

1
2
3
4
5
6
7
8
9
10
11
12
13
14
15
16
17
18
19
20
21
22

A VARIANT NERVE THAT MIMICS THE LEFT RECURRENT LARYNGEAL NERVE: A
CASE STUDY IN HUMAN ANATOMY

Dickran Altounian¹, Cathy Tran², Christina Tran¹, Allison Spencer¹,
Alexandra Shendrik³, Brian P. Kraatz¹, Mathew J. Wedel^{1,3}

¹ College of Osteopathic Medicine of the Pacific, Western University of Health Sciences,
Pomona, CA USA

² College of Dental Medicine, Western University of Health Sciences, Pomona, CA, USA

³ College of Podiatric Medicine, Western University of Health Sciences, Pomona, CA USA

SHORT TITLE: Recurrent laryngeal nerve mimic

KEYWORDS: anatomy, larynx, nervous system, pharynx, variation

CORRESPONDING AUTHOR: Mathew J. Wedel
College of Osteopathic Medicine of the Pacific
309 East 2nd Street
Pomona, CA USA 91766
Email: mathew.wedel@gmail.com
Office: 909-469-6842

23 **ABSTRACT**

24

25 We describe a variant nerve in a human cadaver patient that parallels the course of the left
26 recurrent laryngeal nerve (RLN). Like the normal left RLN, the variant nerve branches from the
27 vagus nerve and wraps around the arch of the aorta, but it passes anterior and medial to the
28 ligamentum arteriosum (= fetal ductus arteriosus) instead of behind it like the normal RLN. After
29 recurring around the aorta, the variant nerve joins the esophageal plexus and also appears to
30 connect to the cervical sympathetic chain. The bilaterally paired RLNs supply innervation not
31 only to the larynx but also to the upper parts of the trachea and esophagus, in particular those
32 parts derived from the 4th-6th pharyngeal arches. We hypothesize that in this case, some of the
33 nerve fibers to the trachea and esophagus were pulled down into the torso by the 4th embryonic
34 aortic arch (= the arch of the aorta in adults), but passed cranial to the 6th embryonic aortic arch
35 (= fetal ductus arteriosus). From where it recurs around the aorta to join the esophageal plexus,
36 the variant nerve is very similar to the pararecurrent nerve in dogs, so there is at least a partial
37 precedent in another placental mammal. Understanding the relationships of the embryonic
38 pharyngeal and aortic arches and their adult derivatives is crucial for correctly identifying the
39 RLN, especially when imposter nerves, like the one documented here, are present.

40

41

42 INTRODUCTION

43 The recurrent laryngeal nerve is one of the most interesting gross structures in the human
44 body. It is equally famous in evolutionary biology and in clinical practice. In an evolutionary
45 context, the nerve is commonly referenced as an example of a morphological inefficiency caused
46 by a developmental constraint (Berry and Hallam, 1986; Darwen and Yao, 1993; Forsdyke,
47 1993; Coyne, 2009; Dawkins, 2009; Kinsella and Marcus, 2009; Wedel, 2012). During early
48 development, the nerves that link the brainstem to the 6th pharyngeal arches pass caudal to the
49 embryonic aortic arches. When the head and the heart are separated by the formation of the neck
50 later in development, the great vessels derived from the caudal aortic arches descend into the
51 thorax. The nerves to the 6th pharyngeal arches are dragged along by these vessels and forced to
52 assume a recurrent course back up the neck to their innervation targets in the pharynx and larynx;
53 they are referred to recurrent laryngeal nerves (RLNs), or, less commonly, recurrent pharyngeal
54 nerves. The long, inefficient pathway of the RLNs is present in all tetrapods, even long-necked
55 taxa such as ostriches and giraffes (Owen, 1841), and it appears to be an unbreakable
56 developmental constraint.

