Clinical guidelines on intraoperative neuromonitoring during thyroid and parathyroid surgery

Hui Sun^{1*}, Wen Tian^{2*}, Kewei Jiang³, Fengyu Chiang⁴, Ping Wang⁵, Tao Huang⁶, Jingqiang Zhu⁷, Jianwu Qin⁸, Xiaoli Liu¹

¹China-Japan Union Hospital of Jilin University, Changchun 130033, China; ²General Hospital of the People's Liberation Army (PLA), Beijing 100853, China; ³People's Hospital of Peking University, Beijing 100044, China; ⁴Institute of Clinical Medicine, Kaohsiung Medical University, Kaohsiung, Taiwan; ⁵Second Affiliated Hospital of Medical College of Zhejiang University, Hangzhou 310009, China; ⁶Union Hospital Affiliated to Huazhong University of Science and Technology, Wuhan 430074, China; ⁷West China Hospital, Sichuan University, Chengdu 610041, China; ⁸Affiliated Cancer Hospital of Zhengzhou University, Zhengzhou 450008, China

*These authors contributed equally to this work.

Correspondence to: Hui Sun, MD. China-Japan Union Hospital of Jilin University, Changchun 130033, China. Email: sunhui1229@163.com; Wen Tian, MD. General Hospital of the People's Liberation Army (PLA), Beijing 100853, China. Email: tianwen301_cta01@163.com.

Submitted Jul 03, 2015. Accepted for publication Aug 12, 2015. doi: 10.3978/j.issn.2305-5839.2015.08.21 View this article at: http://dx.doi.org/10.3978/j.issn.2305-5839.2015.08.21

Introduction

Recurrent laryngeal nerve (RLN) injury is one of the most severe complications of thyroid surgery. Hoarseness due to unilateral RLN injury and breathing disorders and even asphyxia due to bilateral RLN injury can impose serious impacts on the patient's life. It is estimated that the incidence of RLN injury during thyroid surgeries ranges 0.3-18.9% (1-5), making RLN protection a great concern among thyroid surgeons. Intraoperative neuromonitoring (IONM) combines both the functional and anatomic techniques and has the following features: intra-operative navigation and rapid identification of the RLN distribution; predicting nerve variation and protecting of the functional integrity of RLN; clarifying the mechanism and lowering the incidence of RLN injury; and easy to perform. It is a helpful adjunct for complicated surgeries (6,7).

With an attempt to help the surgeons thoroughly understand IONM and carry out this technique in a standardized and reasonable manner, the Chinese Thyroid Association established the Clinical Guidelines on IONM during Thyroid and Parathyroid Surgeries (China Edition).

Basic principles of IONM

Application of IONM in thyroid surgeries was initially proposed by Shedd in 1966 and by Flisberg in 1970. Based on the principle of electrophysiology, the motor nerve is stimulated by electricity during the surgery, and then the nerve impulses are formed and transferred to the dominant muscles to produce myoelectric signals, forming waves and alerts in electromyography (EMG), which will help the surgeons to judge the functional integrity of nerves (8,9) (*Figure 1*).

Clinical significance and technical advantages of IONM

For doctors who are transforming from generalist to specialist, junior doctors, and surgeons who are facing a complicated thyroid surgery, IONM undoubtedly is a good aid (10-12).

Facilitates the identification and locating of RLN

Before RLN exposure, the nerve can be accurately located at its distribution area using the cross method, which enables the doctor to rapidly determine the anatomic range of RLN as well as its rare anatomic variations such as nonrecurrent RLN (13).

Facilitates the exposure and dissection of RLN

During the dissection of RLN, continuous monitoring together with naked-eye observation enables the

Page 2 of 9

differentiation between the monitored nerve and its surrounding non-nerve tissues and the accurate tracing of the nerve and its functional branches. Also, the intraoperative navigation is also helpful for the complete resection of lesions.

