

SAFETY OF DIAGNOSTIC EVALUATION UNDER ANAESTHESIA WITHOUT ENDOTRACHEAL INTUBATION FOR HEAD NECK SQUAMOUS CELL CARCINOMA PATIENTS- WHEN SHOULD WE AVOID

Rajeev Chugh¹, Devendra Kumar Gupta², Rahul Yadav³, Sanajeet Singh⁴, Sunil Goyal⁵, Ravi Roy⁶, Seema Pati⁷

¹Head and Neck Oncosurgeon, Department of ENT, Army Hospital R & R, Dhaula Kuan, New Delhi, India.

²Head and Neck Oncosurgeon, Department of ENT, Army Hospital R & R, Dhaula Kuan, New Delhi, India.

³Professor, Department of ENT, Army Hospital R & R, Dhaula Kuan, New Delhi, India.

⁴HOD, Department of ENT, Army Hospital R & R, Dhaula Kuan, New Delhi, India.

⁵Classified Specialist ENT, Department of ENT, Army Hospital R & R, Dhaula Kuan, New Delhi, India.

⁶Classified Specialist ENT, Department of ENT, Army Hospital R & R, Dhaula Kuan, New Delhi, India.

⁷Resident, Department of ENT, Army Hospital R & R, Dhaula Kuan, New Delhi, India.

ABSTRACT

BACKGROUND

Airway management is a very important aspect during head and neck squamous cancer (HNSCC) diagnostic procedures under anaesthesia, as the upper airway is shared by the anaesthesiologist as well as the surgeon. Adequate airway management is essential during these procedures due to safety issues and cost effectiveness and also to prevent an airway emergency. There is always a question of which patients need endotracheal intubation and in whom we should avoid. We wanted to assess the merits and demerits of upper airway diagnostic procedures in HNSCC under general anaesthesia with or without endotracheal intubation and determine the factors which may lead to an airway emergency or endotracheal intubation.

METHODS

This is a retrospective observational study of consecutive HNSCC patients, who underwent diagnostic or therapeutic rigid laryngoscopy under anaesthesia between Dec 2015 till Jul 2017 (retrospective) and Aug 2017 till December 2018 (prospectively maintained database) at a tertiary referral centre. Consecutive patients of HNSCC who underwent diagnostic endoscopy under anaesthesia for biopsy or mapping of the disease were included in the study. 393 patients' data was screened, out of which 303 patients were eligible for our study. 90 patients were excluded from the study, 30 were in stridor and 60 had a prior tracheostomy at the time of diagnostic evaluation. Various factors were evaluated for endotracheal intubation.

RESULTS

303 consecutive patients' database was analysed. Male:Female ratio was 93:210 (1:2.3); age ranged from 27 years to 81 years; mean age was 43 years; body mass index ranged from 17-33 with median 20. The site of lesion (glottis) and prior treatment were found to be significant factors for elective intubation. Micro-laryngoscopic procedure was found to be significant for endotracheal intubation. Rest all procedures can be safely performed with total intravenous anaesthesia (TIVA).

CONCLUSIONS

Direct laryngoscopy with or without biopsy for UADT SCC is safe without intubation barring few cases. We can do a DL Scopy next day of visit to our centre in per primum, thereby reducing the waiting period and establishing diagnosis early and hence reducing overall treatment time. It is cost effective. Patients with prior cancer directed treatment need intubation because of altered anatomy and neck tissue fibrosis. ML scopy takes longer time; hence, apnoea technique is not recommended due to CO₂ retention; intermittent intubation through the laryngoscope is acceptable.

