## EFFECT OF INTRACUFF MEDIA-ALKALINISED LIGNOCAINE, SALINE, AND AIR ON ENDOTRACHEAL TUBE INDUCED EMERGENCE PHENOMENA: A RANDOMIZED CONTROLLED STUDY

Indu S<sup>1</sup>, Arun M. G<sup>2</sup>, Taznim Mohamed<sup>3</sup>, Suvarna K<sup>4</sup>

<sup>1</sup>Additional Professor, Department of Anaesthesiology, Government Medical College, Kozhikode. <sup>2</sup>Assistant Professor, Department of Anaesthesiology, Government Medical College, Kozhikode. <sup>3</sup>Assistant Professor, Department of Anaesthesiology, Government Medical College, Kozhikode. <sup>4</sup>Associate Professor, Department of Anaesthesiology, Government Medical College, Kozhikode.

## ABSTRACT

## CONTEXT

Emergence from general anaesthesia is associated with post extubation cough, hoarseness, sore throat, and dysphagia, which may affect the smoothness of extubation. Prophylactic interventions have been studied to reduce these tracheal morbidities with varying results.

## AIMS

To compare the efficacy of air, alkalinised lignocaine and saline in maintaining intracuff pressure and reducing postoperative cough (PEC) and sore throat (POST).

## SETTINGS AND DESIGN

A randomised controlled study conducted in a teaching hospital.

### METHODS AND MATERIALS

105 patients scheduled for elective surgeries were randomly allocated into groups of 35 each. The endotracheal tube (ETT) cuffs were inflated with air, alkalinised lignocaine, or saline. The intracuff pressure (ICP) was initially set to 25-30 cm of  $H_2O$ ; measured every 30 minutes and before extubation; the minimum volume for occlusion (MOV) noted. The incidence PEC and POST were monitored.

### STATISTICAL ANALYSIS

Data analysed using Chi-square test, Fisher's exact test; Bonferroni method allowed multiple comparisons. A p value <0.05 was considered significant.

#### RESULTS

Pre-lubricated ETT cuff inflation with liquid media maintained an acceptable ICP. Saline and alkalinised lignocaine were effective in reducing PEC and POST. Alkalinised lignocaine provided smoother extubation and fared better in the early postoperative period.

## CONCLUSIONS

Pre-lubricated ETT cuffs with liquid media reduced PEC and POST. Alkalinised lignocaine showed better profile than saline. Optimum ICP reduces tracheal morbidity.

## KEYWORDS

Alkalinised lignocaine, saline, intracuff pressure, extubation cough, sore throat.

#### **KEYMESSAGES**

Monitoring and maintenance of optimum intracuff pressure is desirable following intubation. Intracuff media has a bearing on emergence phenomena.

**HOW TO CITE THIS ARTICLE:** Indu S, Arun MG, Mohamed T, et al. Effect of intracuff media-alkalinised lignocaine, saline, and air on endotracheal tube induced emergence phenomena: a randomized controlled study. J. Evid. Based Med. Healthc. 2016; 3(59), 3173-3177. DOI: 10.18410/jebmh/2016/689

Financial or Other, Competing Interest: None. Submission 30-06-2016, Peer Review 10-07-2016, Acceptance 21-07-2016, Published 25-07-2016. Corresponding Author: Dr. Indu S, Additional Professor, Department of Anaesthesiology, Government Medical College, Kozhikode. E-mail: indusatish2000@gmail.com DOI: 10.18410/jebmh/2016/689 **INTRODUCTION:** Reported incidence of PEC and POST is 38-96%.<sup>1</sup> Non-lubricated ETT, non-humidified gases, N<sub>2</sub>O diffusion, or multiple intubation attempts are probable causes. Standard techniques or media are not suggested for cuff inflation and maintenance. Recommended ICP is 20-30 cm H<sub>2</sub>O in the ICU.<sup>2</sup> Overinflation causes impairment of tracheal mucosal perfusion, underinflation predisposes to aspiration; therefore, ICP monitoring is indicated.<sup>3</sup>

# Jebmh.com

Usual cuff inflations are with air and monitoring is not routinely practiced. Saline and varying doses of NaHCO<sub>3</sub> with lignocaine have been tried for cuff inflation. In this study, 7.5% NaHCO<sub>3</sub> in high dilutions with lignocaine and saline were compared with air.

