



Comparative study between lidocaine and Metoprolol on hemodynamic attenuation during laryngoscope and endotracheal intubation

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Abstract :

Haemodynamic stability is an integral and essential goal of any anaesthetic management plan. Laryngoscopy and intubation can cause striking changes in haemodynamics. **Objective** to compare lidocaine group with metoprolol group on the attenuation of hemodynamic responses to laryngoscopy and tracheal intubation, undergoing general anesthesia.

This randomized controlled study was conducted at collage health and medical technologist in Teaching Baghdad Hospital, from (NOV._)2017. The study was conducted in 30 patients posted for elective surgery under general anesthesia. The patients were randomized into two equal groups to receive metoprolol 0.075 mg/kg (Group M = 15 patients) and lidocaine 1mg/kg (Group L = 15 patients). Following study drugs, anesthesia induced with Ketamine 1 mg / kg, propofol 2mg /kg and Rocuronium 0.6mg / kg IV. After 3 minutes of injection of Rocuronium, Endotracheal intubation was done. Anesthesia was maintained with (1.2%) isoflurane, oxygen and Rocuronium. All intubation were done within 30 seconds then SBP, DBP, MAP and HR were recorded just after intubation, 1 minute, 3 minutes, 6 minutes and 10 minutes of intubation. We concluded metoprolol has more effect than lidocaine on hemodynamic attenuation during four times interval (1min, 3min, 6min and 10min)

Keywords: Hemodynamic parameters , Lidocaine , Metoprolol .

Introduction:

Laryngoscopy and tracheal intubation are essential in providing general anesthesia, but they produce sympathetic over drive by catecholamine release resulting in hypertension and tachycardia [1]. The hemodynamic stability can be simply defined as the stable blood flow, it means that a person has a stable heart pump and good circulation of blood, while hemodynamic instability is defined as any instability in blood pressure which can lead to inadequate arterial blood flow to organs, so critically ill patients need to be carefully monitored to avoid complications due to hemodynamic instability[2].

The main cause of transient hemodynamic instability and interruption of patient air way reflex is laryngoscopy and intubation[3]. The reflex cardiovascular responses to laryngoscopy and tracheal intubation were known to anaesthesiologists since a long time, so the haemodynamic stability is an integral and essential goal of any anaesthetic management plan , as well as endotracheal intubation stimulates the laryngeal and tracheal sensory receptors, resulting an increase of sympathetic stimulation[4].

The sympathetic stimulation results in tachycardia and elevation of blood pressure that may produce an exaggerated hypertensive response, especially in hypertensive patients, which may lead to cardiac arrest or cerebral stroke. Hemodynamic responses to laryngoscopy incubation were first described by ried and brace in 1940, and these responses are of little significance in healthy patients but may be fatal in patients with heart diseases and high blood pressure [5], the effect of drugs on the haemodynamic_responses can be known by monitoring the heart rate(HR), respiratory rate(RR), Blood pressure, Mean arterial

pressure(MAP), electrocardiography(ECG) and by calculating the Rate pressure product, Heart rate and the rate pressure product as determinants of cardiovascular risk in patients with hypertension.

There are several methods in modifying the hemodynamic responses: Lidocaine has a suppressive effect on the circulatory responses in patients undergoing laryngoscopy and tracheal intubation [6]. whilst the β -adrenergic receptor antagonist like metoprolol decrease occurrence of arrhythmias and tachycardia caused by stressful stimulus during anaesthesia and reduce the neuroendocrine responses to surgical stimuli[7].

Aim of study: To assess the comparative effect between lidocaine and Metoprolol on hemodynamic changes systolic blood pressure(SBP); diastolic blood pressure(DBP); pulse rate(PR) and mean arterial blood pressure(MAP) attenuation during direct laryngoscopy and endotracheal intubation through the 4 time intervals(1min,3min,6min,10min).