KEY WORDS

Head Neck Squamous Cell Cancer, Direct Laryngoscopy, Micro-Laryngoscopy, Total Intravenous Anaesthesia

HOW TO CITE THIS ARTICLE: Chugh R, Gupta DK, Yadav R, et al. Safety of diagnostic evaluation under anaesthesia without endotracheal intubation for head neck squamous cell carcinoma patients- when should we avoid. J. Evolution Med. Dent. Sci. 2019;8(30):2390-2394, DOI: 10.14260/jemds/2019/521

BACKGROUND

Head and neck squamous cell carcinoma is the most common malignancy in Indian male.⁽¹⁾ Purpose of direct laryngoscopy using Jackson or Negus laryngoscope is to assess the exact extent of tumour in terms of involvement of vallecula,

oropharynx, tonsil, post cricoid, oesophagus, arytenoids, interarytenoid mucosa, subglottis and trachea. This assessment is of utmost importance for surgical planning of the patient amongst widely available surgical options from TOLS to open partial laryngectomies to total laryngectomy. This evaluation also is important to decide the type of reconstruction contemplated for that case as lifting the larynx off post pharyngeal wall will exactly reveal the mucosal extent and resection and its reconstruction in terms of primary closure, patch pharyngoplasty or a tubed reconstruction.

Subglottic extension and arytenoid involvement are very important in terms of partial laryngectomy. Extension in to trachea may require more extensive resection of trachea

'Financial or Other Competing Interest': None.

Submission 20-06-2019, Peer Review 15-07-2019,

Acceptance 17-07-2019, Published 29-07-2019.

Corresponding Author:

Dr. Devendra Kumar Gupta,

Head and Neck Oncosurgeon,

Army Hospital R & R,

Dhaulta Kua, Delhi Cantt, New Delhi-10, India.

E-mail: docdk2000@yahoo.com

DOI: 10.14260/jemds/2019/521



which may not be possible in certain cases transcervically.⁽²⁾ (Figure 1, 2, 3)

Airway management is a very important aspect of head neck cancer evaluation as the upper airway is shared by the anaesthesiologist as well as the surgeon. The optimum outcome of the procedure entails a perfect understanding as well as discussion of limitations between the surgeon and the anaesthesiologist for each case in question. Moreover, otorhinolaryngologist or head and neck surgeon is the first point of contact for the patient, hence expected post-operative outcome need to be explained to the patient. We analysed over 300 patients who underwent general anaesthesia for head and neck diagnostic procedures.

The usual protocol for the diagnostic or short therapeutic direct laryngoscopy/ micro-laryngoscopy necessitate securing airway in terms of endotracheal or nasotracheal intubation.⁽³⁾ However, this practice has its demerits also as the total duration of the procedure gets prolonged leading to an overall treatment time prolonged due to waiting list especially in a high volume centre for establishing a diagnosis as well as treatment planning. Endotracheal tube in situ also obscures the common field and hence compromises mapping especially, posterior part of laryngopharynx which need to be assessed in terms of laryngeal preservation procedures. The other technique is to carry out these procedures in short apnoea which involves brief period of apnoea supplemented with continuous oxygen insufflation, without endotracheal intubation. The advantage is that it takes less time and becomes a day care procedure, thus reducing costs. But it may not be considered absolutely safe as airway is not secured during the procedure and time available to the surgeon is also limited.^(4,5,6) A very good understanding between the surgeon and anaesthesiologist is a must for this technique. We present our experience in 303 such patients and the methodology adopted for the same.

METHODS

IEC clearance was obtained to retrospectively evaluate consecutive HNSCC (Head neck squamous cell carcinoma) patients, who underwent diagnostic or therapeutic rigid laryngoscopy under anaesthesia between Dec 2015 till Jul 2017 (retrospective) and Aug 2017 till December 2018 (Prospectively maintained database) at a tertiary hospital

Aims and Objectives

1. To assess the merits and demerits of upper airway diagnostic procedures in HNSCC Head and Neck Squamous cell carcinoma under general anaesthesia with or without endotracheal intubation.
2. To derive factors which may lead to an airway emergency or endotracheal intubation.

