

# Risk Factors for Thyroid Surgery–Related Unilateral Vocal Fold Paralysis

Hung-Chun Chen, MD ; Yu-Cheng Pei, MD, PhD; Tuan-Jen Fang, MD

**Objectives/Hypothesis:** We aimed to identify the risk factors for iatrogenic unilateral vocal fold paralysis (UVFP) caused by thyroid surgery, to allow the identification of patients requiring nerve-protection procedures and monitoring technologies.

**Study Design:** Retrospective case study in a medical center.

**Methods:** Patients who underwent thyroid surgery from April 2011 to February 2016 and who were diagnosed with UVFP by laryngoscopy and laryngeal electromyography were included. Patient demographics, types of surgery, and characteristics of the thyroid lesions were analyzed.

**Results:** Sixty (2.1%) of 2,815 patients who received thyroid surgery developed UVFP. The risk of UVFP was higher in patients over 60 years old (odds ratio, 1.89; 95% confidence interval, 1.01-3.26;  $P = .01$ ). Involvement of the external branch of superior laryngeal nerve (EBSLN) occurred in 19 (31.7%) of the 60 UVFP patients, and was more likely to occur in patients with diabetes mellitus (odds ratio, 14.19; 95% confidence interval, 3.80-52.94;  $P < .001$ ). The incidence of UVFP and involvement of the EBSLN differed among surgery types, and was the highest among patients undergoing total thyroidectomy with neck dissection (TTND) (10/158, 6.3% and 5/158, 3.2%, respectively).

**Conclusions:** The risk of thyroid surgery–related UVFP is higher in older patients. EBSLN involvement is more likely in patients with diabetes mellitus. TTND is associated with higher risks of UVFP and EBSLN injury than other types of surgery, implying the need of intraoperative nerve monitoring in these high-risk characteristics.

**Key Words:** Unilateral vocal fold paralysis, thyroid surgery, external branch of superior laryngeal nerve, intraoperative nerve monitoring, laryngeal electromyography.

**Level of Evidence:** 4

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## INTRODUCTION

Unilateral vocal fold paralysis (UVFP) causes glottic insufficiency, resulting in a breathy voice, voice fatigue, and aspiration, further limiting the patient's quality of life.<sup>1</sup> Surgical injury is the most common cause of UVFP.

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The prevalence of UVFP is thus increasing with the increasing frequency of surgeries for various diseases.<sup>2–5</sup> Although previous studies have shown recent increases in the proportion of iatrogenic UVFP induced by surgeries such as cardiac, carotid, esophageal, and pulmonary surgeries, thyroid surgery remains the major cause of UVFP.<sup>2,4</sup> The incidence of vocal fold paralysis caused by thyroid surgery is reported to be 1.5% to 5.3%, among whom 15% to 17% of cases will have permanent vocal fold palsy.<sup>6–10</sup>

Laryngeal electromyography (LEMG) is the gold standard for identifying the lesioned nerve in patients with UVFP, providing diagnostic, functional, and prognostic information.<sup>11–15</sup> Quantitative LEMG has been proposed to evaluate the severity of denervation further, by analyzing the frequency of turns that reflect the remaining muscle recruitment.<sup>16</sup> Simultaneous external branch superior laryngeal nerve (EBSLN) injury can also be revealed by LEMG evaluation of cricothyroid (CT) muscle.

Thyroid surgery–related UVFP has a distinct clinical presentation, including greater involvement of the EBSLN, compared with UVFP caused by other surgeries.<sup>17,18</sup> However, thyroid surgery comprises a wide spectrum of diagnoses from benign to malignant lesions, and the types of surgery range from minimally invasive to radical procedures, such as total thyroidectomy with neck dissection (TTND). The incidence of UVFP may be different among the procedures but has not been concluded. Results from Ahn et al. showed that the UVFP rate is

higher in specific thyroidectomy with neck dissection but not noted in Sancho's report.<sup>19,20</sup> The combining of EBSLN has also not been reported in the above two reports due to lack of LEMG confirmation.

Novel intraoperative nerve monitoring (IONM) for laryngeal nerves is being developed and may reduce the risk of UVFP following thyroid surgery.<sup>21,22</sup> Regularly performing IONM during thyroid surgery was suggested, but its cost-effectiveness remains unclear.<sup>23</sup> A more effective application of IONM is reasonable in patients with higher risks.<sup>24</sup> To this end, this study aimed to identify the risk factors for thyroid surgery-related UVFP by analyzing patient demographics, comorbidities, diagnosis, and surgery type. We also used comprehensive assessments, including quantitative LEMG and quality of life assessments, to characterize the disease presentations in these patients.

## MATERIALS AND METHODS

### Human Subjects

This study included two subject populations: patients who received thyroid-related surgery from April 2011 to February 2016 in a medical center in Taiwan and were reviewed retrospectively by searching the Health Insurance Surgical Orders and by chart review; and patients with symptoms or signs of vocal fold palsy who presented after thyroid surgery and were referred to the otolaryngology outpatient clinic for further evaluation. Written informed consent was obtained from each participant in the second population before examination. UVFP and bilateral vocal fold paralysis (BVFP) were confirmed by transnasal laryngoscopy, and UVFP was further confirmed by videolaryngostroboscopy and needle LEMG. The two patient populations were combined for analysis. Patients were excluded if they had an uncertain diagnosis or follow-up, were younger than 20 years, or had BVFP. All aspects of the study were approved by the Human Studies Research Committee of Chang Gung Medical Foundation.

### Procedures

Patients with thyroid surgery-related UVFP underwent assessments including LEMG with quantitative analysis, videolaryngostroboscopy, the Voice Outcome Survey (VOS) questionnaire for voice-related quality of life, and the Short Form-36 (SF-36) Health Survey quality-of-life questionnaire. These assessments were performed within 2 weeks of the day of LEMG assessment.