Materials and Methods:

This randomized controlled study was conducted at collage of Health and Medical Technologist in Baghdad Teaching Hospital, from(NOV._)2017. The study design was approved from anesthetic department and informed consent was received from each patient after detailed explanation of the procedure. Thirty patients were selected with American Society of Anesthesiologists (ASA) physical status I and II, aged between 15 to 60 years, and weighed 45 - 100 kg who are undergoing abdominal and pelvic surgery under general anesthesia were randomly allocated. The patients were divided into two groups: Lidocaine group (n = 15) and Metoprolol group (n = 15).

Exclusion criteria:1- Patients have a history of cardiovascular disease 2_cerebrovascular disease 3-psychosis 4-recent drug abuse 5-

respiratory problem 6- difficulty in intubation or repeated attempting of intubation were excluded from the study. Pre-anesthetic check-up was done in all the patients one day before surgery .after arrival at the operation theater, baseline parameters like heart rate (HR), systolic blood pressure (SBP) and the diastolic blood pressure (DBP) were measured by noninvasive blood pressure(NIBP).In addition, SpO₂ and ECG were monitored continuously. IV cannulation was done with 20 gauge cannula. Patients received one of the two standardized treatment regimens prior to laryngoscopy, lidocaine group was received 1mg / kg iv lidocaine , and metoprolol group was received 0,075mg / kg metoprolol based on institutional protocol. All the drugs were prepared to the total volume of 12 ml to prevent biase. Drug preparation and the patients were done by the assistant anesthetic who was not involved in the study.

The patients and the researcher were blinded with the stress blunting agents given. Induction was done with Ketamine 1 mg / kg, propofol 2mg /kg and Rocuronium 0.6mg / kg IV. After 3minutes of injection of Rocuronium , Endotracheal intubation was done. Anesthesia was maintained with(1.2%)isoflurane with oxygen and Rocuronium. All intubation were done within 30 seconds then SBP, DBP, MAP and HR were recorded just after intubation, 1 minute,3minutes, 6 minutes and 10 minutes of intubation.

Result:

Table(1):comparative effects between the studied groups(lidocaine group VS metoprolol group) on the heart rate in different time interval.

Group Statistics				
Heart rate	Drugs	Mean	Std.	P Value

			Deviation	
baseline	Lidocaine	90.07	8.72	0.568
	Metoprolol	91.73	6.99	
1 min	Lidocaine	94.87	7.86	0.002
	Metoprolol	84.8	8.42	
3min	Lidocaine	91.6	8.71	0.000
	Metoprolol	75.73	7.85	
6min	Lidocaine	89.13	8.44	0.000
	Metoprolol	66.4	6.48	
10min	Lidocaine	88.07	9.04	0.000
	Metoprolol	55.6	5.29	

- There was no significant difference in heart rate between two groups (lidocaine group VS metoprolol group) at base line (p-value=0.568), While there was significant difference in heart rate between two groups at(1min,3min,6min and 10min) whereas metoprolol significantly reduced the heart rate as compere to lidocaine .

Table(2):comparative effects between the studied groups(lidocaine group VS metoprolol group) on the mean atrial blood pressure in different time interval

Group Statistics				
Mean artrial blood pressure	Drugs	Mean	Std. Deviation	P Value

Baseline	Lidocaine	94.67	7.12	0.004
	Metoprolol	101.4	4.5	
1 min	Lidocaine	99.13	6.8	0.001
	Metoprolol	91.13	4.66	
3min	Lidocaine	91.07	7.06	0.000
	Metoprolol	79.8	3.12	
6min	Lidocaine	86.47	6.44	0.000
	Metoprolol	70.93	2.02	
10min	Lidocaine	83.47	5.94	0.000
	Metoprolol	63.13	3.78	

- Metoprolol decreased mean blood pressure after (1,3,6,10min)significantly while in Lidocaine group elevated B.p was recorded after 1.minute ,then followed by decrease in B.P after 3,6,10min .
- There was significant difference in mean arterial blood pressure between two groups (lidocaine group VS metoprolol group) at (base line,1min,3min,6min and 10min) .

Table(.3):comparative effect between the studing groups(lidocaine group VS metoprolol group) on the dystolic blood pressure in different time interval.