Participants

Consecutive patients of HNSCC who underwent diagnostic endoscopy under anaesthesia for biopsy or mapping of the disease. 393 patients' data was screened out of which 303 patients were eligible for our study. 90 patients were excluded from the study, 30 were in stridor and 60 had a prior tracheotomy at the time of diagnostic evaluation.

Exposure

We retrospectively analysed the prospectively maintained database of patients of upper aero-digestive tract malignancy who underwent diagnostic or therapeutic direct laryngoscopy, micro-laryngoscopy or evaluation under anaesthesia with or without biopsy from Dec 2015 till Jul 2017 (Retrospective) and Aug 2017 till December 2018 (Prospectively maintained database) at a tertiary care head and neck cancer treatment center. Patients with prior tracheostomy and those in stridor were excluded from the study.

There were two sets of patients, first who underwent above diagnostic procedures with a standard endotracheal intubation and second group had patients without endotracheal intubation. These patients were subjected to Apnoea technique after induction with total intravenous anaesthesia. After supine positioning and attachment of all standard monitors (ECG, NIBP, EtCO₂, Pulse Oximeter and Temperature), Patients were administered 100% oxygen (For pre-oxygenation) and intravenous propofol 1.5 - 2.5 mg/kg was administered as intravenous induction agent. After ascertaining adequate bag and mask ventilation, succinylcholine (Depolarising muscle relaxant) 2 mg/kg intravenous was administered and patient was handed over to surgeon for direct laryngoscopy/ microlaryngoscopy. During the procedure nasal prongs were used to deliver high flow oxygen @ 10-15 litres/minute as continuous insufflation. The procedure took approximately 3-5 minutes.

Various preoperative and intraoperative factors were analysed which might affect the difficulty of direct laryngoscopy or suspension laryngoscopy as well as requirement of intubation during diagnostic or therapeutic endoscopy of larynx, hypopharynx, oropharynx and oral cavity. The factors analysed were site of lesion (Oral tongue, tonsil, base of tongue, supraglottis, glottis, subglottis, marginal zone, pyriform sinus, post cricoid), per primum or post cancer directed treatment (Post radiotherapy, chemoradiotherapy, post-surgery or post-surgery + adjuvant treatment), various anaesthesia and laryngoscopy indices (Mallampati, Cormack Lehane, thyromental distance, 3:2:2 ratio), type of procedure (Direct laryngoscopy with or without biopsy, micro-laryngoscopy with or without therapeutic procedure, evaluation under anaesthesia). (Table 1)

Statistical Analysis

The factors evaluated were analysed using SPSS version 20 and cross tabulation were run to find correlation between factors and intubation.

RESULTS

Consecutive patients of HNSCC who underwent diagnostic endoscopy under anaesthesia for biopsy or mapping of the disease. 393 patients' data was screened out of which 303 patients were eligible for our study. 90 patients were excluded from the study, 30 were in stridor and 60 had a prior tracheotomy at the time of diagnostic evaluation.

Intubation Required	No – 254 (83.8%)	Yes – 49 (16.2%)	FOB assisted intubation – 10 (20.4% of total intubation n-49) all were post cancer directed treatment					
Fibre Optic assessment performed	No- 203 (66.3%)	Yes – 100 (32.2%)						
Mallampati Class	I-109(36%)	II- 100(33%)	III- 48 (15.8%)	IV- 46 (15.2%)				
Thyro- mental distance (cm)	>6.5 – 101(33.3%)	6- 6.5- 141 (46.5%)	< 6 – 61 (20.1)					
Cormack Lahane Score	86 (28.4%)	2 - 101 (33.3%)	3- 58 (19.1%)	4- 58 (19.1%)				
Tracheostomy	Anticipated-56 (18.5%)	Not anticipated- 247 (81.6%)	Performed – 9 (16% of anticipated n-56)	Not required& not anticipated				
Cord fixity	Yes- 77 (25.4%)	No- 226 (74.6%)						
Prior Treatment	Treatment naïve 225(74.3%)	Post RT 36(11.9%)	Post Concurrent Chemo RT 12 (4.0%)	Post Major head neck Surgery 18 (5.9%)	Post-Surgery + CCRT 12 (4%)			
Site/ Procedure	Oral Tongue/EUA 26 (8.6%)	Base of Tongue/ EUA + DLS 59 (19.5%)	Pyriiform Sinus/ DLS 93 (30.7%)	Supraglottis/ DLS 50 (16.5%)	Post Cricoid/DLS 17 (5.6%)	Post Pharyngeal wall/ DLS 8 (2.6%)	Glottis/MLS 34 (11.2%)	Tonsil/EUA + DLS 16 (5.3%)
Procedure	Direct Laryngoscopy and biopsy 187(61.8%)	Direct Laryngoscopy 30 (9.9%)	Microlaryngoscopy and Biopsy 28 (9.2%)	Microlaryngoscopy and LASER excision 15 (4.9%)	Evaluation under anaesthesia + DL Scopy with or without biopsy 43 (14.2%)			

