# **Review**

# The Anatomical Relationship of Inferior Thyroid Artery and Recurrent Laryngeal Nerve: A Review of the Literature and Its Clinical Importance

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## Abstract

**Background:** Thyroidectomy is a quite common daily operation in general surgery. However, the anatomical structures of the region, mainly the relationship of recurrent laryngeal nerve (RLN) and inferior thyroid artery (ITA) makes the procedure challenging. The current review of the literature aims to report the anatomical variations of this relationship.

**Methods:** The preferred reporting items for systemic reviews and meta-analyses (PRISMA) guidelines were used for the systematic review of the articles found after an extensive research through Pub-Med, Science Direct, EMBASE and Web of Science. A total of 16 studies were included for the statistical analysis.

**Results:** The results showed that the most common type of RLN was posterior to the ITA. However, according to Higgins  $I^2$  statistics the heterogeneity of the studies was quite high.

**Conclusions:** The relationship between the RLN and the ITA is quite variable. Anatomical knowledge of the region is vital for attempting to eliminate the risk of injuring the nerve during thyroidectomy.

**Keywords:** Inferior thyroid artery; Recurrent laryngeal nerve; Anatomy; Thyroidectomy

### Introduction

The anatomical relationship between the recurrent laryngeal

Manuscript submitted July 23, 2020, accepted August 4, 2020 Published online September 21, 2020

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doi: https://doi.org/10.14740/jocmr4296

nerve (RLN) and the inferior thyroid artery (ITA) is volatile (Fig. 1). The ITA usually arises from the thyrocervical trunk and supplies the thyroid gland, while the RLN, which is a mixed nerve, is one of the branches of the vagus nerve. The RLN most commonly runs anterior to the ITA on the right side and posterior to the ITA on the left side. Sporadically, the RLN passes between the branches of the ITA.

During thyroidectomy, the RLN is at risk of iatrogenic injury. This is one of the major complications of thyroidectomy surgery, occurring in 2-5% of patients without nerve variations, and it results in voice and swallowing problems. To decrease these complications, a precise surgical technique is necessary to identify the ITA, which is the landmark for recognizing the RLN. The relationship between the artery and the nerve was studied during the past century, and several classification systems were developed. The aim of our study is to provide an evidence-based assessment of the anatomical relationship between the ITA and the RLN to reduce the incidence of iatrogenic injuries and postoperative complications of thyroid surgery.

## **Materials and Methods**

#### Search method

An extensive research was performed of the following databases, which met the inclusion criteria for the analysis: PubMed, Science Direct, EMBASE, and Web of Science. The search terms used were "RLN", "ITA", "thyroid surgery", "compli-cations", "anatomy", and "variations". For the analysis, we included articles published in English from 2005 to 2019. No review protocol existed. Case reports, letters to the editors, and conference abstracts were excluded from the study, and only original articles were used. The analysis was performed according to the preferred reporting items for systemic reviews and meta-analyses (PRISMA) guidelines and conducted independently by two separate reviewers, who were responsible for the data extraction. The data included the study, type of patients (cadavers or perioperative finding), and number of RLNs and their relationship to the ITA. Further statistical consultation was conducted if there was any disparity regarding the data.

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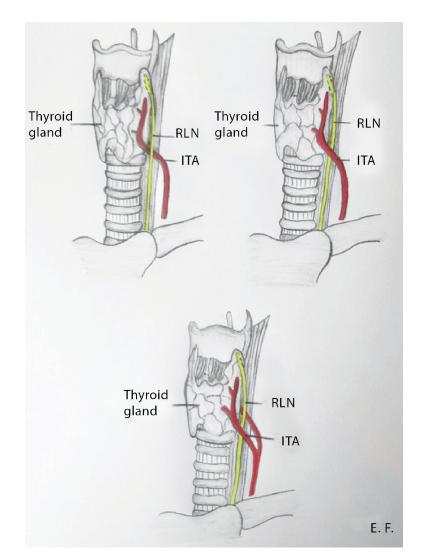


Figure 1. The anatomical relationships between the RLN and the ITA. RLN: recurrent laryngeal nerve; ITA: inferior thyroid artery.

