INTRODUCTION

Recurrent laryngeal nerve (RLN) injury is one of the most important and serious complications of thyroid surgery, with a reported incidence of 0%–11% [1]. Visual identification of the RLN is the gold standard for finding the RLN and avoiding RLN palsy [2,3]. However, even expert thyroid surgeons cannot be entirely free from this complication. Therefore, various types of intraoperative neuromonitoring (IONM) have been developed to facilitate the identification of the RLN and to verify its functional integrity [4-6].

Electrical stimulation to identify the RLN during thyroidectomy was introduced in the late 1960s and has become widely used [7,8]. In the early period, surgeons proposed various IONM methods. Some surgeons palpated the movement of the larynx (especially the posterior hypopharyngeal wall) during thyroidectomy, and some recorded the pressure of an endotracheal tube (ET) cuff that was placed between the vocal cords. Attempts were also made to obtain electromyography (EMG) signals from the laryngeal muscle using needle electrodes or surface electrodes in different positions [9-12]. Initially, needle electrodes were used due to their stability. However, the electromyography (EMG) ET with a surface electrode was developed and released commercially in the 1990s [11,13]. EMG ETs have been widely used for thyroid and parathyroid surgery and have emerged as a standard method for IONM because they are easy to set up [5,8,14]. A standardized process for IONM that systematically identifies signals from the vagus nerve and the RLN in order (V1, R1, R2, V2) was published.
MATERIALS AND METHODS

We prospectively collected EMG data from IONM of 54 nerves at risk (NARs) in 38 patients during thyroidectomy. All surgeries were performed by a single surgeon (YBJ) from July 2017 to December 2018. The indications of IONM were recurrent thyroid node metastasis, extracapsular invasion, and diffuse thyroid goiter included. We decide the indications of IONM according to the recommendation of Korean national medical insurance system.

Vocal cord movement was checked with flexible laryngofiberscopy before and after surgery in all patients. The standard IONM procedure suggested by the International Intraoperative Monitoring Study Group was followed in all surgeries [15]. V1 was defined as an initial EMG signal from the vagus nerve before RLN dissection, and R1 was the initial signal from RLN which was identified at the tracheoesophageal groove. R2 and V2 signals were obtained after thyroidectomy was completed.

All surgeries were performed under general anesthesia with orotracheal intubation. A nondepolarizing muscle relaxant (rocuronium 0.5–0.6 mg/kg) was used only once at the induction of anesthesia, and there was no additional use of muscle relaxants during surgery. Each patient was intubated with an EMG-reinforced ET (Medtronic Xomed, Jacksonville, FL, USA). The size of the EMG tube was chosen to fit as tightly as possible against both sides of the vocal cords. The position of the ET was confirmed routinely after neck extension with a portable flexible laryngofiberscope before the tube fixation. After the placement of the ground electrode, we verified that the impedance value was less than 5 kΩ. A monopolar stimulator probe was used for nerve stimulation.

During the surgery, after the pyramidal lobe was dissected, the thyroid cartilage was exposed. A 22-mm twisted pair of needle electrodes (Medtronic Xomed) was inserted through the perichondrium and thyroid cartilage at the mid-portion of both thyroid alar cartilage, and they were fixed with suture ties (Fig. 1). When the needles were difficult to insert due to calcification of thyroid cartilage, they were inserted into a soft portion in another part of the cartilage or a perichondrium.

Impedance (<5.0 kΩ) and impedance imbalance (<1.0 kΩ) were checked to ensure appropriate mucosal contact. Pulsatile neural stimulation was used at 4 Hz, duration 100 ms at 1 to 2 mA during thyroidectomy with a standard monopolar stimulating probe (Medtronic Xomed). The intensity of stimulation was increased up to 3–5 mA if there was no response at 1–2 mA, especially for the vagus nerve stimulations. The event threshold was preset at 100 μV.

In all cases, EMG signals from both ET and transcartilaginous (TC) electrodes were recorded simultaneously (Fig. 2) to compare two methods in the same patient. We compared the characteristic of EMG signal at each of the four steps of the standard procedure (V1, R1, R2, V2) and efficacy of both techniques. The appropriate event threshold was set to 100 μV. LOS was noted when the largest amplitude was less than 100 μV [15].