Facilitates the judgment of the functional integrity and injury mechanisms of nerves

Electrophysiological monitoring provides quantitative indicators for judging the functional integrity of nerves, can accurately locate the "injury point" of a nerve (14) (*Figure 2*), and help the operator to analyze the injury mechanism and timely recognize and resolve the injury during surgery. Thus, it can markedly reduce the incidence of RLN injury.

Indications of IONM (15-18)

IONM is a preferred option in the following patients, while its application in the other patients should be upon the doctor's recommendation: (I) the mass is located at the dorsal side of the thyroid, with suspected recent capsular hemorrhage or thyroid cancer; (II) in patients with hyperthyroidism, pre-operative ultrasound indicates large gland and rich blood supply; (III) in patients with malignant thyroid tumor, neck lymph node dissection is required, especially in patients with swollen central lymph nodes; (IV) patients receiving a second thyroid surgery but with disordered anatomic structures and severe adhesions; (V) patients with retrosternal goiter and/or large thyroid mass, along with suspected RLN dislocation; (VI) pre-operative imaging indicated the presence of transposition of viscera or subclavian artery variation, along with suspected nonrecurrent RLN; (VII) patients with unilateral vocal cord paralysis, with the contralateral lobe requiring surgical treatment; (IX) patients requiring total thyroidectomy, in particular an endoscopic surgery; (X) patients requiring a surgery to repair RLN injury; (XI) patients requiring a parathyroid surgery; and (XII) patients with special requirements on sound and tone and thus requiring IONM.

Notably, (I) if intra-operative exploration shows that the thyroid cancer has completely infiltrated the RLN, nerve preservation will unavoidably result in residual tumor; thus, the invaded nerve must be resected to thoroughly remove the tumor. Under such circumstances, the post-operative hoarseness can not be avoided even after the application of IONM; (II) in patients with pre-operative vocal palsy, IONM

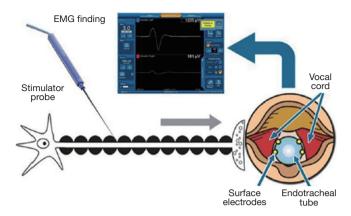


Figure 1 Basic principles of IONM. IONM, intraoperative neuromonitoring.

can be applied to search for the injury site, and nerve repair can be completed with the assistance of IONM; however, it is difficult to completely restore the nerve function.

Since it is impossible to predict all the complex cases before a thyroid or parathyroid surgery, the indications of IONM may be widened if condition allows. The same recommendation has also proposed in international literature: since it is difficult to predict RLN variation before surgery, IONM may be routinely applied if condition allows.

Essential monitoring devices of IONM

The IONM devices can be divided into recording side (recording electrode and its ground electrode) and stimulation side (stimulator probe and its loop electrodes) as well as EMG monitor, interface-connector box, anti-jamming silence detector, and patient simulators (19) (*Figure 3*).

Needle-like electrode and intubation surface electrode are two typical recording electrodes, with the latter being routinely recommended (20).

The stimulation probes can be divided into monopolar and bipolar types, with the monopolar Prass probe with ball tip is routinely recommended (21).

Standardized procedures of IONM

Non-standard application of IONM techniques will cause significant monitoring errors. The main aim of standardized IONM procedures is to guide and improve the quality of IONM techniques and avoid any adverse effect due to improper monitoring operations (22-24). The standard procedure of IONM is shown in *Table 1*.

Annals of Translational Medicine, Vol 3, No 15 September 2015

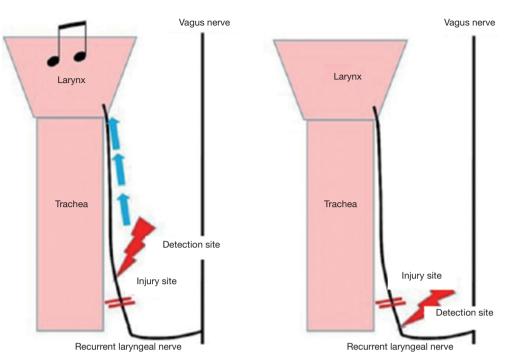


Figure 2 Locating nerve "injury site" by IONM. IONM, intraoperative neuromonitoring.