57 Clinically, physicians have known for two millennia that the RLNs must be identified and
58 protected during thyroid surgery to avoid deinnervating the patient's larynx (Kaplan et al., 2009).
59 Despite this long history of knowledge and study, injury to the RLNs is still one of the most
60 common complications from thyroid surgery. In a meta-analysis involving 16,448 operations and
61 29,998 RLNs at risk, Dralle et al. (2004: table 3) found permanent RLN paralysis in 0.84% of
62 cases overall, and in 5%-25% of patients in certain subgroups and treatments. Even
63 electromyographic monitoring of the RLN during surgery did not reduce the incidence of

64 paralysis below 0.8%. Other recent studies have found similar incidences of RLN paralysis
65 following thyroid surgery (e.g., Chan et al., 2008). The high rate of RLN injury even at the hands
66 of competent, knowledgeable surgeons is a product of two factors: first, the delicacy of the RLNs
67 themselves, which can experience physiological damage even if they are apparently intact at the
68 gross level (Crile 1932), and second, the high frequency of extra-laryngeal branching as the RLN
69 approaches the larynx (Sun et al., 2002; Yalcin et al., 2006, 2008), which led Cernea et al. (2009)
70 to suggest that the RLN should be referred to as a plexus rather than a nerve.

71 Occasionally overlooked is the fact that the RLNs innervate not only the larynx but also
72 other parts of the pharynx. In general, the portions of the pharynx that pass through the
73 embryonic pharyngeal arches are innervated by the nerves that serve each pair of arches. The
74 RLNs serve the 6th pharyngeal arches, so in addition to innervating much of the larynx, they also
75 innervate the upper portions of both the esophagus and trachea (Kuo and Urma, 2006). Of the
76 extra-laryngeal branches of the RLNs, branches to the esophagus are typically prominent (Sun et
77 al., 2002; Yalcin et al., 2006, 2008),

78 Our goals in this work are to review the normal anatomy of the RLN, and to describe and
79 illustrate a variation related to the RLN that we have not seen documented previously: a second
80 large nerve on the left side that takes a recurrent course around the aorta, mirroring the path of
81 the left RLN, albeit on the opposite side of the ligamentum arteriosum.

82

83 **BACKGROUND**

84 The earliest surviving description of the RLN comes from the 2nd century writings of the
85 Roman physician Galen. To demonstrate the nerve's involvement in vocal activity, Galen

86 famously exposed the RLNs in a live, squealing pig, and then severed both nerves, rendering the
87 pig mute. In the 1500s, Vesalius produced the oldest surviving drawings that map out the course
88 of RLN and its major branches. Throughout the post-Roman history of Western medicine,
89 knowledgeable surgeons have attempted to avoid damage to the RLNs during thyroid surgery,
90 thereby preserving their patients' voices (Kaplan et al., 2009).

91 The following description of the paths of the RLNs is based on Steinberg et al. (1986).

92 The RLN originates from the vagus nerve (cranial nerve X). In the middle and lower part of the
93 neck, the vagus nerve courses bilaterally alongside the common carotid artery and internal
94 jugular vein, typically inside the posterior wall of the carotid sheath. On the right side, the right
95 RLN branches from the vagus nerve at the level of the right subclavian artery. The right RLN
96 loops below and around the proximal end of the right subclavian artery to ascend up the neck in
97 the right tracheo-esophageal groove. The right RLN may cross superficially, deep, or between the
98 branches of the inferior thyroid artery as it proceeds superiorly. The left RLN arises lateral to the
99 arch of the aorta, and passes below it lateral to the ligamentum arteriosum (= fetal ductus
100 arteriosus), and ascends in the left tracheo-esophageal groove.