RESULTS

The patients’ characteristics and surgical details are summarized in Table 1. Among the 38 patients, there were 25 women and 13
men, with an average age of 53.8 years. In total, 21 patients underwent total thyroidectomy and 17 underwent thyroid lobectomy. The average time from intubation to V1 signal detection was $42.2 \pm 22.6$ minutes. A V1 amplitude $>500 \mu V$ was achieved in 19 NARs (35.2%) with ET electrodes and 47 NARs (87.0%) with TC electrodes ($P<0.001$). Significantly higher mean EMG amplitudes were recorded by TC electrodes than by ET electrodes in all four-step procedures (Table 2).
Table 1. Clinicopathologic characteristics of the patients

| Variable          | Value (n=38) |
|-------------------|-------------|
| Sex               |             |
| Male              | 13 (34.21)  |
| Female            | 25 (65.79)  |
| Age (yr)          | 53.8±11.6   |
| Pathology         |             |
| Papillary carcinoma | 20 (52.63) |
| Follicular carcinoma | 1 (2.63)  |
| Medullary carcinoma | 1 (2.63)  |
| Benign            | 16 (42.11)  |
| Type of surgery   |             |
| Total thyroidectomy | 21 (55.26) |
| Thyroid lobectomy  |             |
| Time interval between V1 and intubation (min) | 42.2±22.6 |

Values are presented as number (%) or mean±standard deviation.

Prediction of postoperative nerve function

LOS occurred in five NARs (9.26%) in the EMG signals measured with ET electrodes, but only two of them showed significant interval changes in the EMG signal measured with TC electrodes. The other three NARs exhibited no significant interval changes in the EMG signal measured with TC electrodes. Postoperative laryngoscopy revealed that the two NARs exhibiting LOS with both the ET and TC electrodes had RLN palsy. In contrast, the three NARs that displayed LOS only with the ET electrodes had intact vocal cord movement. Thus, the sensitivity, specificity, and positive and negative predictive values of LOS for the EMG signals from the ET electrodes were 100%, 94%, 40%, and 100%, respectively. However, these values were all 100% for the TC electrodes (Table 3).

Impending adverse EMG signals were found in two NARs that showed significant decreases in R2 amplitude (>50% from the R1 amplitude) with the ET electrodes. However, the amplitudes from the TC electrodes did not decrease in those cases, none of which showed vocal cord palsy on postoperative laryngeal examinations. There were no TC-electrode-related complications, such as bleeding, hematoma, or other injuries.

DISCUSSION

This TC neuromonitoring technique using a needle electrode was helpful for identifying the RLN, and we were able to obtain higher-amplitude EMG signals. Furthermore, this technique provided reliable IONM during thyroidectomy through a stable EMG amplitude regardless of the position of the patient or tube. Visual identification of the RLN alone is not sufficient to avoid RLN injury during thyroidectomy, because sometimes the RLN cannot be identified, and functional preservation of the RLN cannot be guaranteed even if the nerve seems grossly intact. For this reason, IONM during thyroid and parathyroid surgery was introduced [8,21]. When IONM was first used in thyroid surgery, several methods were employed, including needle electrode-based techniques [22-24]. Among these techniques, those involving commercially available ET electrodes became widely used. The International Neuromonitoring Study Group proposed standardized monitoring guidelines and recommended that surgeons follow a defined troubleshooting algorithm in the event of EMG signal problems during ET electrode-based surgery, because such problems may be a false signal from various causes, including ET problems [15].

However, whether IONM reduces the rate of postoperative vocal cord paralysis has remained controversial [25,26]. Some authors have reported that IONM is helpful for nerve identification and dissection [25,27-29]. A multivariable analysis of RLN palsy in 1,757 patients revealed that IONM protected against permanent RLN palsy, with an odds ratio [OR] of 0.43 (95% confidence interval [CI], 0.19 to 0.93; P=0.033), although it did not reach statistical significance for protecting against transient RLN palsy (OR, 0.67; 95% CI, 0.44 to 1.01; P=0.061) [25]. However, other authors have reported that IONM did not reduce the vocal cord palsy rate, or that it was only effective in complicated cases, such as reoperations or those with a huge goiter [30,31]. Several meta-analyses have demonstrated that the use of IONM is associated with a reduced rate of transient RLN palsy, but not permanent RLN palsy [32-34].