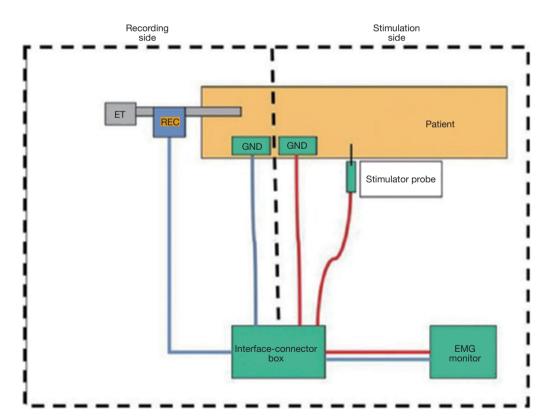


Figure 3 Basic monitoring equipment setup. ET, endotracheal tube; REC, recording electrodes; GND, ground electrodes; EMG, electromyography.

Page 3 of 9

Page 4 of 9

Sun et al. Intraoperative neuromonitoring during thyroid and parathyroid surgery

Procedures	Note
Record vocal cord movement before surgery	Use of fiberoptic laryngoscope
Recommended anesthesia method for IONM	Intermediate- or short-acting muscle relaxant is recommended for preoperative anesthesia induction. The dosage of intermediate-acting muscle relaxant is smaller than that used for routine anesthesia induction. 1x ED95 intermediate-acting non-depolarizing muscle relaxant is recommended; avoid adding muscle relaxant before the termination of intraoperative monitoring
Body position and tube placement	Anesthesia intubation is performed after positioning (intubation under video laryngoscope is recommended)
Device connections and checking	
The ground electrodes are routinely placed	
under the skin at the shoulders or xiphoid	
Confirm the monitoring system has been effectively established	
Check electrode impedance and differences in impedance values	Electrode impedance <5 k $\Omega,$ with deviations <1 k ω
Check initial EMG	Initial fluctuations: about 10 μV
Set up event thresholds The current intensity of stimulator probe	Typically 100 μV
should be routinely set at 1-3 mA	
The monitoring device should be placed	
far away from electro-surgical devices and be connected with anti-jamming silence detectors	
The recording electrode positions should be confirmed during surgery	Directly detect, locate, and record the electrode depth at the antemedial laryngeal line using stimulator probe
IONM four-step method	
Step 1: V1 signal	Obvious bipolar EMG signal is obtained at the ipsolateral vagus nerve at the plexas thyreoidea inferior level (point B), confirming the successful establishment of the monitoring system. If signal is absent at point B, detect the vagus nerve at the plexas thyroid superior level (point A); the presence of signal at point A confirms the presence of non-recurrent laryngeal nerve (25) (<i>Figure 4</i>)
Step 2: R1 signal	Before the exposure of RLN, its EMG signal is located using the Cross method by applying the probe vertical to trachea at its traveling area and then parallel to trachea
Step 3: R2 signal	Continuous monitoring is applied during the dissection of RLN and the signal change is compared in a real-time manner. After the RLN is exposed, the most proximal end of the exposed part is detected for EMG signal
Step 4: V2 signal	After complete hemostasis is achieved at the surgical field, the EMG signal of the vagus nerve is detected before closing the incision
Signal analysis	-
Basic EMG parameters	The biphasic waveform should be differentiated from the monophasic non-EMG artifacts

Table 1 The standard procedures of IONM

Annals of Translational Medicine, Vol 3, No 15 September 2015

Table 1 (continued)

