drug was administered for attenuating sympathetic response to laryngoscopy and intubation, the second group received 2mg/kg lignocaine i.v. bolus and the third group received 0.4mg/1 spray of sublingual nitroglycerin, 3 minutes before laryngoscopy and intubation. HR, systolic, diastolic blood pressure were recorded noninvasively before induction, postinduction, 1,3,5, 7 and 10 minutes from the onset of laryngoscopy.' ANOVA' test was used for statistical analysis. Results: Post intubation incidence of tachycardia (HR>100/min) was significantly higher in control and nitroglycerin group (p<0.05) than in lignocaine group. Rise in SBP and DBP were also statistically significant in control and nitroglycerin group than in lignocaine group (p<0.05) Interpretation and Conclusion: Attenuation of pressor response is seen both with lignocaine and nitroglycerin. Of the two drugs lignocaine 2mg/kg i.v. bolus provides a consistent, reliable and effective attenuation as compared to sublingual nitroglycerin spray 0.4mg.

Key words: Attenuation of pressor response, laryngoscopy, intubation, lignocaine, nitroglycerin.

Introduction

The induction of anaesthesia, laryngoscopy, tracheal intubation and surgical stimulation often evoke cardiovascular responses characterised by alteration in systemic arterial blood pressure, heart rate and cardiac rhythm. The response following laryngoscopy and intubation peaks at 1 to 2 minutes and return to baseline within 5 to 10 minutes. Though the sympathoadrenal responses are probably of little consequence in healthy patients, it is

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hazardous to those patients with hypertension, coronary heart disease, cerebrovascular disease, intracranial pathology and hyperactive airways. In such cases, reflex circulatory responses such as an increase in heart rate, systemic arterial pressure and disturbances in cardiac rhythm need to be suppressed.

Prof. King *et al.* ^{[1],} documented myocardial ischemic changes due to reflex sympathoadrenal responses immediately following laryngoscopy and intubation with a mean increase in systolic pressure of 40mmHg even in normotensive individuals. Pyrs Roberts *et al.* ^{[2],} showed exaggerated form of this response in hypertensive patients.

Various systemic as well as topical agents have been used to reduce these untoward hemodynamic responses during laryngoscopy. When compared to systemic agents, administration of local anaesthetic solutions is likely to be of limited value. The commonest strategies adopted are narcotics, vasodilators, β -blockers, calcium channel blockers, lignocaine, clonidine and other sympatholytics. In our study, we have compared intravenous lignocaine, oral nitroglycerin spray, and placebo in suppressing stress responses to laryngoscopy and intubation.

Since intravenous lignocaine, oral nitroglycerin spray have been known to blunt sympathetic responses to intubation, their efficacy has been compared with control (placebo) in the Department of Anaesthesiology, JJMMC, DAVANGERE.

Aim of the Study

For the safe conduct of anaesthesia, the hemodynamic responses to laryngoscopy and intubation should be abolished or at least attenuated to balance the myocardial oxygen supply and demand. This study was done to compare the efficacy of intravenous lignocaine, oral nitroglycerin spray and placebo in attenuating the hemodynamic stress responses to laryngoscopy and intubation.

Materials And Methods

90 patients of ASA physical status 1 or 2 undergoing elective surgical procedure under general anaesthesia with endotracheal intubation were included in this study. Patients belonging to age group 20 and 50 years of both the sexes were included. It is a prospective single blinded randomized controlled study. The study was approved by our institution ethical committee and after obtaining written, informed consent from the patient, this study was conducted. The study was done during the period from May 2022 to March 2024 in the Department of Anaesthesiology, JJMMC.