Group Statistics				
Dystolic blood pressure	Drugs	Mean	Std. Deviation	P Value
Baseline	Lidocaine	79.33	7.58	0.011
	Metoprolol	85.87	5.32	
1 min	Lidocaine	83.87	7.61	0.012

	Metoprolol	77.47	5.17	
3min	Lidocaine	75.8	7.85	0.013
	Metoprolol	69.07	5.92	
6min	Lidocaine	71.27	6.97	0.000
	Metoprolol	61.67	3.11	
10min	Lidocaine	68.93	6.75	0.000
	Metoprolol	54.27	3.51	

- Metoprolol decreased diastolic blood pressure after (1,3,6,10min) significantly while in Lidocaine group elevated B.p was recorded after 1.minute ,then followed by decrease in B.P after 3,6,10min .

-There was significant difference in dystolic blood pressure between two groups (lidocaine group VS metoprolol group) at (base line,1min,3min,6min and 10min).

Table(4):comparative effect between the studing groups(lidocaine group VS metoprolol group) on the systolic blood pressure in different time interval.

Group Statistics				
Systolic blood pressure	Drugs	Mean	Std. Deviation	P Value
baseline	Lidocaine	126.93	8.69	0.026
	Metoprolol	133.87	7.45	
1 min	Lidocaine	131.4	8.38	0.000
	Metoprolol	117.8	7.89	
3min	Lidocaine	122.33	7.65	0.000
	Metoprolol	101.47	4.84	

6min	Lidocaine	117.93	7.05	0.000
	Metoprolol	91.07	4.27	
10min	Lidocaine	113.8	6.13	0.000
	Metoprolol	84.73	4.53	

Metoprolol decreased systolic blood pressure after (1,3,6,10min) significantly while in Lidocaine group elevated B.p was recorded after 1.minute ,then followed by decrease in B.P after 3,6,10min .

-There was significant difference in systolic blood pressure between two groups (lidocaine group VS metoprolol group) at (base line,1min,3min,6min and 10min).

Discussion:

The hemodynamic changes stemming from airway instrumentation are due to sympathoadrenal discharges caused by epipharyngeal and par pharyngeal stimulation[49]. Tracheal intubation is associated with an increase in heart rate and blood pressure, however, at the emergence of anesthesia additional haemodynamic responses to pain makes tracheal extubation more complicated[50].

Reid and Brace were the first to report the circulatory response to laryngeal and tracheal stimulation in anaesthetized man as tachycardia and increase in arterial blood pressure.Lidocaine and metoprolol have a suppressive effect on the circulatory responses in patients undergoing laryngoscopy and tracheal intubation [51].

The present study showed that both groups lidocaine group (1 mg/kg) and metoprolol group (0.075 mg/kg) were unequally effective in decreasing hemodynamic parameters (systolic, diastolic, mean blood pressures and heart

rate) in both group (lidocaine and metoprolol) which statistically highly significant ($P < 0.000$).

lidocaine's effect on cardiomyocytes is inadequate to prevent heart repolarization[52]. Due to its anti-arrhythmic characteristics, lidocaine is used in the treatment of heart rate disturbances of ventricular origin [53]. In an earlier study indicates that effect of lidocaine could be associated with protective activation of the sympathetic system secondary to airway manipulation and, thus, inhibition of the prolonged repolarization[54].

The metoprolol more effective than lidocaine in attenuation the heart rate ,systolic blood pressure , diastolic blood pressure and mean arterial pressure during four times interval(1min,3min,6min and 10min) . Because metoprolol blocks sympathetic nervous system and decrease in heart rate and blood pressure[33].

Conclusions: Based on results of present study it may be concluded that both group (lidocaine group and metoprolol group) prevent the hemodynamic changes, However Clinically and statistically it can be concluded that metoprolol may be a better agent for hemodynamic attenuation .Lidocaine may be prefers for cases in which the Beta -Blockers are contraindicated.