MATERIALS AND METHODS: After obtaining institutional ethical committee approval and written informed consent, 105 patients of either sex scheduled for elective surgery were randomly allocated into 3 groups of 35 each. They belonged to ASA PS I or II were aged 20 to 50 years and weighing 50 to 70 Kg. No additional risk or financial constraints were placed on them. Patients with difficult airway, hyper reactive airway, surgery of airway, those who needed nasogastric tube insertion, or more than one attempt at intubation were excluded. After adequate fasting, patients were premedicated with inj. morphine 0.1 mg/kg, inj. ondansetron 0.01 mg/kg, and inj. midazolam 0.2 mg/kg. Monitoring included ECG, pulse oximetry, capnography, and blood pressure. Patients were induced with inj. thiopentone sodium 3-6 mg/kg followed by inj. vecuronium bromide 0.12 mg/kg, and inj. lignocaine 1.5 mg/kg before intubation.

ETT of sizes 7/7.5 mm ID for females, 8/8.5 mm ID for males, prelubricated with water soluble jelly were used. Minimum Occlusion Volume (MOV), which is the volume of air or liquid injected into the cuff that eliminates an audible inspiratory leak with positive pressure ventilation was determined.

Group A received air in their cuffs, group B alkalinised lignocaine, and group C 0.9% saline. Lignocaine 2% was alkalinised using 7.5% NaHCO<sub>3</sub> in the ratio 9:1 so as to make 10 mL of solution with a pH around 8. Initial ICP was ensured to be 25-30 cm of H<sub>2</sub>O using cuff pressure manometer. Ventilation was maintained with closed circuit to ensure normocarbia. N<sub>2</sub>O:O<sub>2</sub> in 70%-30% ratio, 0.4-1% isoflurane, and vecuronium were used for maintenance. ICP was measured every 30 minutes and if found high (>40 cm H<sub>2</sub>O), was brought back to the normal range. At the end of surgery, neuromuscular blockade was reversed and ventilation assisted manually till extubation before which the final ICP was measured.

The extubation was graded based on the duration of PEC as: grade 0=no cough; grade 1=cough lasting <15 seconds; grade 2=cough lasting >15 seconds. The incidence of POST at 1 hr, 2 hr, and at 24 hr were observed and graded as follows: grade 0=No sore throat; grade 1=Tolerable pain; grade 2=Intolerable pain. Complications like laryngospasm, bucking on tube, restlessness were noted and treated. The observed data was analysed by SPSS version 18 and the variables compared using Pearson's Chi-square test, or Fisher's exact test as indicated. Bonferroni method allowed multiple comparisons and a p value <0.05 considered statistically significant.

RESU	JLT	S:
RESU		5:

Intracuff media	Air	*Alk. Lignocaine	Saline			
Age (Y)	43.63±13.66	43.03± 12.57	46.55 ±13.68			
Height (cm)	160.50±7.61	160.85±7.48	161.70±7.96			
Weight (Kg)	58.52±9.47	60.88±9.20	61.27±9.89			
ETT size (mm)	7.48±0.53	7.51±0.54	7.4±0.50			
Surgery Duration (min)	126.77±30.04	124.21±27.40	125.67±28.06			
Table 1: Demographic Profile						

Patient parameters comparable:\* Alk-Alkalinise All patients were comparable with respect to age, weight, sex, ETT size, and duration of surgery [Table 1].

Group	Number	MOV-male	MOV-female	Mean	S.D	
Air	30	7.27 mL	3.55 mL	4.54 mL	±1.85	
Alk. lignocaine	33	6.85 mL	3.90 mL	4.71 mL	±1.61	
Saline	33	7.45 mL 3.99 mL 4.83 mL		4.83 mL	±1.84	
Table 2: Comparison of Minimum Occlusion Volume (MOV)						

MOV was higher in males.