### LEMG Examination

The standard protocol for needle LEMG was performed by a board-certified otolaryngologist (T.-J.F.) and a physiatrist (Y.-C.P.). The diagnosis of denervation changes of the recurrent laryngeal nerve (RLN) or concomitant EBSLN was based on abnormal findings in the thyroarytenoid-lateral cricoarytenoid (TA/LCA) and CT muscles, respectively. Patients rested on an examination chair designed specifically for LEMG, with their neck extended and head supported by the adjustable head rest. The LEMG needle-insertion site was injected subcutaneously with 0.3 to 0.5 mL of 2% lidocaine before the examination. LEMG was performed using a concentric needle electrode with a surface-ground electrode adhered to the forehead.

For the TA/LCA muscle complex, the electrode was inserted through the cricothyroid membrane at an angle 15° superiorly

and 30° laterally and off-midline. The patient was asked to produce three series of /i/ sounds at three different intensities (low, moderate, and as high as possible), with each sound lasting at least 400 ms and each inter-/i/ interval lasting approximately 200 ms. For the CT muscle, the electrode was angled 50° laterally and inserted about 5 mm lateral to the midline at the level of the center of the cricothyroid membrane. The patient was asked to produce three upward glissando /i/ sounds at conversational loudness, with each inter-/i/ interval lasting about 3 sec. Raw LEMG traces were recorded for offline quantitative analysis with sweep speeds of 10 ms per division and a gain of 200 μV per division.

Spontaneous activity and insertional activity were evaluated first. Recruitment analysis and semiquantitative motor-unit analysis were then performed, specifically when the rise time of a motor unit action potential was <0.6 ms, indicating a close proximity to the recorded motor unit. A Nicolet Viking Select (Cardinal Health, Dublin, OH) was used with a band-pass filter set between 20 and 10 kHz. An abnormal LEMG was defined as the occurrence of spontaneous activities (such as positive sharp wave, fibrillation, or complex repetitive discharge), > 30% polyphasia, or decreased interference pattern (reduced, discrete, or no interference pattern).

### Quantitative LEMG Analysis

We designed a MATLAB (MathWorks, Natick, MA)-based program to analyze the raw LEMG data. The raw LEMG waveforms were initially binned into nonoverlapping epochs. The epoch duration for the TA/LCA was 200 ms. The timing of each turn and its amplitude were localized using an automatic algorithm. Specifically, a turn was defined by a change in polarity with an amplitude ≥100 μV before and after the change, to exclude noise-related peaks. Turn frequency was computed for each epoch as the number of turns divided by the epoch duration. For each muscle, the turn frequencies for the epochs were averaged with turn frequencies that ranked among the top three epochs, to yield the peak turn frequency.

### Statistical Analysis

Differences between groups were compared using Student *t* tests for continuous data and  $\chi^2$  or Fisher exact tests for nominal data (such as sex, preoperative diagnosis of thyroid lesion, comorbidity, and type of surgery). All variables that were significant in univariate analyses were further analyzed by using a stepwise multivariate logistic regression model to estimate odds ratios and 95% confidence intervals. The level of significance was defined as *P* < .05.

## RESULTS

This study reviewed patients who received thyroid-related surgeries. None received IONM during the thyroid surgeries. The procedures were performed by general surgeons. A total of 3,527 patients receiving thyroid surgery were reviewed, among whom 704 patients were excluded because of uncertain diagnosis or follow-up (*n* = 689) or because they were under 20 years old (*n* = 15) (Fig. 1).

Sixty patients (12 males and 48 females) were noted to suffer UVFP, and eight developed BVFP, yielding overall UVFP and BVFP incidences of 2.1% and 0.3%, respectively (Fig. 2). Due to the limited sample size, the eight