101 There are typically two divisions of the RLN: the cricopharyngeal nerve, which
102 innervates part of the inferior pharyngeal constrictor muscle, and the laryngeal nerve, which
103 innervates much of the larynx. The laryngeal branch of the RLN further divides into an anterior
104 motor branch and a posterior sensory branch. The sensory branch supplies sensory filaments to
105 the laryngeal mucosa below the vocal folds. Occasionally the sensory branch blends with the
106 internal laryngeal nerve, itself a branch of the superior laryngeal nerve (SLN). If present, this
107 interweaving of nerve fibers from the internal laryngeal nerve (SLN) and the laryngeal branch of

108 the RLN is known as the ramus anastomoticus or Galen's anastomosis (Sanders et al., 1993).
109 Sanders et al. (1993) also found other, finer connections between the SLN and RLN, particularly
110 in the area of the interarytenoid muscle, where fibers of both nerves form a tiny plexus.

111 The motor branches of the RLNs supply all of the intrinsic muscles of the larynx except
112 the cricothyroids. In particular, the RLNs innervate the posterior cricoarytenoid muscles, which
113 are the only muscles to abduct the vocal folds (Standring, 2008). Unilateral RLN palsy will cause
114 hoarse speech, and if both RLNs are compromised, the patient will be left mute and may
115 experience difficulty breathing since there will be no way to abduct the vocal folds, as Galen
116 demonstrated to the elders of Rome by vivisecting a pig and severing its RLNs (Kaplan, et al.
117 2009). King and Gregg (1948) suggested that variant vocal cord paralysis could be explained by
118 the presence of adductor (anterior) and abductor (posterior) branches of the recurrent laryngeal
119 nerve. Furthermore, Nemiroff and Katz (1982) found that the posterior branches were frequently
120 smaller than the anterior branches; this in turn increases the possibility that these smaller
121 branches—which innervate the all-important posterior cricoarytenoid muscles—can be
122 overlooked or inadvertently injured during thyroid surgery, potentially leading to transient or
123 permanent vocal cord paralysis (Nemiroff and Katz, 1982). The small but crucial posterior
124 branches of the RLN will be especially vulnerable if they originate outside the larynx; as early as
125 1948, King and Gregg noted that the branches of the RLN were relatively safe from surgical
126 injury above the inferior margin of the thyroid cartilage.

127 The inferior thyroid artery is sometimes used as an anatomic landmark in search for the
128 RLN (Sun et al., 2001), but this is problematic, because the relationship of the RLNs to the
129 inferior thyroid arteries is so variable, not only from patient to patient but even between the left

130 and right sides in a single individual. Reed (1943) reported that the right and left recurrent
131 laryngeal nerve were alike in only 43 of 253 cadavers (17%). Reed also identified three possible
132 paths for the RLN relative to the inferior thyroid artery: (a) superficial to the artery and its
133 branches (18.6%), (b) deep to the artery and its branches (39.1%), and (c) between the branches
134 of the artery (36.5%). Yalcxin (2006) and Bergman (2011) both found that the right RLN more
135 commonly lies anterior or superficial to the inferior thyroid artery, whereas the left RLN
136 frequently passes posterior or deep to the artery.

137 As noted in the Introduction, the RLNs also innervate portions of the esophagus and
138 trachea. The development and innervation of the esophagus are complex; in the following
139 summary we have drawn from the works of Goyal et al. (1999) and Kuo and Urma (2006). The
140 esophageal plexus contains parasympathetic fibers from the vagus nerve and sympathetic fibers
141 from the cervical and thoracic sympathetic trunks. Parasympathetic innervation comes from the
142 nucleus ambiguus and dorsal motor nucleus of the vagus nerve. Both nuclei serve as the origin
143 of the motor functions of the upper esophagus. The esophagus can be divided into three portions:
144 cervical, thoracic, and abdominal. The cervical esophagus extends from the pharyngoesophageal
145 junction to the suprasternal notch, and is bordered anteriorly by the trachea, posteriorly by the
146 vertebral column, and laterally by the carotid sheaths. The upper portion of the esophagus
147 consists of striated muscle, and is derived from mesenchyme of pharyngeal arches 4 and 6 (Kuo
148 et al., 2006); the 5th arch is rudimentary in humans and has no known adult derivatives
149 (O’Rahilly and Tucker, 1973). The middle and lower portions of the esophagus are smooth
150 muscle, and are derived from the mesenchyme of somites surrounding the foregut. The upper
151 esophageal sphincter is derived in part from the 6th pharyngeal arch, and it is therefore innervated

152 by the RLNs, which serve that arch. The RLNs therefore supply the upper esophageal sphincter
153 (in part) and the cervical portion of the esophagus, and the vagus nerves contribute to the
154 esophageal plexus that innervates the remainder of the esophagus.