Five in group A, 2 each in groups B and C were excluded as they needed nasogastric tube insertion, intravenous lignocaine for treating stress response, or multiple attempts at intubation. The MOV were comparable in all the groups with liquids requiring a slightly higher volume, which was not statistically significant [Table 2]. The occlusion volume in males (mean 7.2 mL) was higher when compared to females (mean 3.8 mL) [Table 2]. The initial ICP was 25-30 cm H<sub>2</sub>O. ICP steadily increased with time, the air group showing significantly higher values (46.5 cm $\pm$ 3.54 H<sub>2</sub>O). For alkalinised lignocaine, it was 33.17 $\pm$ 4.21 cm H<sub>2</sub>O and for saline, 33 $\pm$ 3.83 cm H<sub>2</sub>O. The rate of rise ICP and its magnitude was greatest in the first 30 minutes; this was statistically significant the air group. The ICP gradually increased with the duration of surgery in all the groups.

Time in Minutes	No	Air (In cm H <sub>2</sub> O)	No	Alkalinised Lignocaine (In cm H <sub>2</sub> O)	No	Saline (In cm H <sub>2</sub> O)	
Start	30	24.43±1.65	33	24.70±1.43	33	24.88±1.50	
30 m	30	33.07±2.99	33	26.97±2.21	33	26.45±1.54	
60 m	30	36.37±3.67	33	28.39±2.81	33	27.82±2.22	
90 m	30	38.93±3.97	33	29.48±2.76	33	28.94±2.63	
120 m	15	42.33±4.03	15	30.93±3.53	16	31.12±2.82	
150 m	10	45.70±3.86	6	33.17±4.21	9	32.44±2.70	
180 m	2	46.50±3.54	2	31.00±7.07	4	33.00±3.83	
Final	30	43.57±5.63	33	30.94±3.74	33	31.45±4.61	
	Table 3: Intracuff Pressure (ICP)						

Steady increase in ICP; the air group showing significantly higher values. For the liquid group, there was a steady minimal rise in ICP; but was within acceptable range [Table 3]. Patients with no cough (PEC grade 0) or cough less than 15 seconds (PEC grade II) and were considered to have smooth extubation. The extubation was smooth in 18 out of 30(60%) for the air group, 29 out of 33(87.9%) for the lignocaine group, and 22 out of 33(60.7%) for the saline group.

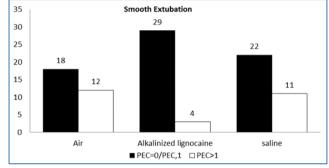


Fig. 1: PEC Grade 0 and Grade1 were Considered as Smooth Extubation (p Value 0.04)

Alkalinised lignocaine offered reduced incidence of cough and smoother extubation (p value 0.04) [Figure1].

Time	Total No.	Air	Alkalinised Lignocaine	Saline	P value	
1 h	30	13(43.3%)	3(9.1%)	10(30.3%)	0.008	
2 h	33	20(66.7%)	15(45.5%)	14(42.4%)	0.115	
24 h	33	25(83.3%)	21(63.6%)	24(72.7%)	0.214	
Table 4: Postoperative Sore Throat (POST)						

POST least for alkalinised lignocaine (p value 0.008) at 1 hour. The incidence of POST in first hour was significantly lower for alkalinised lignocaine 9.1% (p value of 0.008) [Table 4]. In the second hour, 20 out of 30(66.7%) in air group, 15 out of 33(45.5%) in the lignocaine group, and 14 out of 33(42.4%) in the saline group had POST.

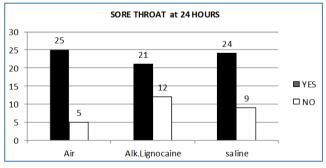


Fig. 1: Sore Throat in 24 Hours (P Value 0.214); Lignocaine Group with A Better Profile

At 24 h, 25 out of 30(83.3%) in the air group, 21 out of 33(63.6%) in lignocaine group, 24 out of 33(72.7%) patients in saline group had POST [Figure 2]. At two hours,

alkalinised lignocaine and saline produced almost similar effects but fared better than air.

Amongst the 24 hour incidence of POST, the least was noted with the lignocaine group followed by saline and then air although statistically not significant.