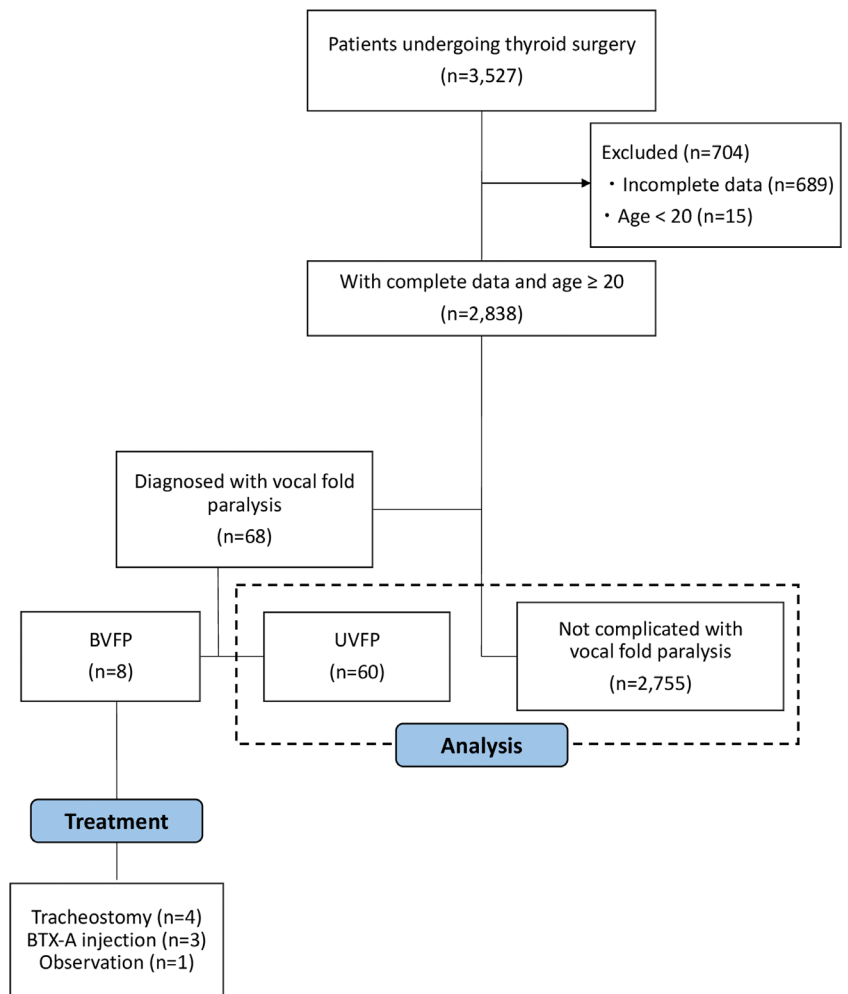


Fig. 1. Flowchart of the database search for thyroid surgery-related UVFP. BTX-A = botulinum toxin type A; BVFP = bilateral vocal fold paralysis; UVFP = unilateral vocal fold paralysis. [Color figure can be viewed in the online issue, which is available at [www.laryngoscope.com](http://www.laryngoscope.com).]

patients with BVFP were analyzed by case characteristics and were not included in the statistical analysis.

### Demographics

Regarding the surgery type, 1,273 patients underwent lobectomy (LB), 158 TTND, 982 total thyroidectomy (TT), 57 unilateral subtotal thyroidectomy (UST), and 345 bilateral subtotal thyroidectomy (BST). Patients with UVFP (UVFP group) were older than those without UVFP (no-UVFP group) ( $57.0 \pm 13.3$  vs.  $50.9 \pm 13.9$ ;  $P < .001$ ). The UVFP group had a higher proportion of patients with thyroid malignancy (benign/malignant: 35/25) than the no-UVFP group (benign/malignant: 2011/744) ( $P = .02$ ). Patients in the UVFP group were also more likely to have hypertension (6/60, 10.0% vs. 105/2,755, 3.8%;  $P = .03$ ) compared with the no-UVFP group (Table I).

According to the different types of surgery, UVFP developed in 18 with LB (1.4%), 10 patients with TTND (6.3%), 25 with TT (2.5%), 0 with UST (0%), and seven with BST (2.0%) (Fig. 3A), with the highest incidence in the TTND group and the lowest in the UST group. Among the patients undergoing TTND ( $n = 10$ ), five underwent central neck dissection, four central with lateral neck dissection,

and one lateral neck dissection. By multivariate logistic regression analysis, the risk of UVFP was higher in patients over 60 years old (odds ratio, 1.89; 95% confidence interval, 1.01-3.26;  $P = .01$ ) and highest in patients undergoing TTND (odds ratio, 3.47; 95% confidence interval, 1.42-8.44;  $P = .006$ ).

### UVFP in Patients With Thyroid Malignancies

Among 769 patients with thyroid malignancies, 143 underwent TTND and 626 underwent thyroidectomy without neck dissection. The incidence of UVFP was significantly higher in patients receiving concurrent neck dissection than in those without (13/143, 9.1% vs. 27/626, 4.3%;  $P = .03$ ).

### Patients With and Without EBSLN Involvement

All of the 2,815 patients were further divided into two groups according to the presence of UVFP combining EBSLN injury revealed by LEMG (Table II). Patients with EBSLN injury included a higher proportion of patients with thyroid malignancies (benign/malignant: 9/10) than the no-EBSLN injury group (benign/malignant: 2037/759) ( $P = .02$ ). They were