Procedures	Note
No obvious decrease in R2 and V2 signals	The basic EMG parameters include amplitude, latency and duration (26) (<i>Figure 5</i>); the RLN has intact function
Loss of R2 and V2 signals	If the RLN is injured during the surgical operation, detect the "injury site" and search for the injury cause*
Photo recording the exposed RLN during	To confirm the RLN continuity (visual integrity)
surgery	
Postoperative laryngoscopy	

*, If no "injury site" is detected, it is important to determine whether there is a "real" loss of signal. (I) stimulate the nerve and then observe the contact between endotracheal tube electrode and vocal cord via a laryngoscope; (II) detect signals at vagus nerve and RLN again before closing the incision. IONM, intraoperative neuromonitoring; RLN, recurrent laryngeal nerve.

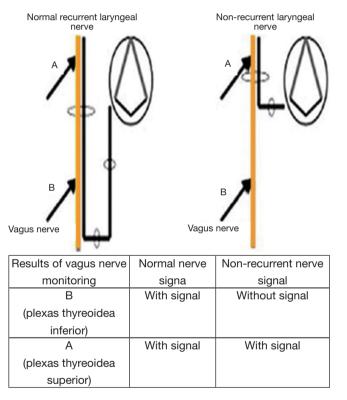


Figure 4 Monitoring setup of non-recurrent laryngeal nerve.

Causes and solutions of common IONM errors

The incidence of IONM error ranges 3.8-23.0% (27). IONM errors can bring great psychological pressure to the operator, delay the surgical process, and even cause the operator make wrong decision. Therefore, it is important to master the causes and solutions of common IONM errors, so as to ensure the surgery is performed in a safe and smooth

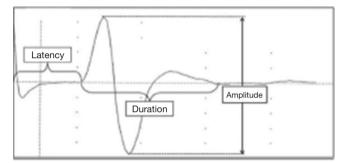


Figure 5 Basic parameters of the EMG wave of IONM. IONM, intraoperative neuromonitoring.

manner. *Table 2* lists the causes and solutions of common IONM errors, and the troubleshooting process (28) is displayed in *Figure 6*.

The standard application of IONM can help 85% of beginners to smoothly locate the laryngeal nerves, provide quantitative indicators of nerve function for surgeons, assist the surgeons to deal with the complicate anatomic structures and skillfully keep away from the dangerous areas. Also, by using both eyes and ears (by listening to the alerts), the operators can accurately resect the tumor. Thus, IONM has become an effect adjunct for the golden standard of nakedeve protection. With "simple, effective, and practicable" as the basic principles, the Clinical Guidelines on IONM during Thyroid and Parathyroid Surgeries (China Edition) established by Chinese Thyroid Association elucidates the standardized operation procedures and decision-making steps of IONM. Before the application of IONM, the surgeons need to fully know neuromonitoring systems and thoroughly understand the Guidelines and receive corresponding training and verification in a standardized monitoring base.