Table 1. Factors Evaluated

Factors	Intubation Required No.(Percent)	Intubation Not Required No. (Percent)	Percent of intubation required	P Value (95% CI)
Site				
Oral Tongue (n-25)	1	24	4	P = 0.13 (0.2% to 1.8%)
Base Of Tongue (n-63)	6	57	9.5	P = 0.17 (0.3% to 1.7%)
Tonsil (n-16)	0	16	0	P = 0.19 (0.6% to 1.5%)
Pyriiform Sinus (n-94)	7	87	7.4	P = 0.20 (0.7% to 2.8%)
Post Cricoid (n-17)	0	17	0	P = 0.33 (0.5% to 3.8%)
Post Pharyngeal wall (n-9)	0	9	0	P = 0.16 (0.8% to 1.8%)
Supraglottis (n-49)	7	42	14.3	P = 0.43 (0.7% to 2.9%)
Glottis(n-32)	28	4	87.5	P = 0.03 (0.03% to 1.8%)
Prior Treatment				
Treatment Naive(n-211)	28	183	13.3	P = 0.66 (0.7% to 2.8%)
Post Radiotherapy (n-42)	9	33	21.4	P = 0.05 (0.3% to 1.9%)
Post Concurrent Chemo-RT (n-15)	5	10	33.3	P = 0.05 (0.3% to 1.9%)
Post Major Head Neck Surgery (n-20)	2	18	10	P = 0.05 (0.3% to 1.9%)
Post-Surgery + Adjuvant RT/ CCRT (n-15)	5	10	33.3	P = 0.05 (0.3% to 1.9%)
Procedure				
Direct Laryngoscopy with biopsy(n-187)	9	178	4.8	P = 0.45 (0.7% to 2.9%)
Direct Laryngoscopy without biopsy(n-30)	2	28	6.7	P = 0.43 (0.7% to 2.9%)
Microlaryngoscopy (n-28)	21	7	75	P = 0.001 (0.01% to 0.9%)
Microlaryngoscopy and TOLS (n-15)	15	0	100	P = 0.001 (0.01% to 0.9%)
Evaluation under anaesthesia without direct laryngoscopy(n-43)	2	41	4.7	P = 0.41 (0.6% to 1.9%)
Laryngoscopy Indices				
Mallampati I(n-106)	16	90	15.1	P = 0.43 (0.7% to 2.9%)
Mallampati II (n-103)	15 (4.9)	88 (29)	14.6	P = 0.40 (0.8% to 2.6%)
Mallampati III (n-49)	8 ((2.6)	41 (13.5)	16.3	P = 0.39 (0.5% to 1.7%)
Mallampati IV (n-45)	10 (3.3)	35 (11.5)	22.2	P = 0.35 (0.6% to 2.9%)
Thyromental Distance >6.5 cm (n-101)	20 (6.6)	81 (26.7)	19.8	P = 0.28 (0.2% to 1.5%)
Thyromental Distance 6-6.5 cm (n-141)	19 (6.2)	122 (40.2)	13.5	P = 0.17 (0.9% to 1.9%)
Thyromental Distance < 6 cm (n-61)	10 (3.3)	51 (16.8)	16.4	P = 0.40 (0.4% to 2.1%)
Cormack Lehane I (n-87)	11 (3.6)	76 (25.0)	12.6	P = 0.31 (0.6% to 1.0%)
Cormack Lehane II (n-101)	16 (5.2)	85 (28.0)	15.8	P = 0.40 (0.8% to 1.9%)
Cormack Lehane III (n-58)	10 (3.3)	48 (15.8)	17.2	P = 0.43 (0.5% to 3.9%)
Cormack Lehane IV (n-57)	12 (3.9)	45 (14.8)	21.1	P = 0.23 (0.6% to 1.8%)