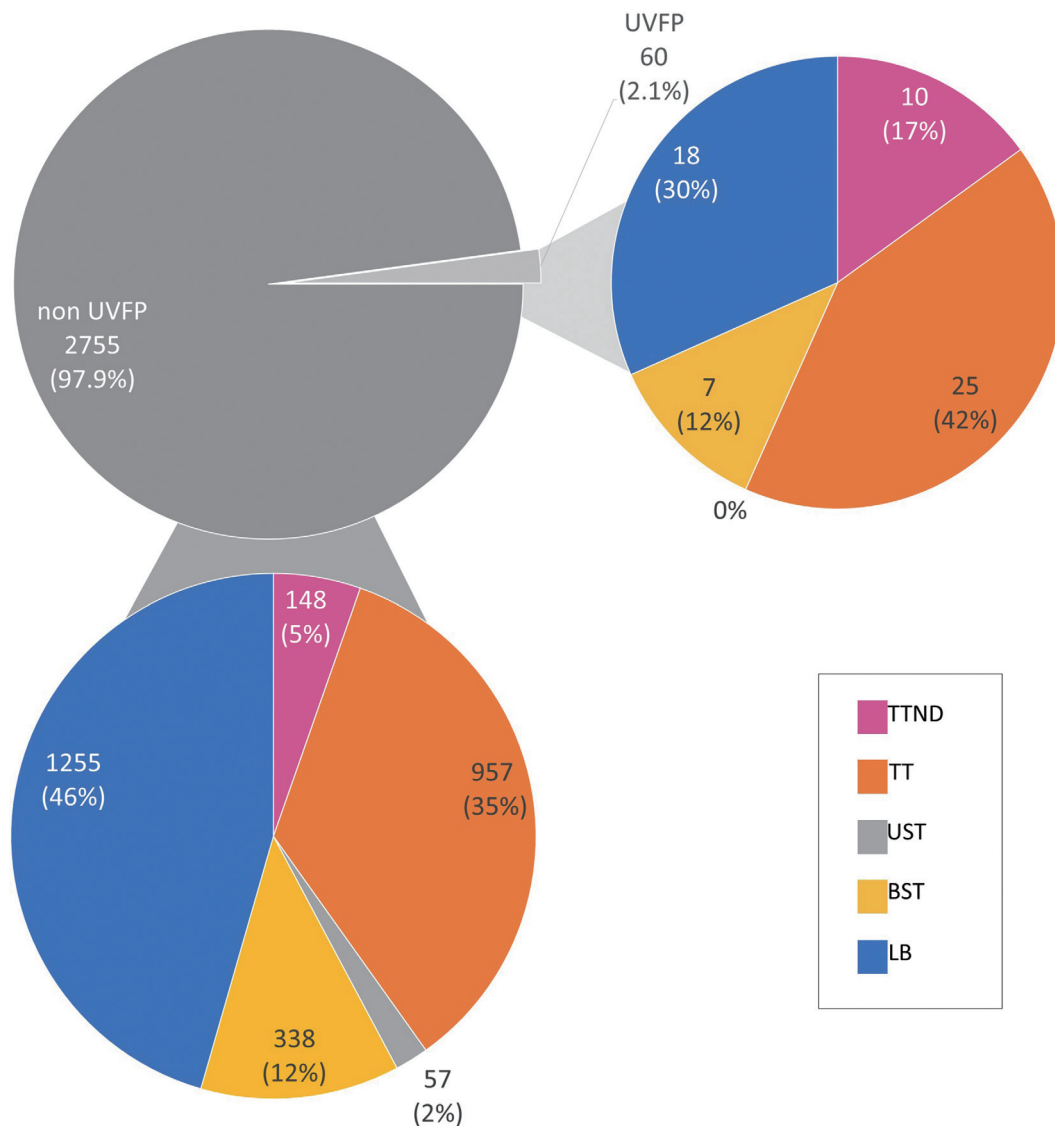


Fig. 2. Distribution of all patients undergoing thyroid surgery. Data are presented as case number (percentage). BST = bilateral subtotal thyroidectomy; LB = lobectomy; TT = total thyroidectomy without neck dissection; TTND = total thyroidectomy with neck dissection; UST = unilateral subtotal thyroidectomy; UVFP = unilateral vocal fold paralysis. [Color figure can be viewed in the online issue, which is available at [www.laryngoscope.com](http://www.laryngoscope.com).]

also more likely to have diabetes mellitus (DM) (3/19, 15.8% vs. 43/2796, 1.5%;  $P = .003$ ). The incidences of EBSLN involvement among patients with UVFP differed among surgery types, being highest in the TTND group (5/158, 3.2%) and lowest in the UST group (0/57, 0%) (Fig. 3B). By multivariate logistic regression analysis, the risk of UVFP combining EBSLN injury was higher in patients with DM (odds ratio, 14.19; 95% confidence interval, 3.80-52.94;  $P < .001$ ) and highest in patients undergoing TTND (odds ratio, 6.77; 95% confidence interval, 1.51-30.29;  $P = .007$ ).

#### UVFP Patients With and Without EBSLN Involvement

The 60 patients with UVFP were divided into two subgroups according to the LEMG findings: patients with only RLN and those with RLN + EBSLN injuries

(Table III). Nineteen of the 60 UVFP patients (31.7%) had concomitant EBSLN injury. Most of the EBSLN injuries ( $n = 16$ , 84.2%) were at the same side as RLN injury, whereas one (5.3%) was on the contralateral side and the other two (10.5%) on both sides. There was no difference between the RLN and RLN + EBSLN groups in terms of sex, age, or the side of UVFP.

#### Quantitative LEMG

Quantitative recruitment analysis revealed no difference in turn frequency or turn ratio of the lesioned TA/LCA muscle complex between patients with RLN and those with RLN + EBSLN injuries, indicating an equal level of RLN denervation. The turn frequency of the CT muscle on the lesioned side was significantly lower in the RLN + EBSLN group compared with the RLN group

TABLE I.  
Demographics of Patients Undergoing Thyroid Surgery

	No-UVFP Group	UVFP Group	P Value
No. of cases	2,755 (97.9)	60 (2.1)	
Sex (male/female)	561/2,194	12/48	1.00
Age, yr	50.9 ± 13.9	57.0 ± 13.3	<.001*
Malignancy status (benign/malignant)	2,011/744	35/25	.02 <sup>†</sup>
Comorbidity			
Hypertension (yes/no)	105/2,650	6/54	.03 <sup>†</sup>
Diabetes mellitus (yes/no)	43/2,712	3/57	.07
Heart disease (yes/no)	10/2,745	1/59	.21
Stroke (yes/no)	7/2,748	0/60	1.00
Type of surgery			
LB	1,255 (45.6)	18 (30.0)	<.001*
TTND	148 (5.4)	10 (16.7)	
TT	957 (34.7)	25 (41.7)	
UST	57 (2.1)	0 (0)	
BST	338 (12.3)	7 (11.7)	
Preoperative diagnosis			
Benign neoplasm	466 (16.9)	13 (21.7)	
Nontoxic uninodular goiter	317 (11.5)	1 (1.7)	
Nontoxic multinodular goiter	497 (18.0)	7 (11.7)	
Nontoxic diffuse goiter	14 (0.5)	0 (0)	
Toxic nodular goiter	6 (0.2)	0 (0)	
Toxic multinodular goiter	41 (1.5)	1 (1.7)	
Toxic diffuse goiter	637 (23.1)	12 (20.0)	
Thyrotoxicosis	30 (1.1)	2 (3.3)	
Thyroiditis	2 (0.1)	0 (0)	
Malignant neoplasm	745 (27.0)	24 (40.0)	

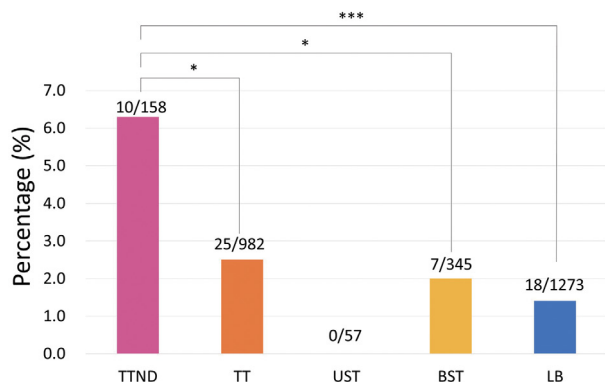
Data are presented as mean ± standard deviation or as the number (percentage) of patients.