Table 2 Causes and solutions of common IONM	ONM errors	
Common IONM errors	Causes	Solutions
Electrode impedance is too high	The subcutaneous electrodes have not completely removed after its withdrawal from patients	Check whether the subcutaneous electrodes fall off and keep the electrode clean
Subcutaneous electrodes >10	The impedance of the electrode itself is too high	Replace the electrodes and fix them with tape
Stimulator probe electrodes >25 kΩ	The electrode core does not connect well with the interface-connector box The interface-connector box does not connect well with the monitor	Check the connections of the interface-connector box
Recording electrodes	The recording electrode does not connect well with vocal cord	Adjust the tube depth and angle under fiberoptic laryngoscope
Single electrode impedance >5 k Ω	The surface electrode of endotracheal tube is displaced	Indwell the tube under conventional video laryngoscope
Impedance deviation>1 kΩ	Application of insulating lubricant before intubation	Avoid the application of insulating medium at the recording electrodes
Electrode impedance is zero	Two subcutaneous electrodes contact with each other	Re-indwell the subcutaneous electrodes, with the inter-electrode distance of >1 cm
Electrosurgical interference	The probe of anti-jamming detector is not connected	Circle the cable of the electrosurgical device, with the anti-jamming detector clipped on the twisted cable
After the establishment of standard monitoring system	Preoperative vocal cord palsy	Re-check the preoperative laryngoscopic records
Before thyroid surgery	The nerve detected by the operator is actually not a vagus nerve	Detect at 1 mA after confirming the exposure of vagus nerve
V1 signal is absent	The vagus nerve is injured during its exposure non-recurrent laryngeal nerve is present	Directly detect the carotid sheath at 3 mA to obtain the V1 signal If the signal of vagus nerve is absent at the plexas thyreoidea inferior level, re-check it at the plexas thyreoidea superior level
	Anesthesia induction is not performed as recommended	Wail until the muscle relaxant wears off or use a proper dose of a muscle relaxant antagonist
	Improper type or dosage of muscle relaxant The detection current is not high enough	Check the matching between the measured stimulation (as displayed on monitor) and the stimulation setup value Re-check the connections between the electrodes and the interface-
		connector box Check whether the fuse in the interface-connector box has been burned out

Table 2 (continued)

Table 2 (continued)		
Common IONM errors	Causes	Solutions
	The frequency of stimulus pulse is too low	Stimulus pulse frequency: 4 times/s by default
	The setup of event threshold is too high	Routinely 100 µV; avoid changing this parameter value casually
	The selected monitoring mode, channel, and	Re-check the monitoring mode, channel, and volume setup
	volume are improper	Each detection should be maintained at least 1s
	The duration of detection for nerve is too short Avoid reuse	Avoid reuse
	The probe is damaged, with insulation layer	Clear liquids at the detection area
	falling off	Re-check whether the electrodes are off
	The shunt of the nerve detection area is too	The depth of the surface electrodes of endotracheal tube can be
	large	detected and located at the laryngeal anteromedian line

RLN, recurrent laryngeal nerve.

	The selected monitoring mode, channel, and	Re-check the monitoring mode, channel, and volume setup
	volume are improper	Each detection should be maintained at least 1s
	The duration of detection for nerve is too short	Avoid reuse
	The probe is damaged, with insulation layer	Clear liquids at the detection area
	falling off	Re-check whether the electrodes are off
	The shunt of the nerve detection area is too	The depth of the surface electrodes of endotracheal tube can be
	large	detected and located at the laryngeal anteromedian line
	The muscle for detecting neurological effects is	
	detached from the recording electrode	
EMG signal is present while no nerve is	Consecutive "sequence" EMG response cannot	Consecutive "sequence" EMG response cannot Light anesthesia, with spontaneous activity of laryngeal muscle;
detected	be explained	the recording nerve or muscle is tracted by other nerve or muscle
	Artifacts occur in the non-neural traveling area	The surface electrode of endotracheal tube is placed too deeply
	The detection current is too large	Direct detect the nerve trunk (1 mA is recommended)
		Adjust according to the anatomic structures and EMG signals during
		the surgery
V1 signal is good, confirming the	Intraoperative anesthesia or muscle relaxation	Avoid adding muscle relaxant before the termination of monitoring
establishment of standard monitoring	status changes	Check the nerve continuity
system, whereas there is the decrease	Nerve transection injury	Locate the injury site and analyze the possible injury mechanisms:
of signal by >50% or LOS during the	Nerve injury not visible to the naked eye	traction injury, heat injury, suction injury, and/or thread-cutting injury
dissection of RLN	Monitoring system failure	Re-check the electrode connections to ensure a good circuit
	Recording electrode displacement due to	performance
	changes in head position or body position	Use simulators to re-check the monitor and interface-connector box
	during the surgery	(e.g., fuse)
		Re-check the laryngoscope and adjust the endotracheal tube