Table 2. Intubation Requirement Versus Various Factors

Prior Treatment	Intubation Required	Intubation Not Required	Odds Ratio 1.93
Yes	21	71	
No	28	183	

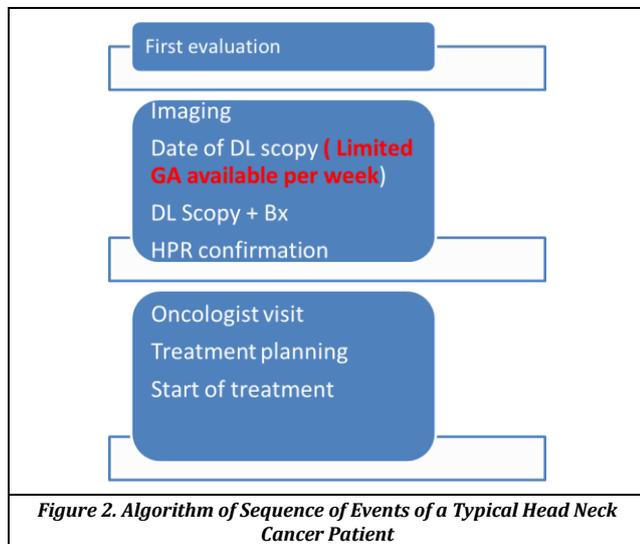
Table 3. Odds Ratio of Prior Treatment for Intubation

	Intubation Required	Intubation Not Required	Total	OR- 97.7
ML Scopy with or without LASER	36	7	43	
Non-MLS procedure	13	247	260	

Table 4. Odds Ratio of Microlaryngoscopy for Intubation



Figure 1. Typical Picture of Office Laryngoscopy in Laryngopharyngeal Cancer Obscuring the Airway and Alarming the Surgeon and Anaesthetist



Oropharynx	Average diameter- 39-51mm	ID- 7-8.5mm	OD- 9.6-11.6 mm	% obscuring - 20-25%
At arytenoids	ETT overlies arytenoids	ID- 7-8.5mm	OD- 9.6-11.6 mm	Complete obscuring the arytenoids
At vocal cords	12-25mm	ID- 7-8.5mm	OD- 9.6-11.6 mm	> 50 % obscuring the post part of vocal cords

Figure 3. Relative Diameter of Endotracheal Tube Obscuring Upper Aerodigestive Tract for Diagnostic Direct Laryngoscopy

213 were male and 90 were female. Body mass index ranged from 18- 36 Kg/m² with a median of 22.5. 254 (83.8 %) did not require endotracheal intubation, however, 49 (16.2%) required intubation. Out of 49 required intubation 10 required a fibreoptic assisted awake intubation in view of difficult airway, all of these were post cancer directed treatment for head neck cancer. Fibreoptic assessment was performed in 203 (66.3%) patients. Tracheostomy was anticipated in 56 (18.5%) and only 9 (3%) required tracheostomy. None of the patients whom tracheostomy was not anticipated required tracheostomy. Majority of the patients were treatment naive 225 (74.3%), 36 (11.9%) were post radiotherapy, rest were post-surgery with or without chemoradiotherapy or concurrent chemoradiotherapy alone. Most of the patients were having a lesion either in hypopharynx or in larynx (Pyriform sinus- 93 (30.7%), supraglottis- 50 (16.5%), Glottis- 34(11.2%), Base of tongue- 59 (19.5%).