\* $P < .001$ .

<sup>†</sup> $P < .05$ .

BST = bilateral subtotal thyroidectomy; LB = lobectomy; TT = total thyroidectomy without neck dissection; TTND = total thyroidectomy with neck dissection; UST = unilateral subtotal thyroidectomy; UVFP = unilateral vocal fold paralysis.

A. UVFP incidence



B. EBSLN injury incidence

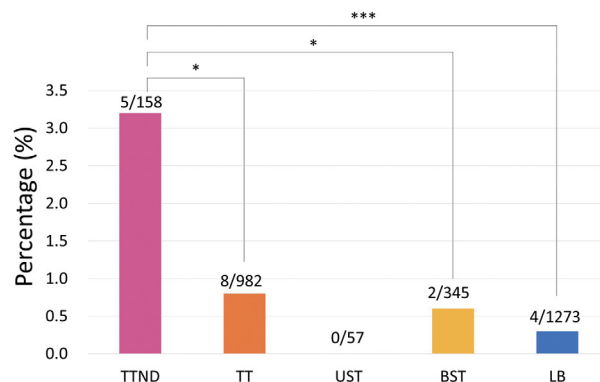


Fig. 3. Comparison of incidence across surgery types. (A) The incidence of UVFP is significantly higher in the TTND group. (B) The incidence of concomitant EBSLN injury is the highest in the TTND group. \* $P < .05$ . \*\*\* $P < .001$ . BST = bilateral subtotal thyroidectomy; EBSLN = external branch of superior laryngeal nerve; LB = lobectomy; TT = total thyroidectomy without neck dissection; TTND = total thyroidectomy with neck dissection; UST = unilateral subtotal thyroidectomy; UVFP = unilateral vocal fold paralysis. [Color figure can be viewed in the online issue, which is available at [www.laryngoscope.com](http://www.laryngoscope.com).]

TABLE II.  
Demographics of Patients Undergoing Thyroid Surgery According to the Presence of EBSLN Injury in UVFP

	No-EBSLN Group	EBSLN Group	P Value
No. of cases	2,796 (99.3)	19 (0.7)	
Sex (male/female)	567/2,229	6/13	.25
Age, yr	51.0 ± 14.0	53.8 ± 15.2	.37
Malignancy status (benign/malignant)	2,037/759	9/10	.02*
Comorbidity			
Hypertension (yes/no)	109/2,687	2/17	.17
Diabetes mellitus (yes/no)	43/2,753	3/16	.003 <sup>†</sup>
Heart disease (yes/no)	11/2,785	0/19	1.00
Stroke (yes/no)	7/2,789	0/19	1.00
Type of surgery			.001 <sup>†</sup>
LB	1,269 (45.4)	4 (21.1)	
TTND	153 (5.5)	5 (26.3)	
TT	974 (34.8)	8 (42.1)	
UST	57 (2.0)	0 (0)	
BST	343 (12.3)	2 (10.5)	
Preoperative diagnosis			
Benign neoplasm	476 (17.0)	3 (15.8)	
Nontoxic uninodular goiter	317 (11.3)	1 (5.3)	
Nontoxic multinodular goiter	503 (18.0)	1 (5.3)	
Nontoxic diffuse goiter	14 (5.01)	0 (0)	
Toxic nodular goiter	6 (0.2)	0 (0)	
Toxic multinodular goiter	42 (1.5)	0 (0)	
Toxic diffuse goiter	646 (23.1)	3 (15.8)	
Thyrotoxicosis	30 (1.1)	2 (10.5)	
Thyroiditis	2 (0.1)	0 (0)	
Malignant neoplasm	760 (27.2)	9 (47.4)	

Data are presented as mean ± standard deviation or as the number (percentage) of patients.

\* $P < .05$ .

<sup>†</sup> $P < .01$ .

BST = bilateral subtotal thyroidectomy; EBSLN = external branch of superior laryngeal nerve LB = lobectomy; TT = total thyroidectomy without neck dissection; TTND = total thyroidectomy with neck dissection; UST = unilateral subtotal thyroidectomy; UVFP = unilateral vocal fold paralysis.

(326.6 ± 221.3 vs. 753.7 ± 244.7;  $P < .001$ ), whereas the turn frequency of the CT muscle on the nonpalsy side was similar in both groups ( $P = .07$ ).

### UVFP-Related Health and General Health: VOS Questionnaire and SF-36

There was no difference in VOS score (31.1 ± 16.7 vs. 29.0 ± 16.8;  $P = .64$ ) between the RLN and RLN + EBSLN groups. There were also no differences between the two groups in any quality-of-life domains measured by the SF-36, including general health perceptions ( $P = .08$ ), vitality ( $P = .462$ ), physical functioning ( $P = .40$ ), bodily pain ( $P = .63$ ), physical role functioning ( $P = .79$ ), emotional role functioning ( $P = .60$ ), social functioning ( $P = .25$ ), and mental health ( $P = .39$ ).

### BVFP Characteristics

Eight of the 68 patients with vocal fold paralysis had BVFP. One of these patients had no dyspnea, whereas

the rest had dyspnea of differing severities. Four of these received elective tracheostomy, and three received intralaryngeal botulinum toxin type A (BTX-A; Allergan, County Mayo, Ireland) injections (Table IV).