155

156 **MATERIALS AND METHODS**

157 The variant nerve described herein was discovered by the students at Table 30 in the
158 Medical Gross Anatomy laboratory at Western University of Health Sciences in the fall of 2011,
159 most of whom are now authors of this work. The cadaver patient is an 82-year-old male who
160 died of adenocarcinoma of the pancreas, and whose remains were donated to the anatomy
161 program at Western University of Health Sciences through the university's willed body program.
162 All of the dissections were performed by the student authors, using standard surgical dissection
163 tools. The work was reviewed and approved by the Institutional Review Board at Western
164 University of Health Sciences (protocol 14/RFD/006).

165

166 **RESULTS**

167 **Description**

168 The variant nerve was discovered by the students at Table 30 in the Medical Gross
169 Anatomy laboratory at Western University of Health Sciences in the fall of 2011, most of whom
170 are now authors of this work. During a routine dissection of the heart and mediastinum of our
171 cadaver patient, we noticed a large nerve that descended from the superior thorax to wrap around
172 the aorta on its anterior aspect (Fig. 1). We will refer to this nerve herein as the variant nerve.
173 The cadaver patient also had a variation in the branching of the great vessels near the heart: the

174 left vertebral artery branched directly from the aortic arch, between the left common carotid and
175 left subclavian arteries (Fig. 2). We identified this extra branch of the aortic arch as the vertebral
176 artery based on subsequent dissections. Although the variant origin of the left vertebral artery is
177 interesting in itself, it is fairly common and well-understood (Uchino et al., 2013) and is
178 probably not related to the variant nerve.

179 The variant nerve branched from the vagus nerve above the level of the subclavian artery
180 (Fig. 2). It wrapped around the aorta anteriorly and inferiorly, taking a similar course to that of
181 the recurrent laryngeal nerve, with one important difference: whereas the recurrent laryngeal
182 nerve passed posterior to the ligamentum arteriosum, the variant nerve passed medially to it and
183 was not closely related to the ligament (Fig. 1, 2, and 3). As the variant nerve passed around the
184 aorta, it contributed branches to the cardiac plexus (Fig. 3). Posterior to the aorta, the variant
185 nerve contributed to and essentially disappeared within the dense and complex esophageal plexus
186 (Fig. 4). At this point, it became impossible to trace all of the possible connections of the variant
187 nerve, although at least one prominent strand of nervous tissue (highlighted in Fig. 4) connected
188 the variant nerve to the stellate ganglion of the left cervical sympathetic chain. We were unable
189 to detect any connections between the esophageal plexus and the larynx, so although the variant
190 nerve did recur around the aorta, it probably did not supply any structures other than the
191 esophagus and trachea.

192 In contrast to the variant nerve, the left recurrent laryngeal nerve passed mostly posterior
193 to the esophageal plexus and appeared to share few connections with it (Fig. 4). We did not find
194 any significant extralaryngeal branches of the left recurrent laryngeal nerve. Superiorly it passed
195 posterior to the inferior thyroid artery before entering the larynx (Fig. 5). The branches of the left

196 recurrent laryngeal nerve inside the larynx were normal and the ramus anastomoticus or Galen's
197 anastomosis (between the recurrent laryngeal nerve and the internal branch of the superior
198 laryngeal nerve) was not present on the left.