Inclusion criteria: 1) ASA 1 or 2 2) Patients with airway with modified mallampatti grade class 1 or 2 3) Age group 20 to 50 years of both sexes. 4) Patients who are willing and able to give informed written consent

Exclusion criteria: 1) Patients with full stomach 2) Patients posted for emergency surgery 3) Patients with difficult airway 4) Hypertension, diabetes, ischemic heart disease and pregnancy 5) Patients with contraindications to study drugs. 6) Patient refusal

Anaesthetic protocol: Pre-operative visit was done to allay anxiety and good rapport was established with the patients. All the patients were given pre-operative night sedation with tab.alprazolam 0.5mg orally. Interventions: Induction of anaesthesia was standardised for all patients. Monitors used were NIBP, ECG, EtCO2 and pulse oximetry. Method: 90 patients of both the sexes of ASA 1 or 2 undergoing surgical procedure were randomly allocated into three groups. Group A – was control group, in this group no drug was administered for attenuating sympathetic response to laryngoscopy and intubation Group B – receiving oral NTG spray (0.4mg/spray) 3 minutes before laryngoscopy and intubation Group C – receiving

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i.v. lignocaine (2mg/kg) 3 minutes before laryngoscopy and intubation. Their pulse rate, blood pressure and sp02 were recorded. They were premedicated with injection Glycopyrrolate 0.01 mg/kg, injection Midazolam 0.05mg/kg and injection Pentazocine 0.5mg/kg Preoxygenation done with 100% oxygen for 3 minutes.

Administration of study drug: The study drug was given 3minutes before laryngoscopy and intubation. Then vital signs were recorded. Two minutes later, patient was induced with injection propofol 2mg/kg body weight intravenously. The vital signs were recorded. Then, injection succinylcholine 2mg/kg body weight was given. Intubation was performed by the same person for all the cases with appropriate sized endotracheal tubes orotracheally. Anaesthesia was maintained with controlled ventilation with N2O/O2 mixture 3:2, Sevoflurane 1 MAC and injection vecuronium 0.05mg/kg and IPPV. No surgical stimulation was permitted for 10 minutes after intubation. At the end of surgery reversal was done with Inj. Neostigmine 0.05mg/kg and Inj. Glycopyrrolate 0.01mg/kg i.v. Patients were monitored throughout the period from entering into operation theatre till recovery and in the immediate post-operative period by means of automated NIBP, pulse oximetry and ECG in a multichannel monitor. ECG was monitored with particular importance to any alteration in rhythm. All patients were extubated and were shifted to post anaesthetic care unit for a follow up for 24 hours.

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Variables	Groups	Ν	Mean	Std. Dev	F value	p value*
AGE	Control	30	28.10	8.880	1.463	.237
	IV Lignocaine	30	32.07	9.028		
	Oral NTG spray	30	31.43	10.909		
	Total	90	30.53	9.700		

RESULTS

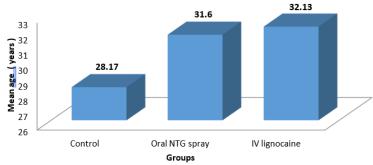


TABLE 1: AGE DISTRUBUTION

Figure 1

Gro ups	Conti	rol		IV Li	IV Lignocaine				NTG	p val ue*	Cont rol and IV	IV an d O ra l	Cont rol and Ora l
MA P	Me an	SD	M D	M ea n	SD	M D	Me an	SD	M D		p valu e [#]	p val ue [#]	p valu e [#]

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Basa	82.	5.0		79.	4.6		81.	4.8	2.5	0.0	.002	.08	.000
1	03	55		67	93		50	55		01		6	
Post	88.	5.7	6.8	84.	4.2	4.7	100	6.4	21.	0.0	0.00	0.0	0.00
ind	87	22	4	37	79	00	.37	78	37	01	1	01	1
1	117	6.8	35.	89.	4.2	9.6	105	4.4	16.	0.0	0.00	0.0	0.00
min	.57	82	533	30	68	33	.13	78	27	01	1	01	1
3	119	8.0	39.	92.	3.0	13.	92.	2.8	8.5	0.0	0.00	1	0.00
min	.60	50	933	93	51	933	87	97	0	01	1		1
5	108	7.9	29.	92.	1.8	3.2	84.	2.9	3.0	0.0	0.00	1	0.00
min	.50	86	500	07	18	00	53	21	3	01	1		1
7	96.	8.1	7.3	85.	1.4	1.0	80.	3.1	-	0.0	0.00	0.5	0.00
min	17	50	00	37	26	00	30	09	37.	01	1	44	1
									27				
10	88.	6.1	4.1	78.	1.8	-	81.	4.8	2.5	0.0	.002	.08	.000
min	53	57	67	87	33	2.6	50	55		01		6	
						33							