### DISCUSSION

Thyroid diseases, including benign and malignant lesions, are common clinical problems, and the incidence of thyroid cancer has increased about twofold over the past 15 years.<sup>25</sup> The 2015 American Thyroid Association (ATA) guidelines suggest that surgical therapy should be considered for malignancies and growing benign nodules with compressive or structural symptoms,<sup>26</sup> though the extent of surgery for differentiated thyroid cancer depends on the tumor stage, cell types, and presenting function of the larynx. Notably, however, thyroid surgery remains the leading cause of UVFP, and it is therefore necessary to identify the risk factors related to UVFP in patients undergoing thyroidectomy.



TABLE III.  
Demographics, Laryngeal Muscle Recruitment, and Quality of Life in Patients With Thyroid Surgery–Related Unilateral Vocal Fold Paralysis

	RLN Group	RLN + EBSLN Group	P Value
No. of cases	41 (68.3)	19 (31.7)	
Sex (male/female)	6/35	6/13	.17
Age, yr	53.0 ± 13.5	55.2 ± 14.2	.57
Side of UVFP (left/right)	19/22	11/8	.58
Preoperative diagnosis (benign/malignant)	26/15	14/5	.56
Recruitment analysis			
Lesioned TA/LCA (turn/s)	306.0 ± 255.1	217.9 ± 203.8	.19
Normal TA/LCA (turn/s)	890.2 ± 287.6	872.5 ± 339.7	.84
Turn ratio of TA/LCA	0.35 ± 0.29	0.32 ± 0.33	.66
Lesion side of CT (turn/s)	753.7 ± 244.7	326.6 ± 221.3	<.001*
Normal side of CT (turn/s)	846.7 ± 327.0	664.6 ± 412.7	.07
Turn ratio of CT	1.01 ± 0.48	0.61 ± 0.58	.006 <sup>†</sup>
VOS	31.1 ± 16.7	29.0 ± 16.8	.64
SF-36			
Vitality	53.3 ± 16.1	49.7 ± 18.8	.46
Physical functioning	77.9 ± 19.5	82.1 ± 14.2	.40
Bodily pain	79.3 ± 19.1	76.5 ± 22.4	.63
General health perceptions	53.8 ± 19.8	44.0 ± 18.9	.08
Physical role functioning	41.3 ± 46.9	44.7 ± 49.0	.79
Emotional role functioning	54.1 ± 46.4	47.3 ± 44.9	.60
Social functioning	57.7 ± 26.3	49.5 ± 23.5	.25
Mental health	63.0 ± 17.9	58.7 ± 17.4	.39

Data are presented as mean ± standard deviation or as the number (percentage) of patients.

\* $P < .001$ .

<sup>†</sup> $P < .01$ .

CT = cricothyroid muscle; EBSLN = external branch of superior laryngeal nerve; RLN = recurrent laryngeal nerve; SF-36 = Short Form-36 Health Survey; TA/LCA = thyroarytenoid/lateral cricoarytenoid muscle complex; UVFP = unilateral vocal fold paralysis; VOS = Voice Outcome Survey.

The present study revealed that the incidence of thyroid surgery–related UVFP was higher in older patients, and patients with DM had higher risk for UVFP combining EBSLN injury. Among the different types of surgery, TTND was associated with the highest

incidences of both UVFP and EBSLN injury. These results will provide useful information for patient consultations before surgery. Aging increases the risk of malignancy and also of UVFP, according to our results. However, the decade-specific disease progression rate

TABLE IV.  
Summary of Eight Patients With Thyroid Surgery–Related Bilateral Vocal Fold Paralysis

Sex	Age (Years)	Diagnosis	Surgery Type	Surgery Duration (Minutes)	Symptoms and Signs	Intervention	Time to Intervention (Days)
Female	45	Follicular adenoma and parathyroid gland hyperplasia	LB	234	Dyspnea, choking	Tracheostomy	66
Female	51	Recurrent PTC with neck metastasis	LB	219	Dyspnea	Tracheostomy	69
Female	54	Nodular hyperplasia	LB	144	Dyspnea	Tracheostomy	6
Female	40	PTC, follicular variant, and nodular hyperplasia	TT	310	Dyspnea	Tracheostomy	28
Female	41	PTC, follicular variant and nodular hyperplasia	TT	237	Dyspnea	BTX-A injection	56
Female	54	Nodular hyperplasia	TT	212	Exertional dyspnea, falsetto	BTX-A injection	638
Female	78	PTC, pT3N1b	TTND	211	Exertional dyspnea, choking	BTX-A injection	348
Female	43	Diffuse hyperplasia with nodular formations	TT	152	Dysphonia, choking	Observation	–

BTX-A = botulinum toxin type A; LB = lobectomy; PTC = papillary thyroid carcinoma; TT = total thyroidectomy; TTND = total thyroidectomy with neck dissection.

declines with age in patients with thyroid microcarcinoma.<sup>27</sup> In light of the increased risk of surgical complications, we agree with Miyauchi<sup>27</sup> in recommending active surveillance rather than aggressive surgery in elderly patients.

The role of neck dissection in thyroid malignancies remained arguable. In a cohort of 155 patients with papillary thyroid cancer, no significant difference in the incidences of vocal fold paralysis between TTND and TT were reported.<sup>28</sup> However, Ahn et al. reported that, in the case of cN0 neck, TTND resulted in a significantly higher rate of vocal fold paralysis than only TT, and they suggested that TTND should be reserved for patients with local or neck recurrences.<sup>19</sup> Similar to the results from Ahn et al.,<sup>19</sup> the present study showed that TTND was associated with a significantly higher risk of UVFP compared with surgery without neck dissections in patients with thyroid malignancies. However, for the oncologic control, the current ATA guidelines suggest TTND for clinically involved neck metastasis and for advanced primary tumors (T3, T4), with or without neck metastasis.<sup>26</sup> Given the success of early temporary injection for UVFP,<sup>29</sup> we suggest that radical surgical treatment should still be considered in patients with high-risk primary thyroid cancers, despite the risk of surgery-related UVFP. However, it is essential to emphasize the risks prior to surgery and try to preserve nerve function without impacting on the oncologic outcomes in those requiring surgery to manage thyroid cancer.