199 The right recurrent laryngeal nerve contributed many small branches to the esophagus,
200 and passed anterior to the inferior thyroid artery before entering the larynx (Fig. 6). As on the
201 left, we did not note any significant extralaryngeal divisions other than those to the esophagus.
202 Inside the larynx, the right recurrent laryngeal nerve was joined with the internal laryngeal nerve
203 to form Galen's anastomosis (Fig. 7).

204

205

206 **DISCUSSION**

207 **Identity of the Variant Nerve**

208 In the absence of neuronal tracing, any identification of the variant nerve can only be a
209 hypothesis based on its gross anatomy. This necessarily tentative identification is subject to
210 further test if similar patterns of nerves are found in living patients or laboratory organisms in the
211 future.

212 Cardiac branches from the vagus nerve and cervical sympathetic trunk commonly course
213 anteriorly over the aortic arch, medial to the left recurrent laryngeal nerve (see, e.g., Mizeres,
214 1963; Standring, 2008: fig. 56.20) or occasionally joined with it (Lemere, 1932). These cardiac
215 nerves typically form a dense plexus on the anterior aspect of the aortic arch, with individual
216 nerves that are quite small (< 1mm in diameter). One possibility is that our variant nerve is
217 simply this group of cardiac nerves, fused into a single connective sheath instead of distributed

218 into a plexus of many fine branches. This is unlikely to be a complete explanation, however,
219 because a comparatively thick trunk of the variant nerve passed around the aorta and took a short
220 recurrent course into the esophageal plexus (Figs. 3 and 4).

221 Proximally, the variant nerve was clearly connected to the vagus nerve, and first appeared
222 to us a branch of the vagus nerve above the subclavian artery (Fig. 2). It also appeared to share a
223 connection to the stellate ganglion of the cervical sympathetic chain—at least, a prominent band
224 of nervous tissue connected the stellate ganglion to the portion of the variant nerve that wrapped
225 around the aorta inferiorly (Fig. 4). Distally (i.e., closer to innervation targets) the variant nerve
226 contributed to both the cardiac plexus (Fig. 3) and the esophageal plexus (Fig. 4). Given this
227 pattern of connections, it is likely that the variant nerve carried both preganglionic
228 parasympathetic fibers derived from the vagus nerve, and postganglionic sympathetic fibers from
229 the stellate ganglion, to both the esophageal plexus and the cardiac plexus. The portion of the
230 nerve that wrapped around the aorta inferiorly might have contained parasympathetic and
231 sympathetic nerve fibers running in opposite directions—parasympathetic fibers from the vagus
232 nerve to the esophageal plexus wrapping around the aorta from front to back, and sympathetic
233 fibers from the cervical sympathetic chain to the cardiac plexus running around the aorta from
234 back to front.

235 Although we have not found any previous reports of a similar division of the recurrent
236 nerves to the pharynx from the rest of the RLN in humans, this division is present in dogs.

237 Lemere (1932: p. 422) described it as follows:

238 “In the dog, immediately after its origin, [the RLN] divides into the recurrent
239 proper and what might be termed the pararecurrent. The first courses without

240 interruption to the muscles of the larynx. The pararecurrent supplies the trachea,
241 esophagus, and infracordal mucosa of the larynx posteriorly. In man, of course,
242 this component remains within the recurrent. The pararecurrent may rarely be
243 included in the recurrent in dogs, but it usually runs as a separate nerve or as a
244 plexus.”

245 This description and the accompanying illustration (Lemere 1932: fig. 1) match very closely the
246 variant nerve documented here. It is interesting that Lemere (1932) found the pararecurrent nerve
247 as a separate branch in most cases in dogs, but apparently blended with the RLN. The opposite
248 situation apparently holds in humans, where normally the esophageal and tracheal branches
249 remain with the RLN, but—at least in this case, and possibly in others not yet documented—they
250 branch off as a separate, pararecurrent nerve. Possibly the pararecurrent nerve is a latent
251 developmental possibility in most mammals, which is expressed at different frequencies in
252 various mammalian lineages.