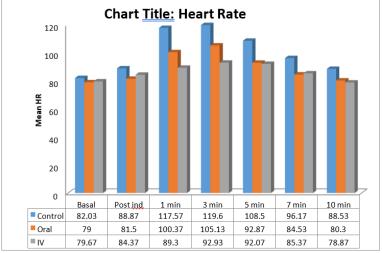


Figure 2

TABLE 3: SYSTOLIC BLOOD PRESSURE

Gro ups	Cont	rol		IV Li	gnocai	ne	Oral	NTG s	pray	p val ue*	Con trol and IV	IV an d O ra l	Con trol and Ora l
MA	Me	SD	M	Me	SD	M	Me	SD	M		p ,	p	p ,
Р	an		D	an		D	an		D		valu e [#]	val ue [#]	valu e [#]
Bas	130	17.		131	12.		131	16.		0.9	1	1	1
al	.33	095		.17	614		.73	484		4			
Post	127	16.	-	124	11.	-	129	16.	-	0.4	1	0.6	1
ind	.27	58	3.0	.87	325	6.3	.67	668	2.0	7		62	
			6						6				
1	140	8.5	10.	132	17.	1.8	139	15.	7.3	0.0	0.12	0.3	1

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min	.67	35	34	.97	713		.1	473	7	98	6	11	
3	143	15.	12.	125	17.	-	130	18.	-	0	0	0.6	0.02
min	.17	024	84	.37	893	5.8	.9	252	0.8			42	
									3				
5	149	14.	19.	118	17.	-	128	18.	-	0	0	0.0	0
min	.6	459	27	.03	664	13.	.1	153	3.6			69	
						14			3				
7	151	13.	21.	115	17.	-	129	17.	-	0	0	0.0	0
min	.8	197	47	.4	316	15.	.6	745	2.1			03	
						77			3				
10	131	14.	1.4	129	10.	-	131	17.	-	0.8	1	1	1
min	.73	617		.53	082	1.6		654	0.7	35			
						4			3				

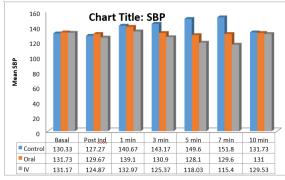


Figure 3

TABLE 4: DIASTOLIC BLOOD PRESSURE

Gro ups	Cont	rol			ignoca			NTG s	pray	p val	Cont rol	IV an	Cont rol
										ue*	and IV	d Or al	and Ora l
MA	Μ	SD	M	Μ	SD	M	Μ	SD	M		p .	p .	р
Р	ea		D	ea		D	ea		D		valu #	val	valu #
	n			n			n				e#	ue#	e#
Basa	75.	8.1		80.	5.3		78.	6.7		0.0	0.01	0.9	0.2
1	5			6			8			15	3		
Post	74.	6.7	-	76.	4.9	-	75.	6	-	0.4	0.70	1	1
ind	47	61	1.0	3		4.3	9		2.9	53	7		
			3										
1	86.	4.4	11	80.	4.8	0.2	83.	9.3	4.9	0.0	0.00	0.2	0.31
min	5	31		83	43	3	7	74		06	4	89	3
3	87.	10.	12.	77.	9.8	-	81.	11.	2.9	0.0	0.00	0.3	0.08
min	63	053	13	4	74	3.2	7	117		01	1	35	8
5	89.	9.1	14.	73.	8.9	-	79.	10.	0.8	0.0	0	0.0	0
min	8	06	3	2	07	7.4	67	981	7	01		35	
7	89.	8.4	14.	71.	8.7	-	80.	10.	1.8	0.0	0	0.0	0.00
min	83	53	33	37	98	9.2	67	61	7	01		01	1
						3							

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10	77.	4.6	2.4	75.	6.0	-	78.	6.7	-	0.2	0.57	0.3	1
min	97	57	7	97	09	4.6	4	6	0.4	37	2	37	
						3							