Annals of Translational Medicine, Vol 3, No 15 September 2015

Sun et al. Intraoperative neuromonitoring during thyroid and parathyroid surgery

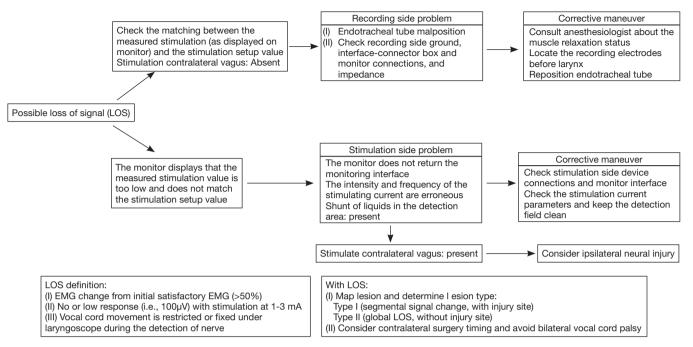


Figure 6 Algorithm for troubleshooting common IONM errors. IONM, intraoperative neuromonitoring.

With the assistance of IONM, an experienced surgeon will be able to further lower the incidence of RLN injury and improve the surgical safety and completeness, which can also be a new trend in laryngeal nerve protection during a thyroid surgery (29).

Acknowledgements

None.

Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

References

- Dralle H, Sekulla C, Haerting J, et al. Risk factors of paralysis and functional outcome after recurrent laryngeal nerve monitoring in thyroid surgery. Surgery 2004;136:1310-22.
- 2. Chiang FY, Wang LF, Huang YF, et al. Recurrent laryngeal nerve palsy after thyroidectomy with routine identification of the recurrent laryngeal nerve. Surgery 2005;137:342-7.
- 3. Chiang FY, Lee KW, Huang YF, et al. Risk of vocal palsy after thyroidecitomy with identification of the recurrent

laryngeal nerve. Kaohsiung J Med Sci 2004;20:431-6.

- 4. Qu XC, Xiao Y, Huang T, et al. Exposure and injury prevention of recurrent laryingeal nerve during thyroid surgery. Chinese Journal of Cancer 2008;18:700-3.
- Liu CP, Huang T. Causes and Treatment of Recurrent Laryngeal Nerve Injury during Thyroidectomy. Chin J Bases Clin General Surg 2008;15:314-7.
- Dionigi G, Barczynski M, Chiang FY, et al. Why monitor the recurrent laryngeal nerve in thyroid surgery? J Endocrinol Invest 2010;33:819-22.
- Liu XL, Sun H, Zheng ZL, et al. Application and advances in the monitoring of recurrent laryngeal nerve during thyroid surgeries. Chinese Journal of general surgery, 2009;18:1187-90.
- Sun H, Liu XL, Lian LX, et al. Principles and application of the monitoring of recurrent laryngeal nerve. News and Reviews (Ear, Nose, and Throat) 2012;27:137-40.
- Zhou G, Jiang KW, Ye YJ, et al. Intraoperative assessment of recurrent laryngeal nerve function by monitoring EMG of lateral cricoaryteoid muscle. Chinese Journal of General Surgery 2012;27:272-5.
- Barczyński M, Konturek A, Cichoń S. Randomized clinical trial of visualization versus neuromonitoring of recurrent laryngeal nerves during thyroidectomy. Br J Surg 2009;96:240-6.
- 11. Dionigi G, Bacuzzi A, Boni L, et al. What is the learning

Annals of Translational Medicine, Vol 3, No 15 September 2015

curve for intraoperative neuromonitoring in thyroid surgery? Int J Surg 2008;6 Suppl 1:S7-12.