253

254 **Developmental Basis**

255 The upper part of the esophagus is derived from pharyngeal arches 4 and 6. In humans,
256 these arches and their derivatives are innervated by the vagus nerve: the 4th arch is served by the
257 superior laryngeal nerve and the pharyngeal plexus, the 5th arch is entirely resorbed and has no
258 persistent derivatives, and the 6th arch is innervated by the recurrent laryngeal nerve (O’Rahilly
259 and Tucker, 1973; Standring, 2008). In early development the pharyngeal arches are located just
260 ventral to the brain and brainstem, and the nerves take a straight path from the central nervous
261 system to the muscles, glands, and other innervation targets in the pharyngeal arches. At this

262 early stage, the nerves that serve the pharyngeal arches become entwined with the embryonic
263 aortic arches, which also serve the pharyngeal arches. Later in development the head and the
264 heart are separated by the formation of the neck, and the heart and the great vessels descend into
265 the thorax—specifically, the great vessels derived from the 4th and 6th aortic arches. When these
266 vessels move down into the thorax, they drag along the nerves that serve the 6th arch. These
267 nerves are forced to grow in length to maintain the connections between brainstem and the
268 pharynx, and take on a recurrent course. These nerves serve all of the derivatives of the 6th
269 pharyngeal arches, including not only the larynx and its muscles and mucosa but also the
270 portions of the esophagus and trachea associated with the 6th arch. The name 'recurrent laryngeal
271 nerve' does not reveal the full scope of the activities of this nerve, because it omits the
272 innervation of the relevant portions of the esophagus and trachea. 'Recurrent pharyngeal nerve'
273 would be more accurate, and indeed the nerve in question is occasionally referred to by that
274 name, but mostly in older literature (e.g., Hooper, 1885; Messerklinger and Propst, 1953).

275 The function of the recurrent laryngeal nerve in innervating part of the esophagus is
276 relevant to current case. In all humans and indeed in all vertebrates, we would expect that some
277 of the neurons innervating the proximal third of the esophagus would be forced to take a
278 recurrent course around the great vessels near the heart. Normally these neurons are bundled by
279 connective tissue into the gross structure that we recognize as the recurrent laryngeal (or
280 recurrent pharyngeal) nerve. On the right side, this can hardly fail to happen; the only remnant of
281 embryonic aortic arches 4-6 is the right subclavian artery (4th arch), so there is one vascular
282 'hook' to pull the nerves to pharyngeal arches 4-6 down into the chest. But on the left, there are
283 two such 'hooks': the arch of the aorta (4th embryonic arch), and the ductus arteriosus (6th arch).

284 So it is at least theoretically possible that vagal branches to portions of the pharynx other than
285 those derived from the 6th arch could pass distal to the 4th embryonic aortic arch (= arch of the
286 aorta in adults), but proximal to the 6th aortic arch (= ductus arteriosus, or ligamentum
287 arteriosum in adults). We suspect that this happened in the current case; it seems to be the only
288 explanation that is fully consistent with embryology that also explains all of the morphological
289 features of the variant nerve.

290

291 **Clinical Implications**

292 The main clinical implication of the variant nerve documented here is that it is so large
293 and its course is so similar to that of the left RLN that the two could potentially be confused. The
294 variant nerve differs from the left RLN in three respects: (1) it branches from the vagus much
295 farther superiorly, near the root of the neck, whereas the normal RLN branches from the vagus as
296 the latter nerve passes the arch of the aorta; (2) the variant nerve passes anterior and medial to
297 the ligamentum arteriosum, instead of behind it; and (3) the variant nerve apparently innervated
298 only portions of the esophagus and trachea, but not the larynx, whereas the left RLN in this case
299 served the larynx but had few connections to the esophageal plexus (in contrast to the numerous
300 esophageal branches of the right RLN—see Figure 6). The number and fineness of the nerve
301 fibers in the esophageal plexus on both sides (see Figures 4 and 6) should be of interest to all
302 students of anatomy, especially surgeons.