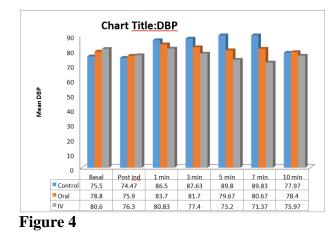
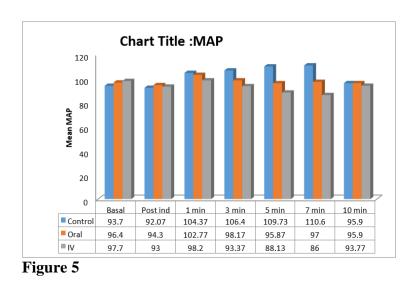


TABLE 5: MEAN ARTERIAL BLOOD PRESSURE

Gro		Con			IV			Or		р	Con	IV	Con
ups		trol						al		val ue*	trol and	an d	trol and
											IV	O ra l	Ora l
MA	Me	SD	Μ	Μ	SD	Μ	Me	SD	Μ		р	р	р
Р	an		D	ea n		D	an		D		valu e [#]	val ue [#]	valu e [#]
Bas al	93. 7	10.2		97. 7	7.4		96. 4	9.3		0.2 33	0.28 6	1	0.76 8
Post	92.	8.97	-	93	7	-	94.	8.4	-	0.5	1	1	0.89
ind	07	5	1.6 3			4.7	3		2.1	8			9
1	104	4.72	10.	98.	8.3	0.5	102	9.8	6.3	0.0	0.01	0.0	1
min	.37	3	67	2	72		.77	6	7	1		86	
3	106	9.72	12.	93.	11.	-	98.	12.	1.7	0.0	0	0.3	0.02
min	.4	9	7	37	845	4.3 3	17	785	7	01		31	1
5	109	9.27	16.	88.	11.	-	95.	12.	-	0.0	0	0.0	0
min	.73		03	13	206	9.5 7	87	566	0.5 3	01		25	
7	110	8.25	16.	86	11.	-	97	12.	0.6	0.0	0	0	0
min	.6	7	9		095	11. 7		216		01			
10	95.	6.91	2.2	93.	6.8	-	95.	9.5	-	0.4	0.89	0.8	1
min	9	5		77	41	3.9 3	9	46	0.5	82		9	

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Discussion

The sequence of induction anaesthesia, laryngoscopy and tracheal intubation are associated with marked haemodynamic changes and autonomic reflex activity which may be a cause of concern in many high-risk patients.^[3]

Laryngoscopy and intubation is associated with increase in heart rate, blood pressure and cardiac arrhythmias. These changes disappear within 5 minutes of laryngoscopy^[4] Although these responses of blood pressure and heart rate are transient, they may be detrimental in high risk patients especially in patients with cardiovascular disease and increased intracranial pressure.^[5]

Multiple factors influence the cardiovascular changes associated with laryngoscopy and intubation. Age, drugs, duration and type of procedures, depth of anaesthesia, hypoxia, hypercarbia etc., influence the pressor response. Heart rate changes decrease with increasing age, young patients show more changes. Prominent changes in haemodynamic responses are often seen in geriatric patients^{.[6]}

In our study we selected the age range of 20 to 50 years. Patients who are on antihypertensive drugs may show decrease in pressor response and so, we excluded the patients on antihypertensive medications from our study.

the combination of drugs used for premedication, induction, and maintenance of anaesthesia can alter the sympathetic response to laryngoscopy and intubation.

Midazolam at a dose of 0.2mg/kg/i.v decreases the blood pressure and increases the heart rate. However, premedication with 0.05mg/kg i.m. of midazolam has no effect on sympathetic response to laryngoscopy and intubation.^[7] Pentazocine an opioid agonist antagonist may increase the blood pressure, heart rate and catecholamine levels. Glycopyrrolate premedication can moderately increase the heart rate. Succinyl choline has negative inotropic and chronotropic effect. It acts on the muscarinic receptors of SA node. A marked noradrenergic response was noted when intubation was performed under succinylcholine^[8] Nitrous oxide may increase the tone of sympathetic nervous system. The direct action of nitrous oxide is negative inotropism which is offset by increased sympathetic tone.