Recommandation:

1. Metoprolol is highly recommended for attenuation of sympathetic response to direct laryngoscopy as a sole agent.
2. Detailed studies with larger samples are recommended to emphasize on the benefits of using metoprolol in patients with ischaemic heart disease.
3. Comparative effect study between lidocaine and Metoprolol in asthmatic patients>

References

- 1- Raju Basu S M, Pramanik. *Duration and degree of circulatory changes following laryngoscopy and intubation and study of a method of attenuation of these changes.*Ind. J. Anaesth 1988;26:29-33.,2013.
- 2- halevy O, Nadel M, Barak, Rozenboim. *Hemodynamic and catecholamine response to tracheal intubation: direct laryngoscopy compared with fiberoptic intubation.* J Clin Anesth, 15 (2) (2003), pp. 132-136.
- 3- Bruder N, Ortega D, Granthil C. *Consequences and prevention methods of hemodynamic changes during laryngoscopy and intratracheal intubation.* Ann Fr Anesth Reanim. 1992;11:57-71.
- 4- Basu S M, Pramanik. *Duration and degree of circulatory changes following laryngoscopy and intubation and study of a method of attenuation of these changes.*Ind. J. Anaesth 1988;26:29-33.
- 5- Imani F, Alebouyeh MR, Taghipour Anvari Z, Faiz SH. *Use of remifentanil and alfentanil in endotracheal intubation: a comparative study.* Anesth Pain Med. 2011;1(2):61–5. doi: 10.5812/kowsar.22287523.2130.
- 6- Kim WY, Lee YS, Ok SJ, Chang MS, Kim JH, Park YC, et al. *Lidocaine does not prevent bispectral index increases in response to endotracheal intubation.* Anesth Analg. 2006;102(1):156–9.
- 7- Zachariah, S.K. and Ninan ,S.(2015). *Hemodynamic Responses to Microlaryngoscopy in ENT Patients -A Comparative Study of Premedication with Oral Clonidine and Metoprolol . Int J Med Health Sci., Vol-4; Issue-4).*
- 8- "Lidocaine Hydrochloride (Local)". *The American Society of Health-System Pharmacists.* Archived from the original on 2015-09-06.
- 9- Picard J, Ward SC, Zumpe R, Meek T, Barlow J, Harrop-Griffiths W (February 2009). *"Guidelines and the adoption of 'lipid rescue' therapy for local*

anaesthetic toxicity". Anaesthesia. 64 (2): 122–5. doi:10.1111/j.1365-2044.2008.05816.x.

10- Gulihar, Abhinav; Robati, Shibby; Twaij, Haider; Salih, Alan; Taylor, Grahame J.S. (December 2015). "Articular cartilage and local anaesthetic: A systematic review of the current literature". *Journal of Orthopaedics*. 12: S200 S210.

11- Carterall, William A. (2001). "Molecular mechanisms of gating and drug block of sodium channels". *Sodium Channels and Neuronal Hyperexcitability. Novartis Foundation Symposia*. 241. pp. 206–225.

12- Sheu SS, Lederer WJ (Oct 1985). "Lidocaine's negative inotropic and antiarrhythmic actions. Dependence on shortening of action potential duration and reduction of intracellular sodium activity". *Circulation Research*. 57 (4): 578–90.

13- Lewin NA, Nelson LH (2006). "Chapter 61: Antidysrhythmics". In Flomenbaum N, Goldfrank LR, Hoffman RL, Howland MD, Lewin NA, Nelson LH. *Goldfrank's Toxicologic Emergencies (8th ed.)*. New York: McGraw-Hill. pp. 963–4. ISBN 0-07-143763-0.

14- "Local anaesthetic drugs".

15- Cepeda MS, Tzortzopoulou A, Thackrey M, Hudcova J, Arora Gandhi P, Schumann R (2010). "Adjusting the pH of lidocaine for reducing pain on injection". *Cochrane Database Syst Rev*(12): CD006581.

16- Derry, S; Wiffen, PJ; Moore, RA; Quinlan, J (24 July 2014). "Topical lidocaine for neuropathic pain in adults". *The Cochrane Database of Systematic Reviews*. 7:CD010958.

17- Lidocaine/prilocaine spray for premature ejaculation". *Drug and therapeutics bulletin*. 55 (4): 45–48. April 2017.