#### Statistical analysis

The random effects model MetaXL 5.3 (freely available software for meta-analysis in Microsoft Excel) was used to examine the heterogeneity of the pooled data. The pooled prevalence and Cochran's Q test were estimated to identify heterogeneity among the studies. If the confidence intervals (CIs) of the pooled prevalence estimate overlapped poorly, this was strong evidence of statistical heterogeneity. The I<sup>2</sup> statistics (Higgins statistics) were calculated to describe the percentage of the variability in effect estimates that was due to heterogeneity rather than sampling error.

#### Results

An initial search of the databases identified a total of 578 records; 511 articles did not meet the inclusion criteria, and 38 were duplicates. Twenty-nine full-text articles were assessed for eligibility. Of these, 13 articles were excluded due to incomplete data and statistical issues, and finally, 16 articles were used in the qualitative and quantitative analysis (Fig. 2). A total of 12 studies were performed on cadavers, and the other four were from intraoperative findings (Table 1 [1-16]).

A total of 4,198 RLNs (2,099 patients) were assessed for the statistical analysis. The most common type of RLN was posterior to the ITA, with a prevalence estimate of 52.08% (95% CI: 46.5 - 57.6), while the second most common was anterior to the ITA, with a pooled prevalence of 23.91% (95% CI: 16.8 - 31.7). The least common type was between the branches of the ITA, with a prevalence estimate of 18.94% (95% CI: 13.3 - 25.6). In addition, there were differences between the left and right sides, with the posterior type being the most common, occurring in 35.73% (95% CI: 32.57 - 38.95) of the nerves on the left side and 20.78% (95% CI: 14.2 - 28.1) of those on the right one (Table 2). The data from the included studies were not statistically sufficient in order to conclude any sex or race related difference in the local anatomy of ITA and the RLN.

In all the above cases, according to the Higgins  $I^2$  statistics, the heterogeneity was quite high, except in the case of the rela-

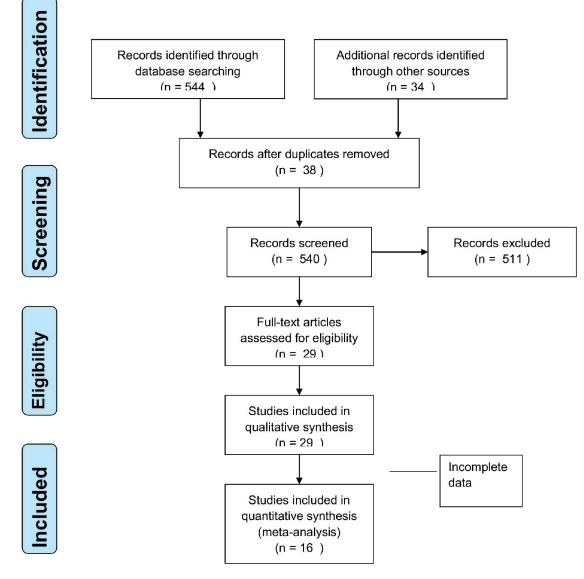


Figure 2. Flow diagram according to PRISMA guidelines. PRISMA: the preferred reporting items for systemic reviews and metaanalyses.

tionship of the RLN running posterior to the ITA at the left side. Despite the relatively large sample size more studies should be conducted to achieve more reliable conclusions (Figs. 3 - 6).

# Discussion

The first well-described enlargement of the thyroid gland goes back to the ages of the Roman Empire; later, a detailed explanation of the thyroid anatomy was given in the 16th century AD by Leonardo da Vinci [17]. Surgery for thyroid diseases has evolved over the years; however, despite the vast decrease in morbidity, this is still an issue, and injury of the RLN can have a detrimental impact on patients' quality of life and cause psychosocial distress [18]. According to several studies, injury of RLN during thyroidectomy varies from 1.5% to 14% [19-21]. Recovery takes place within 1 year in more than 80% of cases [22]. The most common cause of litigation related to performed surgical operations worldwide is iatrogenic vocal cord palsy [23]. The main mechanisms of damaging the RLN during thyroidectomy are pressure, laceration, thermal damage, division, ligation, ischemia, and manipulation [24-26]. Additional lack of surgical plan during dissection, caused by fibrosis and scar tissue resulting from previous interventions, is a well-known reason for iatrogenic damage of the RLN. The result is that 12.5% of patients suffer from temporary vocal cord paralysis, and unfortunately, 3.8% will experience permanent paralysis [27].