In a study conducted by Singh S and Smith JE^[9] nasotracheal intubation comprises of three distinct phases a) nasopharyngeal intubation b) direct laryngoscopy to identify the vocal cords and c) Passage of tracheal tube into the trachea. Nasopharyngeal intubation causes significant pressor response. This response is exaggerated by the passage of tracheal tube in

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the larynx and trachea. Direct laryngoscopy did not increase the response significantly. In our study we included only direct laryngoscopy and orotracheal intubation.

Stanley Tam *et al.* ^[10] studied that the optimal time of injection of lignocaine before tracheal intubation for effective attenuation of circulatory response to intubation. Lignocaine 1.5mg/kg i.v. was administered 1,2,3, and 5 minutes before intubation in different groups of nonhypertensive patients belonging to ASA I or II. Lignocaine i.v given 3 minutes before intubation was found effective.

Miller CD and Warren SJ ^[11] studied the effectiveness of lignocaine to attenuate cardiovascular response to laryngoscopy and tracheal intubation. Patients were allocated into groups to receive lignocaine 1.5mg/kg i.v 1,2, and 3 minutes before laryngoscopy and failed to show any significant differences between any of these groups.

Wilson IG *et al.*^[12] studied that the varying time of prior doses of lignocaine 1.5mg/kg i.v on the cardiovascular response to tracheal intubation. There was significant rise in mean arterial pressure in all the groups. However Placebo showed rise in mean arterial pressure of 19.1% compared to lignoacaine group.

Robert K. Stoelting *et al.*^{[13} [,] in his study concluded that a short duration of direct laryngoscopy combined with laryngotracheal lidocaine administered just before intubation minimises pressor responses.

Inadae, Cullen DJ Nemeskar AR^[14] compared the efficacy of labetalol 10mg and lidocaine 2mg/kg intravenous injection just before the induction of anaesthesia and found lidocaine is a safe and cost effective mean of preventing tachycardia but not hypertension in response to laryngoscopy and intubation.

Madhuri Gopal, Sangitha^[15] in a comparative study of pressor response to laryngoscopy and intubation with oral spray of nitroglycerine and oropharyngeal spray of lignocaine concluded that Nitroglycerine oral spray in the dose of 0.8mg given 30 seconds before induction can be a better alternative in attenuating hemodynamic responses to laryngoscopy and intubation.

Binod Pegu *et al.*^[16] studied that the attenuation of stress response to laryngoscopy and intubation: sublingual nitroglycerin spray v/s intravenous fentanyl and sublingual nitroglycerin spray and concluded that combination of intravenous Fentanyl plus Nitroglycerin spray is more effective than NTG alone in attenuating the stress response following laryngoscopy and intubation.

Many strategies have been recommended which include minimising the duration of laryngoscopy to less than 20 seconds, topical application of local anaesthetics^{[17],} iv Betablockers, calcium channel blockers, clonidine, Sodium Nitroprusside, lignocaine. No single drug or technique is satisfactory^{[18].} Each technique has its advantages and disadvantages, the most obvious being that the prevention often outlasts the stimulus.

Attenuation of diastolic blood pressure is very significant in both lignocaine and Nitroglycerin groups as compared to control group until the end of 7 minutes (p<0.05). Among the two study groups lignocaine showed a better attenuation of diastolic blood pressure compared to nitroglycerin. Similarly mean arterial pressure was increased by 16.03% in control group while it increased by 6.37% in nitroglycerin group and only by 0.5% in lignocaine group compared to preinduction values by 1 minute post laryngoscopy. Attenuation of mean arterial pressure is significant in lignocaine group as compared to both nitroglycerin and control group (p>0.05). The efficiency of intravenous lignocaine over oral nitroglycerin spray in attenuation of cardiovascular responses has been verified.

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Conclusion

Based on present clinical comparative study the following conclusions can be made. Oral nitroglycerin spray significantly attenuates the sympathetic responses to laryngoscopy and tracheal intubation but Intravenous lignocaine is more efficient than oral nitroglycerin spray in attenuating the sympathetic responses to laryngoscopy and intubation. Intravenous lignocaine at a bolus dose of 2mg/kg administered 3 minutes before laryngoscopy and intubation can be recommended to attenuate the sympathetic response to laryngoscopy and intubation.

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