Majority of patients underwent direct laryngoscopy with or without biopsy - 227(71.7%), 43 (14.2%) underwent evaluation under anaesthesia for carcinoma tongue for posterior extent along with direct laryngoscopy. 43 (14.1%) underwent microlaryngoscopy with or without LASER resection of lesion in glottis. They were uniformly distributed in various anaesthesia indices like Mallampati, Cormack Lahane and thyromental distances. (Table 2)

On univariate analysis using SPSS version 20, site of lesion (Glottis), prior cancer directed treatment and type of

procedure (Microlaryngoscopy) correlated significantly with requirement of intubation. However, on multivariate analysis microlaryngoscopy and prior treatment were the only factors which significantly affected the decision of intubation amongst this subset of patients. (Table 2)

The Odds ratio (HR) of prior treatment and microlaryngoscopy for requirement of intubation were 1.93 and 97 respectively.

DISCUSSION

Various indices which have been described may serve as important tools to preoperatively predict the airway and accordingly prepare the operative room for difficult intubation or surgical airway. The same may also help in better counseling of the patients. These tools are simple to comprehend and measure.^(7,8) All head neck surgeons should be well versed with their use to predict difficulty of direct laryngoscopy, as well as predict the exposure at microlaryngoscopy. However, for the requirement of intubation these indices did not affect significantly.

Altered anatomy is the most common factor for difficult intubation^(9,10) and was significant factor in our study also. Prior cancer directed treatment for the recurrent head neck cancer patients required intubation because of altered anatomy and neck fibrosis which significantly affected the requirement of intubation in our study (P=0.05 95 % confidence interval 0.3% to 1.9%). Since this was a retrospective review at a single centre hence there are chances of selection bias, however treatment naïve patients can be safely taken up for diagnostic laryngoscopy without intubation.

Site of Lesion

Preoperative indirect laryngoscopy and imaging should be performed to know the site, size and extent of the lesion. It should be performed ideally by the surgeon who is going to perform diagnostic or therapeutic laryngoscopy. This indirect laryngoscopy provides the operating surgeon with the necessary information to plan for any special preparation, if required. However, other than glottis none of the other sites significantly affected the requirement of intubation-

- 1. Base of Tongue-** Lesions on base of tongue will obscure the view of airway below and hence an apprehension to lose airway by the anaesthesiologist. If indirect laryngoscopy is done for intubation, there is a possibility of torrential bleed or the friable tumour may dislodge in the airway leading to airway emergency; hence a rigid direct laryngoscope (Kleinesaucer type) by the head neck surgeon/ ENT surgeon who has evaluated the lesion prior will have lesser chances of airway emergency.
- 2. Tonsil-** Tonsillar lesions can be biopsied easily in an OPD setting under local anaesthesia. However, trans-oral resection with or without Robot/LASER has emerged as a major breakthrough in oropharyngeal malignancies and evaluation under anaesthesia to know the exact extent and depth is essential before undertaking the procedure. This is a short procedure and endotracheal intubation may not be necessary, as it might prolong the overall procedure time.
- 3. Supraglottis/Glottis-** These lesions will obscure the view of vocal cords and lower airway making intubation

difficult. Secondly, endotracheal tube will obscure the mapping of disease; hence evaluation by the surgeon without intubation is preferred. If intubation becomes necessary, the same can be done through direct laryngoscope after the mapping.