- Sun H, Liu XL, Zhang DQ, et al. Clinical application of recurrent laryngeal nerve protection and monitoring during thyroidectomy. Chin J Bases Clin General Surg 2010;17:768-71.
- Chiang FY, Lu IC, Chen HC, et al. Anatomical variations of recurrent laryngeal nerve during thyroid surgery: how to identify and handle the variations with intraoperative neuromonitoring. Kaohsiung J Med Sci 2010;26:575-83.
- Chiang FY, Lu IC, Kuo WR, et al. The mechanism of recurrent laryngeal nerve injury during thyroid surgery-the application of intraoperative neuromonitoring. Surgery 2008;143:743-9.
- Sun H, Liu XL, Fu YT, et al. Application of intraoperative neromonitoring during complex thyroid operation. Chinese Journal of Practical Surgery 2010;30:66-8.
- Qin JW, Hei H, Zhang ST, et al. Locating and protecting recurrent laryngeal nerve in minimally invasive videoassisted thyroidectomy. Cancer Research and Clinic 2010;12:804-5.
- Wei T, Li ZH, Zhu JQ. Real-Time Monitoring of Recurrent Laryngeal Nerve During Thyroid Reoperation. Chin J Bases Clin General Surg 2010;17:772-4.
- Wang P, Yan HC. Prevention and treatment of complications after complete endoscopic thyroidectomy. Journal of Laparoscopic Surgery 2012;17:806-9.
- Randolph GW, Dralle H, International Intraoperative Monitoring Study Group, et al. Electrophysiologic recurrent laryngeal nerve monitoring during thyroid and parathyroid surgery: international standards guideline statement. Laryngoscope 2011;121 Suppl 1:S1-16.
- 20. Liu XL, Sun H. Principles and clinical application of intraoperative monitoring of recurrent laryngeal nerve.

Cite this article as: Sun H, Tian W, Jiang K, Chiang F, Wang P, Huang T, Zhu J, Qin J, Liu X. Clinical guidelines on intraoperative neuromonitoring during thyroid and parathyroid surgery. Ann Transl Med 2015;3(15):213. doi: 10.3978/j.issn.2305-5839.2015.08.21

Chinese Journal of Practical Surgery 2012;32:409-11.

- Sun H, Liu XL, Zhao T, et al. New methods for identifying recurrent laryngeal nerve during thyroid surgery. News and Reviews (Ear, Nose, and Throat) 2010;25:46-8.
- 22. Chiang FY, Lee KW, Chen HC, et al. Standardization of intraoperative neuromonitoring of recurrent laryngeal nerve in thyroid operation. World J Surg 2010;34:223-9.
- 23. Randolph GW, Kamani D. The importance of preoperative laryngoscopy in patients undergoing thyroidectomy: voice, vocal cord function, and the preoperative detection of invasive thyroid malignancy. Surgery 2006;139:357-62.
- Liu XL, Sun H. Optimization and interpretation of intraoperative neuromonitoring of recurrent laryngeal nerve in thyroid operation. News and Reviews (Ear, Nose, and Throat) 2010;25:152-4.
- Sun H, Liu XL, Zhao T, et al. Intraoperative neuromonitoring in identification of non-recurrent laryngeal nerve: experience of 6 cases. Chin J Endocr Surg 2010;4:402-4.
- Dralle H, Sekulla C, Lorenz K, et al. Intraoperative monitoring of the recurrent laryngeal nerve in thyroid surgery. World J Surg 2008;32:1358-66.
- 27. Chan WF, Lo CY. Pitfalls of intraoperative neuromonitoring for predicting postoperative recurrent laryngeal nerve function during thyroidectomy. World J Surg 2006;30:806-12.
- Sun H, Liu XL. The preservation method of the recurrent laryngeal nerve and superior laryngeal nerve in the thyroid surgery. Chinese Journal of Practical Surgery 2012;32:356-9.
- Tian W, Luo J. Comparison of the guidelines on the management of thyroid nodules and differentiated thyroid cancer between China and the United States. Chinese Journal of Practical Surgery 2013;33:475-9.