**DISCUSSION:** In our study, we used either air, alkalinised lignocaine (1 mL 7.5% NaHCO<sub>3</sub>: 9 mL 2% lignocaine) or 0.9% normal saline for endotracheal cuff inflation and compared their effects on ICP and the quality of extubation. Buffering lignocaine with 7.5% NaHCO<sub>3</sub> in this ratio produces a solution with a pH of about eight. This is likely to give an adequate concentration of unionized fraction of lignocaine for trans-cuff diffusion providing optimum clinical effects. The results obtained were in good correlation with studies done using lower concentrations of (1.4%) NaHCO3 to alkalinise 2% lignocaine.

At higher concentrations, NaHCO3 may precipitate and is likely to block the pilot balloon during cuff inflation. The duration of surgery in our study was around 120 to 130 minutes. The rise in ICP was high with air as media especially in the initial period (30 minutes), but was gradual and within acceptable limits when liquids were used for cuff inflation. Each time, the ICP reached above 40 cm  $H_2O$ . It was brought back to the normal range. The volume needed for cuff inflation (MOV) was comparable in all the three groups and was on an average 4-6 mL.

This was higher for males. The maintenance of ICP was better in those with alkalinised lignocaine and saline in their cuffs. When compared with saline, alkalinised lignocaine had a better profile in the quality of extubation in maintaining the ICP and in reducing POST. This is in concordance with previous studies. Considering a remote possibility of cuff rupture, saline may have a theoretical advantage over alkalinised lignocaine. The main goal of anaesthesiologist is to provide smooth extubation. During emergence, patient may experience restlessness and cough, which may coexist with or precipitate bronchospasm, laryngospasm, or even negative pressure pulmonary oedema. This can also increase the intraocular, intragastric, intrathoracic, and intracranial pressures.<sup>4</sup>

An important factor maybe direct irritation and resultant inflammatory reaction within the tracheal mucosa caused by the ETT cuff. High ICP caused by large volumes of media used for cuff inflation along with N<sub>2</sub>O diffusion can cause transient reversible ischemic changes in the adjacent tracheal mucosa.<sup>5</sup> However, newer low pressure-high volume cuffs tend to withstand the rise in cuff pressure due to N<sub>2</sub>O influx to some extent. Topical anaesthetics and steroids have been tried to prevent PEC and POST.<sup>6</sup> ETT prelubricated with water soluble jelly and alkalinised lignocaine cuffs are claimed to reduce POST.<sup>7</sup> Ideal ICP is that which provides an airway seal that would allow positive pressure ventilation with minimal leak.

Tracheal pressure >48 cm H<sub>2</sub>O was seen to reduce capillary blood flow while pressures <18 cm H<sub>2</sub>O increases the risk of aspiration. ICP of 25-40 cm H<sub>2</sub>O protects from aspiration, avoids traumatizing trachea, and reduces PEC and POST.<sup>8</sup> Over inflation of cuff induced by N<sub>2</sub>O intraoperatively can be reduced to some extent by substitution of air with saline resulting in maintaining sustained and optimal ICP.9,10 Lignocaine (4%) spray was also found to be more effective than intracuff alkalinised lignocaine in reducing PEC in surgeries <2 hours.<sup>11</sup> Lignocaine (4%) instillation in ETT cuffs reduced incidence of cough better than intravenous lignocaine in neurosurgical patients.<sup>[12]</sup> PEC in smokers was found to be lesser with 2% alkalinised lidocaine filled ETT cuff.[13] Preoperative nebulization with ketamine was found to reduce the severity of sore throat in a recent study.<sup>[14]</sup>

In a systematic review, it was concluded that both topical and systemic lidocaine were effective in reducing POST.<sup>15</sup> A meta-analytical study confirmed that lignocaine both alkalinised and non-alkalinised instilled in endotracheal tube cuffs were effective in reducing PEC and POST.<sup>[16]</sup> Alkalinising lignocaine allowed better diffusion as the hydrophobic neutral base form of lignocaine was able to diffuse across the cuff membrane (65% of diffusion during 6 h) as against the commercially available charged hydrochloride form where only 1% of total diffuses to block the tracheal receptors. This necessitated a larger dose of

lignocaine hydrochloride (200-500 mg) to exert a clinical effect. Adding 8.4% NaHCO<sub>3</sub> allows a smaller dose of lignocaine to be used (20-40 mg), so that only a very small concentration of lignocaine is attained (<0.08 µg/mL), which is lower than when lignocaine is used topically (0.43-1.5 µg/mL) or intravenously (2-3 µg/mL).<sup>17</sup> In one RCT, it was noted that alkalinised lignocaine instilled cuffs in longer duration surgeries produced better outcome as diffusion across membrane is a function of time.<sup>18</sup>