IONM is helpful in detecting laryngeal nerve during operation, but the benefit of reducing UVFP or the cost-effectiveness of routine use of IONM remains controversial. Visual identification was reported to be more cost-effective than IONM for identifying the RLN during thyroidectomy,<sup>23,30–32</sup> whereas Al-Qurayshi et al. claimed the use of IONM has an economic advantage in cases receiving bilateral thyroid surgery as a result of the avoidance of bilateral nerve injury through surgical staging.<sup>33</sup> The present study revealed that patients with thyroid malignancies who underwent TTND were at higher risk of postoperative UVFP, implying that IONM might be beneficial in this patient population. In addition, given a higher risk of UVFP, IONM might also be considered for older patients and those with DM scheduled for thyroidectomy.

Approximately one-third of patients in the current study who developed thyroid surgery-related UVFP had concomitant EBSLN injury. The risk of EBSLN injury induced by thyroid surgery has gained increasing attention, and the ATA recommends careful dissection of the superior pole of the thyroid gland to preserve the EBSLN.<sup>26</sup> Jansson et al. found that nine of 20 patients undergoing thyroid surgery had superior laryngeal nerve (SLN) injury postoperatively, of whom three patients had partial SLN lesions before surgery,<sup>34</sup> whereas Teitelbaum and Wenig found that one of 20 patients undergoing thyroid surgery had SLN injury.<sup>35</sup> However, these studies did not consider the type of thyroid surgery or which patient groups were more likely to have EBSLN injury. In contrast, our study revealed that patients with DM were at particularly high risk. Furthermore, patients who

underwent more radical procedures were at greater risk of EBSLN injury. These results highlight the need to consider the possible complications before carrying out thyroid surgeries.

The number of UVFP cases may be overestimated because some patients may have malignancy-related UVFP before surgery, which remained undiagnosed because of their silent clinical presentation. Patients with papillary thyroid cancer were reported to have a 50% chance to having RLN paralysis before surgery, though only 64% of them had dysphonia.<sup>36</sup> A previous study showed that 1.3% of 1,923 patients who received thyroid surgery had preoperative vocal fold paralysis, and 76% of these were later proven to have malignancies.<sup>37</sup> Preoperative assessment of vocal fold paralysis should thus be carried out in patients undergoing thyroid surgery, especially for thyroid malignancies.

This study had several potential limitations. First, retrospective data may have been missing, and the Health Insurance Surgical Orders may have been imprecise. Second, the surgeries were not all performed by the same surgeon, which may have resulted in variations in the incidences of nerve injuries. Third, the incidence of postoperative UVFP may have been overestimated because preoperative UVFP was not assessed routinely and may therefore have been underestimated in patients without notable symptoms or signs. It is therefore necessary to conduct a prospective cohort study including regular pre- and postoperative assessments to verify the incidence of thyroid surgery-related nerve injuries.

## CONCLUSION

Patients receiving TTND are at the highest risk of both UVFP and concomitant EBSLN injury. Older patients are at increased risk of developing postoperative UVFP, whereas patients with DM are more likely to have EBSLN injury. The results have implications for necessary intraoperative nerve monitoring in such high-risk groups.

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## BIBLIOGRAPHY

1. Crumley RL. Unilateral recurrent laryngeal nerve paralysis. *J Voice* 1994;8:79–83.
2. Rosenthal LH, Benninger MS, Deeb RH. Vocal fold immobility: a longitudinal analysis of etiology over 20 years. *Laryngoscope* 2007;117:1864–1870.
3. Chen HC, Jen YM, Wang CH, Lee JC, Lin YS. Etiology of vocal cord paralysis. *ORL J Otorhinolaryngol Relat Spec* 2007;69:167–171.
4. Spataro EA, Grindler DJ, Paniello RC. Etiology and time to presentation of unilateral vocal fold paralysis. *Otolaryngol Head Neck Surg* 2014;151:286–293.
5. Takano S, Nito T, Tamaruya N, Kimura M, Tayama N. Single institutional analysis of trends over 45 years in etiology of vocal fold paralysis. *Auris Nasus Larynx* 2012;39:597–600.
6. Alimoglu O, Akdag M, Kaya B, et al. Recurrent laryngeal nerve palsy after thyroid surgery. *Int Surg* 2008;93:257–260.
7. Enomoto K, Uchino S, Watanabe S, Enomoto Y, Noguchi S. Recurrent laryngeal nerve palsy during surgery for benign thyroid diseases: risk factors and outcome analysis. *Surgery* 2014;155:522–528.