18- Martí-Carvajal, AJ; Simancas-Racines, D; Anand, V; Bangdiwala, S .2015. "Prophylactic lidocaine for myocardial infarction". *The Cochrane Database of Systematic ,Reviews*. 8: CD008553.

- 19- Slaughter, LA; Patel, AD; Slaughter, JL (March 2013). "Pharmacological treatment of neonatal seizures: a systematic review". *Journal of child neurology*. 28 (3): 351–64.
- 20- Biller JA (2007). "Airway obstruction, bronchospasm, and cough". In Berger AM, Shuster JL, Von Roenn JH. *Principles and practice of palliative care and supportive oncology*. Hagerstown, MD: Lippincott Williams & Wilkins. pp. 297–307. ISBN 978-0-7817-9595-1. Inhaled lidocaine is used to suppress cough during bronchoscopy. Animal studies and a few human studies suggest that lidocaine has an antitussive effect...
- 21- Birsa LM, Verity PG, Lee RF. "Evaluation of the effects of various chemicals on discharge of and pain caused by jellyfish nematocysts". *Comp. Biochem. Physiol. C Toxicol. Pharmacol*. 151 (4): 426–30.
- 22- Morabito R, Marino A, Dossena S, La Spada G (Jun 2014). "Nematocyst discharge in *Pelagia noctiluca* (Cnidaria, Scyphozoa) oral arms can be affected by lidocaine, ethanol, ammonia and acetic acid". *Toxicon*. 83: 52–8. .
- 23- James G. Adams (2012). "32". *Emergency Medicine: Clinical Essentials*. Elsevier Health Sciences. ISBN 9781455733941. Archived from the original on 2017-09-08.
- 24- Segal MM, Rogers GF, Needleman HL, Chapman CA (Dec 2007). "Hypokalemic sensory overstimulation". *Journal of child neurology*. 22 (12): 1408–10.
- 25- Hakim AJ, Grahame R, Norris P, Hopper C (February 2005). "Local anaesthetic failure in joint hypermobility syndrome". *J R Soc Med*. 98 (2): 84–5.
- 26- "Lidocaine international forms and names". *Drugs.com*. Retrieved 29 October 2017.
- 27- "Lidocaine Hydrochloride (Antiarrhythmic)". *The American Society of Health-System Pharmacists*. Archived from the original on 2015-08-10. Retrieved Aug 26, 2015.
- 28- "Lidocaine". *Epocrates*. Archived from the original on 2014-04-22.

- 29- *"Lidocaine Hydrochloride and 5% Dextrose Injection". Safety Labeling Changes. FDA Center for Drug Evaluation and Research (CDER). January 2014. Archived from the original on 2013-04-03.*
- 30- *"Lidocaine Viscous: Drug Safety Communication - Boxed Warning Required - Should Not Be Used to Treat Teething Pain". FDA Center for Drug Evaluation and Research (CDER). June 2014. Archived from the original on 2014-07-14.*
- 31- *"Lidocaine - N01BB02". Drug porphyrinogenicity monograph. The Norwegian Porphyria Centre and the Swedish Porphyria Centre. Archived from the original on 2014-04-20. strong clinical evidence points to lidocaine as probably not porphyrinogenic*
- 32- *Nielsen, LJ; Lumholt, P; Hölmich, LR (27 October 2014). "[Local anaesthesia with vasoconstrictor is safe to use in areas with end-artries in fingers, toes, noses and ears.]". Ugeskrift for laeger. 176 (44). PMID 25354008.*
- 33- *"Metoprolol". The American Society of Health-System Pharmacists. Archived from the original on 2014-03-12. Retrieved Apr 21, 2014.*
- 34- *Cupp M (2009). "Alternatives for Metoprolol Succinate" (pdf). Pharmacist's Letter / Prescriber's Letter. 25 (250302)..*
- 35- *Suita, Kenji; Fujita, Takayuki; Hasegawa, Nozomi; Cai, Wenqian; Jin, Huiling; Hidaka, Yuko; Prajapati, Rajesh; Umemura, Masanari; Yokoyama, Utako (2015-07-23).*
- 36- *"Metoprolol". The American Society of Health-System Pharmacists. Archived from the original on 25 April 2011. Retrieved 3 April 2011.*
- 37- *MERIT-HF Study Group (1999). "Effect of metoprolol CR/XL in chronic heart failure: Metoprolol CR/XL Randomised Intervention Trial in Congestive Heart Failure (MERIT-HF)". Lancet. 353 (9169): 2001–2007.*
- 38- *Biffi, M.; Boriani, G.; Sabbatani, P.; Bronzetti, G.; Frabetti, L.; Zannoli, R.; Branzi, A.; Magnani, B. (Mar 1997). "Malignant vasovagal syncope: a randomised trial of metoprolol and clonidine". Heart. 77 (3): 268–72.*