In our study, majority of surgeries were of more than 120 minutes duration, which might have contributed to the desired clinical effect. Increasing the pH through alkalinisation increases the unionized fraction available for diffusion. Effective suppression of cough receptors required more than 60 minutes.<sup>19</sup> But, at a higher pH, there is a tendency towards slower release of lignocaine and a chance of injuring the tracheal mucosa in case the cuff ruptures.<sup>20</sup> In another study, alkalinising lignocaine to a pH 7.4 to 7.6 was effective within 30 minutes, but peak action was seen in 300 minutes. Intracuff buffered lignocaine (6 mL lignocaine: 0.5 mL 7.5% NaHCO3) produces a concentration of 155 µg/mL on tracheal receptors in about 90 minutes producing clinical effects after trans-cuff diffusion.<sup>21</sup>

Lowering the concentration of NaHCO<sub>3</sub> to 1.4% for alkalinising 2% lignocaine was also effective in reducing the incidence of emergence phenomenon. In vitro studies confirm that even lower volumes of NaHCO<sub>3</sub> can produce greater release of lidocaine across low volume high pressure cuffs and variations in the volume and concentration of NaHCO3 had no effect on the diffusion of lignocaine.<sup>22</sup> A recent study using 2% lignocaine and 8.4% NaHCO<sub>3</sub> in the ratio 19:1 noted reduced cough, sore throat, and hoarseness in smokers.<sup>23</sup> Significant reduction in the sedation and analgesic requirement was noted in mechanically ventilated patients when alkalinised lignocaine was instilled in their ETT cuffs.<sup>24</sup> According to the review of evidence, there is need for routine ICP monitoring for every anaesthetic as well as in the ICU.<sup>25</sup>

**CONCLUSION:** Maintenance of ICP within acceptable range was better when liquids were used as inflating media. If alkalinised lignocaine is not available, the easy availability of saline makes it a better option than air. The average volume of cuff occlusion was 4 to 6 mL to maintain an ICP of 20-30 cm  $H_2O$ . Extubation was smoother and postoperative sore throat lesser with liquids especially alkalinised 2% lignocaine. As a routine saline or preferably alkalinised, 2% lignocaine need to be used for ETT cuff instillation. Intracuff pressure monitoring and maintenance of optimum ICP significantly reduces emergence phenomena and tracheal morbidity following endotracheal intubation.

## REFERENCES

 Crerar C, Weldon E, Salazar J, et al. Comparison of 2 laryngeal tracheal anaesthesia techniques in reducing emergence phenomena. AANA J 2008;76(6):425-431.