4. **Marginal Zone/Pyriform Sinus-** These are lateral lesions and usually do not obscure the airway. Very large or pedunculated lesions may cause ball valve type obstruction of the airway. In such situation, direct laryngoscopy with removal of the part of the tumour which is overlying the cords is attempted to secure the airway. Intubation again may lead to dislodgement of tumour in the lower airway causing airway emergency.

Apnoea technique with high flow oxygen through nasal prongs- After pre-oxygenation and induction with Propofol and Succinylcholine; direct laryngoscopy was performed with Kleinssaucers direct laryngoscope to map the mucosal extent of disease and to obtain biopsy. Average time taken for each procedure was 150 seconds with a range of 75 seconds to 300 seconds.

Micro-laryngoscopy with or without LASER- suspension laryngoscopy done using direct laryngoscope and microscope at 400 mm focal length. Suspension laryngoscopy with microscope adjustment with or without LASER takes few minutes especially in a post graduate teaching institute, keeping residents training in consideration. Hence all Micro-laryngoscopic procedures were undertaken using 5 to 5.5 size microlaryngoscopy tube. In a few cases where the lesion was extending up to post commissure or inter-arytenoid region, microlaryngoscopy was initiated without securing the airway; with intermittent endotracheal intubation through the suspension laryngoscope which was kept in situ.

Evaluation under Anaesthesia was performed like DL Scopy, and usually the two were combined to assess the lower extent of disease.

CONCLUSIONS

TIVA is safe and provides excellent view of the airway. Patient turn-over in high volume centres is very high & hence cost effective. However, following points should be kept in mind-

1. Preoperative assessment of airway by the evaluating surgeon especially in post cancer directed treatment cases should be done.
2. Pre-procedure counseling of patient and attendants about chance of airway emergency and tracheostomy if deemed necessary, should be done.

3. Fibre optic flexible laryngoscopy along with difficult intubation cart needs to be kept standby in the operating room.
4. Microlaryngoscopy & biopsies and definitive treatment requires longer duration and hence prior or intermittent endotracheal intubation is required.

REFERENCES

- [1] <https://www.uicc.org/news/new-global-cancer-data-globocan-2018>.
- [2] Priest RE, Wesolowski S. Direct laryngoscopy under general anaesthesia. *Trans Am Acad Ophthalmol Otolaryngol* 1960;64:639-48.
- [3] Woodman D. Laryngoscopy under general anaesthesia: apnoeic oxygenation technique. A report of over 100 cases. *Ann Otol Rhinol Laryngol* 1961;70:1113-6.
- [4] Hogue CW Jr, Bowdle TA, O'Leary C, et al. A multicenter evaluation of total intravenous anaesthesia with remifentanyl and propofol for elective inpatient surgery. *Anaesth Analg* 1996;83(2):279-85.
- [5] Succo G, Bussi M, Presutti L, et al. Supratracheal laryngectomy: current indications and contraindications. *Acta Otorhinolaryngol Ital* 2015;35(3):146-56.
- [6] Armstrong M, Mark LJ, Snyder DS, et al. Safety of direct laryngoscopy as an outpatient procedure. *Laryngoscope* 1997;107(8):1060-5.
- [7] Orosco RK, Lin HW, Bhattacharyya N. Safety of adult ambulatory direct laryngoscopy: revisits and complications. *JAMA Otolaryngol Head Neck Surg* 2015;141(8):685-9.
- [8] Paul RR, Varghese AM, Mathew J, et al. Difficult laryngeal exposure in microlaryngoscopy: can it be predicted preoperatively? *Indian J Otolaryngol Head Neck Surg* 2016;68(1):65-70.
- [9] Umesh G, Ellango A, Jasvinder K, et al. Immediate postoperative airway obstruction secondary to airway edema following tumor excision from the neck. *J Anaesth* 2009;23(4):609-12.
- [10] Ebert TJ, Muzi M, Berens R, et al. Sympathetic responses to induction of anaesthesia in humans with propofol or etomidate. *Anaesthesiology* 1992;76(5):725-33.