- 39- Geffner DL, Hershman JM (July 1992). " β -Adrenergic blockade for the treatment of hyperthyroidism". *The American Journal of Medicine*. 93 (1): 61–8.
- 40- Kühlkamp, V; Schirdewan, A; Stangl, K; Homberg, M; Ploch, M; Beck, OA (2000). "Use of metoprolol CR/XL to maintain sinus rhythm after conversion from persistent atrial fibrillation". *J Am Coll Cardiol*. 36 (1): 139–146.
- 41- Pillay (2012). *Modern Medical Toxicology*. Jaypee Brothers Publishers. p. 303. ISBN 9789350259658. Archived from the original on 2017-07-07.
- 42- *Medical Toxicology*. Lippincott Williams & Wilkins. 2004.p. 684. ISBN 9780781728454. Archived from the original on 2017-07-07.
- 43- Metoprolol". *Drugs.com*. Archived from the original on 2010-01-21.
- 44- Cruickshank JM (2010). "Beta-blockers and heart failure". *Indian Heart Journal*. 62 (2): 101–110.
- 45- "Metoprolol (Oral Route) Precautions". *Drug Information*. Mayo Clinic. Archived from the original on 2009-04-16.
- 46- "Prescribing medicines in pregnancy database". Australian Government. 3 March 2014. Archived from the original on 8 April 2014.
- 47- Page C, Hacket LP, Isbister GK (2009). "The use of high-dose insulin-glucose euglycemia in beta-blocker overdose: a case report". *Journal of Medical Toxicology*. 5 (3): 139–143.
- 48- Albers S, Elshoff JP, Völker C, Richter A, Läer S (2005). "HPLC quantification of metoprolol with solid-phase extraction for the drug monitoring of pediatric patients". *Biomedical Chromatography*. 19 (3): 202–207.
- 49- Kovac AL, Masiongale A (2007) Comparison of Nicardipine versus Esmolol in Attenuating the Hemodynamic Responses to Anesthesia Emergence and Extubation. *J Cardiothorac Vasc Anesth* 21: 45-50.
- 50- Lowrie A, Johnston PL, Fell D, Robinson SL (1992) Cardiovascular and plasma catecholamine responses at tracheal extubation. *Br J Anaesth* 68: 261-263.

51- Kim WY, Lee YS, Ok SJ, Chang MS, Kim JH, Park YC. Lidocaine does not prevent bispectral index increases in response to endotracheal intubation. *Anesth Analg.* 2006;102(1):156–9.

52- Newman D - Overview of lidocaine. *Cardiac Electrophysiology Review.* 2000;4:248-250.

53- Khan IA, Gowda RM - Novel therapeutics for treatment of long-QT syndrome and torsade de pointes. *Int J Cardiol.* 2004;95:1-6.

54- Owczuk R, Wujtewicz MA, Sawicka W, (2008). The effect of intravenous lidocaine on QT changes during tracheal intubation. *Anaesthesia*,63:924-931.

55- Mohammadi,S.S.; Maziar,A. and Saliminia,A.2016. Comparing Clonidine and Lidocaine on Attenuation of Hemodynamic Responses to Laryngoscopy and Tracheal Intubation in Controlled Hypertensive Patients: A Randomized, Double-Blinded Clinical Trial, *Anesth Pain Med.*; 6(2): e34271.