8. Serpell JW, Lee JC, Yeung MJ, Grodski S, Johnson W, Bailey M. Differential recurrent laryngeal nerve palsy rates after thyroidectomy. *Surgery* 2014;156:1157–1166.
9. Chiang FY, Wang LF, Huang YF, Lee KW, Kuo WR. Recurrent laryngeal nerve palsy after thyroidectomy with routine identification of the recurrent laryngeal nerve. *Surgery* 2005;137:342–347.
10. Landerholm K, Wasner AM, Jarhult J. Incidence and risk factors for injuries to the recurrent laryngeal nerve during neck surgery in the moderate-volume setting. *Langenbecks Arch Surg* 2014;399:509–515.
11. Dedo HH. The paralyzed larynx: an electromyographic study in dogs and humans. *Laryngoscope* 1970;80:1455–1517.
12. Faaborg-Andersen K, Buchthal F. Action potentials from internal laryngeal muscles during phonation. *Nature* 1956;177:340–341.
13. Blair RL, Berry H, Briant TD. Laryngeal electromyography—techniques, applications, and a review of personal experience. *J Otolaryngol* 1977;6:496–504.
14. Haglund S, Knutsson E, Martensson A. An electromyographic analysis of idiopathic vocal cord paresis. *Acta Otolaryngol* 1972;74:265–270.
15. Guindi GM, Higenbottam TW, Payne JK. A new method for laryngeal electromyography. *Clin Otolaryngol Allied Sci* 1981;6:271–278.
16. Statham MM, Rosen CA, Nandedkar SD, Munin MC. Quantitative laryngeal electromyography: turns and amplitude analysis. *Laryngoscope* 2010;120:2036–2041.
17. Pei YC, Fang TJ, Li HY, Wong AM. Cricothyroid muscle dysfunction impairs vocal fold vibration in unilateral vocal fold paralysis. *Laryngoscope* 2014;124:201–206.
18. Tseng WC, Pei YC, Wong AM, Li HY, Fang TJ. Distinct disease and functional characteristics of thyroid surgery-related vocal fold palsy. *Thyroid* 2016;26:943–950.
19. Ahn D, Sohn JH, Park JY. Surgical complications and recurrence after central neck dissection in cN0 papillary thyroid carcinoma. *Auris Nasus Larynx* 2014;41:63–68.
20. Sancho JJ, Pascual-Damieta M, Pereira JA, Carrera MJ, Fontane J, Sitges-Serra A. Risk factors for transient vocal cord palsy after thyroidectomy. *Br J Surg* 2008;95:961–967.
21. Kandil E, Mohamed SE, Deniwar A, et al. Electrophysiologic identification and monitoring of the external branch of superior laryngeal nerve during thyroidectomy. *Laryngoscope* 2015;125:1996–2000.
22. Deniwar A, Kandil E, Randolph G. Electrophysiological neural monitoring of the laryngeal nerves in thyroid surgery: review of the current literature. *Gland Surg* 2015;4:368–375.
23. Rocke DJ, Goldstein DP, de Almeida JR. A cost-utility analysis of recurrent laryngeal nerve monitoring in the setting of total thyroidectomy. *JAMA Otolaryngol Head Neck Surg* 2016;142:1199–1205.
24. Papaleontiou M, Hughes DT, Guo C, Banerjee M, Haymart MR. Population-based assessment of complications following surgery for thyroid cancer. *J Clin Endocrinol Metab* 2017;102:2543–2551.
25. Siegel RL, Miller KD, Jemal A. Cancer statistics, 2017. *CA Cancer J Clin* 2017;67:7–30.
26. Haugen BR, Alexander EK, Bible KC, et al. 2015 American Thyroid Association management guidelines for adult patients with thyroid nodules and differentiated thyroid cancer: the American Thyroid Association Guidelines Task Force on Thyroid Nodules and Differentiated Thyroid Cancer. *Thyroid* 2016;26:1–133.
27. Miyauchi A. Clinical trials of active surveillance of papillary microcarcinoma of the thyroid. *World J Surg* 2016;40:516–522.
28. Roh JL, Park JY, Park CI. Total thyroidectomy plus neck dissection in differentiated papillary thyroid carcinoma patients: pattern of nodal metastasis, morbidity, recurrence, and postoperative levels of serum parathyroid hormone. *Ann Surg* 2007;245:604–610.
29. Fang TJ, Pei YC, Li HY, Wong AM, Chiang HC. Glottal gap as an early predictor for permanent laryngoplasty in unilateral vocal fold paralysis. *Laryngoscope* 2014;124:2125–2130.
30. Higgins TS, Gupta R, Ketcham AS, Sataloff RT, Wadsworth JT, Sinacori JT. Recurrent laryngeal nerve monitoring versus identification alone on post-thyroidectomy true vocal fold palsy: a meta-analysis. *Laryngoscope* 2011;121:1009–1017.
31. Gremillion G, Fatakia A, Dornelles A, Amedee RG. Intraoperative recurrent laryngeal nerve monitoring in thyroid surgery: is it worth the cost? *Ochsner J* 2012;12:363–366.
32. Pisanu A, Porceddu G, Podda M, Cois A, Ucheddu A. Systematic review with meta-analysis of studies comparing intraoperative neuromonitoring of recurrent laryngeal nerves versus visualization alone during thyroidectomy. *J Surg Res* 2014;188:152–161.
33. Al-Qurayshi Z, Kandil E, Randolph GW. Cost-effectiveness of intraoperative nerve monitoring in avoidance of bilateral recurrent laryngeal nerve injury in patients undergoing total thyroidectomy. *Br J Surg* 2017;104:1523–1531.
34. Jansson S, Tisell LE, Hagne I, Sanner E, Stenborg R, Svensson P. Partial superior laryngeal nerve (SLN) lesions before and after thyroid surgery. *World J Surg* 1988;12:522–527.
35. Teitelbaum BJ, Wenig BL. Superior laryngeal nerve injury from thyroid surgery. *Head Neck* 1995;17:36–40.
36. Roh JL, Yoon YH, Park CI. Recurrent laryngeal nerve paralysis in patients with papillary thyroid carcinomas: evaluation and management of resulting vocal dysfunction. *Am J Surg* 2009;197:459–465.
37. Kay-Rivest E, Mitmaker E, Payne RJ, et al. Preoperative vocal cord paralysis and its association with malignant thyroid disease and other pathological features. *J Otolaryngol Head Neck Surg* 2015;44:35.