Fowler [28], and later, Skandalakis et al [29] concluded that injury to the RLN is more common when the nerve lies

Study	Type of patients	Number of RLN	RA	LA	Α	RP	LP	Р	В
She et al, 2005 [1]	С	99			30.3%			38.4%	31.3%
Uen et al, 2006 [2]	С	120	20.0%	8.33%	14.2%	61.67%	70.0%	65.8%	20.0%
Kuo et al, 2006 [3]	С	91			20.9%			59.3%	19.7%
Sultana et al, 2006 [4]	С	120			27.5%			30.8%	40.0%
Makay et al, 2008 [5]	IP	487	23.8%	18.9%	21.4%	63.66%	70.6%	67.2%	12.9%
Altorjay et al, 2009 [6]	IP	1,023	45.6%	15.4%	30.5%	47.70%	76.1%	61.9%	7.6%
Lee et al, 2009 [7]	С	110	40.0%	23.6%	31.8%	34.55%	56.4%	45.4%	22.7%
Sun et al, 2010 [8]	IP	566			1.94%			64.8%	31.9%
Kaisha et al, 2011 [9]	С	146			36.9%			50.0%	12.3%
Matubis et al, 2011 [10]	С	108			28.7%			59.3%	11.11%
Zhang et al, 2011 [11]	С	80			21.3%			40.0%	36.6%
Lu et al, 2012 [12]	С	100			22.0%			43.0%	35.0%
Asgharpour et al, 2012 [13]	С	246			32.9%			35.8%	31.3%
Tang et al, 2012 [14]	С	160	75.0%	7.50%	41.2%	10.00%	86.3%	48.1%	6.25%
Dai et al, 2014 [15]	IP	342			30.9%			62.9%	6.14%
Ozguner et al, 2014 [16]	С	400	11.5%	7.00%	9.25%	42.00%	65.0%	53.5%	5.00%

Table 1. The Included Studies and the Prevalence of Each Type of RLN to ITA Relationship

C: cadavers; IP: intraoperative finding; RLN: recurrent laryngeal nerve; ITA: inferior thyroid artery; RA: right anterior, LA: left anterior, A: anterior, RP: right posterior, LP: left posterior, P: posterior, B: between.

anterior to or between the ITA. Surgeons should also take care when performing operations of the thyroid on the right side because of the predominance of the anterior configuration [30].

There are studies suggesting that intraoperative nerve monitoring (IONM) has not proven more effective than direct nerve visualization has [31, 32]. However, when there is an anatomical variant, reoperation of the thyroid gland or obscured pathology IONM should be used to prevent iatrogenic injury of the RLN [33].

The purpose of our study is to highlight the variety in the normal anatomical course of RLNs during thyroidectomy to help surgeons perform safer operations and carry out accurate interventions to avoid lifelong complications in patients. In our study, after reviewing a total of 578 publications according to the PRISMA guidelines, we used 16 articles for both quantitative and qualitative analysis. After assessing 2,099 patients, we concluded that the RLN was usually running posterior to the ITA, with a prevalence of 52.08%. In addition, there were differences between the left and right sides of the thyroid gland, with the posterior type being more common on the left, representing an occurrence of 35.73%.

Despite the large sample of patients included in our study and the statistical methods, which are the strength of our analysis, the heterogeneity was quite high according to the Higgins statistics. One explanation for this is the anatomical differences among races regarding the embryological development of the nervous and vascular systems. Another issue was that most of the studies used in the analysis were performed on cadavers and not during an operation. To exclude safe results with low statistical bias, it is necessary to conduct larger studies from the same continent in which surgery is performed on living subjects.

In conclusion, the relationship between the ITA and the RLN is highly variable. Thus, surgeons should always keep in mind the anatomical relationship of the RLN to the ITA and

 Table 2.
 The Number of Studies Included, the Prevalence Estimate and the Statistical Heterogeneity According to Higgins I<sup>2</sup> Statistics

Variables	Number of studies (number of nerves)	Anterior % (95% CI)	I <sup>2</sup> % (95% CI)	Posterior % (95% CI)	I <sup>2</sup> % (95% CI)	Between % (95% CI)	I <sup>2</sup> % (95% CI)
Overall	16 (4,198)	23.9 (16.8 - 31.7)	96.7 (95.6 - 97.4)	52.1 (46.5 - 57.6)	91.7 (88.1 - 94.2)	18.9 (13.0 - 25.6)	96.1(94.8 - 97.1)
Right	6 (955)	16.5 (8.96 - 25.7)	95,9 (93.4 - 97.5)	20.8 (14.2 - 28.2)	93.1 (87.7 - 96.1)		
Left	6 (996)	06.5 (4.31 - 9.10)	77.2 (49.3 - 89.7)	35.7 (32.6 - 38.9)	52.1 (0 - 80.1)		

CI: confidence interval.