# Jebmh.com

- Estebe JP, Dollo G, Le Corre P, et al. Alkalinisation of intracuff lidocaine improves endotracheal tubeinduced emergence phenomena. Anaesth Analg 2002;94(1):227-230.
- 3. Talekar CR, Udy AA, Boots RJ, et al. Tracheal cuff pressure monitoring in the ICU: a literature review and survey of current practice in Queensland. Anaesth Intensive Care 2014;42(6):761-770.
- Nseir S, Brisson H, Marquette CH, et al. Variations in endotracheal cuff pressure in intubated critically ill patients: prevalence and risk factors. Eur J Anaesthesiol 2009;26(3):229-234.
- 5. Scuderi PE. Postoperative sore throat: more answers than questions. Anaesth Analg 2010;111(4):831-832.
- Bensaid S, Duvaldestin P, Lieutaud T, et al. Nitrous oxide increases endotracheal cuff pressure and the incidence of tracheal lesions in anaesthetized patients. Anaesth Analg 1999;89(1):187-190.
- Sumathi PA, Shenoy T, Ambareesha M, et al. Controlled comparison between betamethasone gel and lignocaine jelly applied over tracheal tube to reduce postoperative sore throat, cough, and hoarseness of voice. Br J Anaesth 2008;100(2):215-218.
- Estebe JP, Delahaye S, Le corre P, et al. Alkalinisation of intracuff lidocaine and use of gel lubrication protect against tracheal tube-induced emergence phenomena. Br J Anaesth 2004;92(3):361-366.
- Stewart SL, Secrest JA, Barbara R, et al. A comparison of endotracheal tube cuff pressures using estimation techniques and direct intracuff measurement. AANA Journal 2003;71(6):443-447.
- Shroff, Prerana P, Patil, et al. Efficacy of cuff inflation media to prevent post intubation-related emergence phenomenon: air, saline, and alkalinised lignocaine. European Journal of Anaesthesiology 2009;26(6):458-462.
- 11. Combes X, Schauvliege F, Peyrouset O, et al. Intracuff pressure and tracheal morbidity: influence of filling with saline during nitrous oxide anaesthesia. Anaesthesiology 2001;95(5):1120-1124.
- D'Argon F, Beaudet N, Gagnon V, et al. The effects of lidocaine spray and intracuff alkalinised lidocaine on the occurrence of cough at extubation: a double-blind randomised controlled trial. Can J Anaesth 2013;60(4):370-376.
- Venkatesan T, Korula G. A comparative study between the effects of 4% endotracheal tube cuff lignocaine and 1.5 mg/kg intravenous lignocaine on coughing and haemodynamics during extubation in neurosurgical patients: a randomised controlled double-blind trial. J Neurosurg Anaesthesiol 2006;18(4):230-234.

- 14. Ahuja V, Mitra S, Sarna R. Nebulized ketamine decreases incidence and severity of postoperative sore throat. Indian J Anaesth 2015;59(1):37-42.
- 15. Kori K, Muratani T, Tatsumis, et al. Influence of endotracheal tube cuff lubrication on postoperative sore throat and hoarseness. Masui 2009;58(3):342-345.
- 16. Tanaka Y, Nishimori M, Tsujimura Y, et al. Lidocaine for preventing postoperative sore throat. Cochrane Database Sys Rev 2015;7:CD004081.
- 17. Lam F, Lin YC, Tsai HC, et al. Effect of intracuff lidocaine on postoperative sore throat and the emergence phenomenon: a systematic review and meta-analysis of randomised controlled trials. PLoS ONE 2015;10(8):e0136184.
- Dollo G, Estebe JP, Le Corre P, et al. Endotracheal tube cuffs filled with lidocaine as a drug delivery system: in vitro and in vivo investigations. Eur J Pharm Sci 2001;13(3):319-323.
- 19. Bennett MH, Isert PR, Cumming RG. Postoperative sore throat and hoarseness following tracheal intubation using air or saline to inflate the cuff-a randomised controlled trial. Anaesth Intensive Care 2000;28(4):408-413.
- 20. Fagan C, Frizelle HP, Laffey J, et al. The effects of intracuff lidocaine on endotracheal tube-induced emergence phenomena after general anaesthesia. Anaesth Analg 2000;91(1):201-205.
- 21. Jaichandran VV, Angayarkanni N, Karunakaran C, et al. Diffusion of lidocaine buffered to an optimal pH across the endotracheal tube cuff-an in vitro study. Indian J Anaesth 2008;52(5):536-540.
- 22. Estebe JP, Gentili M, Le Corre P, et al. Alkalinisation of intracuff lidocaine: efficacy and safety. Anaesth Analg 2005;101(5):1536-1541.
- 23. Navarro LH, Lima RM, Aguiar AS, et al. The effect of intracuff alkalinised 2% lidocaine on emergence coughing, sore throat, and hoarseness in smokers. Rev Assoc Med Bras 2012;58(2):248-253.
- 24. Basuni AS. Intracuff alkalinised lidocaine reduces sedative/analgesic requirements for mechanically ventilated patients. S J Anaesth 2014;8(4):451-455.
- 25. Sultan P, Carvallo B, Rose BO, et al. Endotracheal tube cuff pressure monitoring: a review of evidence. J perioper pract 2011;21(11):379-386.