This controversy may be due not only to the low incidence of RLN palsy, but also to the unreliable monitoring of RLN function. Unreliable monitoring causes surgeons to distrust EMG sig-
nal changes and makes it difficult for them to determine a subsequent surgical strategy when LOS occurs. Monitoring may be unreliable for several reasons, but ET displacement is the most common reason for a false adverse EMG signal because ET electrodes are contact electrodes. Several attempts have been made to overcome these limitations, such as by creating standardized guidelines or using different kinds of electrodes [21,35]. Recent reports showed that TC IONM using subdermal needle electrodes could be an effective alternative method [17,20]. Chiang et al. [20] prospectively evaluated an IONM technique in which paired needle electrodes were placed at the subperichondrium of the thyroid cartilage in 205 NARs. They reported that higher and more stable EMG amplitudes and fewer false EMG results were obtained through this technique [20]. An animal study using a porcine model also revealed that placing needle electrodes at the perichondrium of the thyroid cartilage enabled earlier identification of RLN injury and more stable monitoring [17].

We inserted the needle electrodes through the thyroid cartilage and obtained significantly higher EMG amplitudes from TC electrodes than from ET electrodes in all four-step procedures (V1-R1-R2-V2) ($P<0.001$). Many surgeons and guidelines have stated that a sufficient initial V1 amplitude (at least 500 µV) is necessary for accurate monitoring [36,37]. In our study, V1 amplitudes $>$500 µV were obtained from 19 NARs (35.2%) with ET electrodes and from 47 NARs (87.0%) with TC electrodes; thus, a significantly higher proportion of TC electrode measurements met this criterion ($P<0.001$). A high initial amplitude facilitates the early identification and localization of the RLN. During surgery, traction or electrical damage can cause the EMG amplitude to decrease, even when the nerve is grossly intact. Higher EMG amplitudes make it possible to monitor RLN function more sensitively. Thus, surgeons can obtain feedback from IONM, change their procedures, and be more careful when performing procedures near the RLN [37-39].

LOS is defined as a signal amplitude less than 100 µV and can indicate that there has been severe damage to the RLN. However, several factors can influence the EMG signal, resulting in false positive or negative results. This can make decisions difficult, such as when LOS occurs on one side during a contralateral operation [38,39]. Therefore, it is very important to know whether the LOS is a true signal. In our study, the two NARs that exhibited LOS with both the ET and TC electrodes had postoperative RLN palsy. However, vocal cord movement was intact in the three NARs that displayed LOS with only the ET electrodes, so these were false-positive findings. As a result, the positive and negative predictive values were both 100% for the TC electrodes, while they were 40% and 100%, respectively, for the ET electrodes.

The negative predictive value of IONM has reported to be varied from 12% to 88% [24,40-44]. This implies that various factors can cause false-positive LOS findings, but the most common cause is the inherent limitations of ET electrodes [33]. Our finding of a low positive predictive value for ET electrodes supports the notion that the position of the ET can be problematic. As TC electrodes enable stable monitoring without false positive or negative results, surgeons can use them confidently for IONM. Thus, TC electrodes can reduce unnecessary burdens during surgery and help surgeons to decide on a surgical strategy. In addition, TC electrodes have an economical benefit over ET electrodes. An ET with surface EMG electrodes costs about US $300 in Korea, while TC electrodes only require an EMG needle, which costs less than $10.

Despite these advantages, there are some limitations to TC electrodes. Firstly, TC electrodes are more invasive than ET electrodes [17,20]. Bleeding, hematoma, and other needle-related complications can occur. However, none of our patients experienced needle-related complications because the needles were thin. Secondly, needle electrodes can be difficult to insert if the patient's thyroid cartilage is calcified. However, in our study, the needle electrodes were inserted in all cases without much difficulty. Even if the needles cannot pass through the thyroid cartilage, the electrodes can be placed in the perichondrium of the thyroid cartilage, which can also provide enough EMG signal amplitude. In conclusion, TC electrodes are feasible and safe in IONM during thyroid surgery. This technique provides a more stable and reliable EMG signal than ET electrodes. Therefore, TC electrodes can be a good alternative to overcome the limitations of ET electrodes.

**CONFLICT OF INTEREST**

No potential conflict of interest relevant to this article was reported.

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