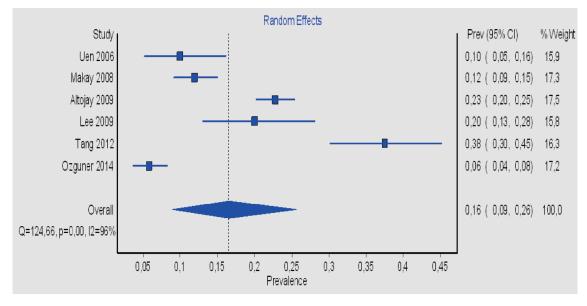


Figure 3. Forest plot for the right-sided prevalence of the RLN anterior to ITA. RLN: recurrent laryngeal nerve; ITA: inferior thyroid artery.

identify any abnormality to reduce morbidity and iatrogenic injury during thyroid surgery.

## **Conflict of Interest**

All authors declare that they have no conflict of interest.

# Acknowledgments

None to declare.

# **Financial Disclosure**

There was no funding for this study.

# **Author Contributions**

All authors contributed to the study conception and design. Material preparation, data collection and analysis were performed by Anastasios Katsourakis and George Karavasilis. The first draft of the manuscript was written by George Noussios and Anastasios Katsourakis, and all the authors commented on pre-

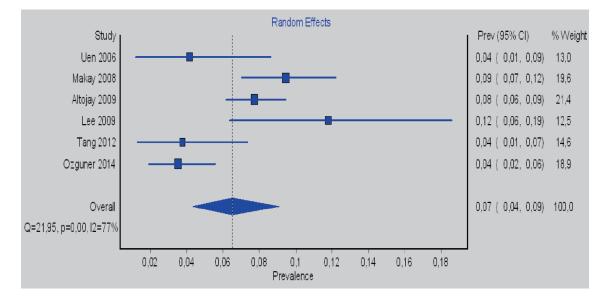


Figure 4. Forest plot for the left-sided prevalence of the RLN anterior to ITA. RLN: recurrent laryngeal nerve; ITA: inferior thyroid artery.

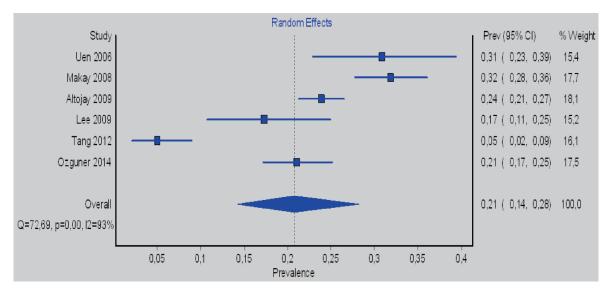


Figure 5. Forest plot for the right-sided prevalence of the RLN posterior to ITA. RLN: recurrent laryngeal nerve; ITA: inferior thyroid artery.

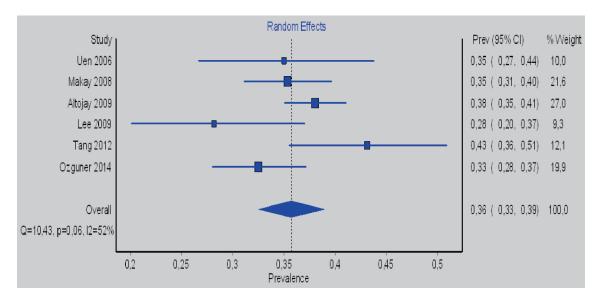


Figure 6. Forest plot for the left-sided prevalence of the RLN posterior to ITA. RLN: recurrent laryngeal nerve; ITA: inferior thyroid artery.

vious versions. All the authors approved the final manuscript.

#### **Data Availability**

The authors declare that data supporting the findings of this study are available within the article.

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