303 Of the three visual criteria for distinguishing the variant nerve from the normal RLN, the
304 relationship of the RLN to the ligamentum arteriosum is the most important, because it is
305 apparently invariant. Non-recurrent inferior laryngeal nerves occur reasonably frequently on the

306 right side, in 0.3%-1.6% of patients. Non-recurrent nerves on the left are much less frequent
307 (0.04%), and they are always associated with situs inversus, in which the normal bilateral
308 asymmetry of the internal organs is reversed from left to right (Toniato et al., 2004). In cases
309 where the inferior laryngeal nerve is non-recurrent, it is always the case that the right subclavian
310 artery (or left subclavian artery, in patients with situs inversus) is formed by a segmental branch
311 from the descending aorta, distal to the contralateral subclavian artery. What this means is that
312 the 4th embryonic aortic arch on the right side (or the left, in situs inversus) was completely
313 resorbed during development, and there was no remaining vascular 'hook' to drag the inferior
314 laryngeal nerve down into the torso (Henry et al., 1988). But even in these cases, the RLN that
315 passes under the arch of the aorta and the ligamentum arteriosum (normally the left RLN, but the
316 right RLN in situs inversus) is still present; there are no known cases of bilaterally non-recurrent
317 inferior laryngeal nerves. Therefore the ligamentum arteriosum is the most crucial landmark for
318 identifying the left RLN (or the right one, in situs inversus). This is particularly important in
319 cases like the one documented here, in which an imposter nerve is also present and takes a
320 recurrent course around the aorta, but not around the ligamentum arteriosum.

321

322 **CONCLUSIONS**

323 In this work we have shown that nerves to other portions of the pharynx (including for
324 our purposes the esophagus and trachea) can also take a recurrent course around the aorta and
325 visually mimic the RLN. In this case, the variant nerve could be distinguished from the normal
326 RLN by its path anterior and medial to the ligamentum arteriosum, rather than posterior and
327 lateral to it. Innervation to the esophagus is complex and involves both recurrent and

328 nonrecurrent fibers of the vagus nerve. Normally the recurrent fibers to the larynx, esophagus,
329 and trachea are all bundled into the RLNs, but in this case some may have passed caudal to the
330 4th embryonic aortic arch (the arch of the aorta in adults) but proximal to the 6th aortic arch (the
331 ductus arteriosus, or ligamentum arteriosum in adults), producing the variant nerve. This variant
332 nerve is very similar to the pararecurrent nerve of dogs, which also recurs around the great
333 vessels to serve the esophagus and trachea. Despite almost 2000 years of study, the RLN remains
334 a challenging structure for anatomists and surgeons because of its complexity, fragility, and
335 variability.

336

337 **ACKNOWLEDGEMENTS**

338 Above all, we are grateful to our donor patient for giving us the opportunity to learn from
339 his cadaveric remains. We thank Nina McCoy of the Western University of Health Sciences
340 Willed Bodies Program for arranging access to the cadaver patient, and Craig Kuehn of the
341 Department of Anatomy at WesternU for assistance and advice during the study.

342

343 **REFERENCES**

344 Bergman, R.A., Afifi, A.K., and Miyauchi, R. (2014). Recurrent Laryngeal Nerve. *Illustrated*
345 *Encyclopedia of Human Anatomic Variation: Opus III: Nervous System: Cranial Nerves and*
346 *Ganglia.*
347 [http://www.anatomyatlases.org/AnatomicVariants/NervousSystem/Text/RecurrentLaryngealNer](http://www.anatomyatlases.org/AnatomicVariants/NervousSystem/Text/RecurrentLaryngealNerve.shtml)
348 [ve.shtml](http://www.anatomyatlases.org/AnatomicVariants/NervousSystem/Text/RecurrentLaryngealNerve.shtml). Retrieved November 20, 2014.

349

350 Berry, R.J. and Hallam, A. 1986. The Collins Encyclopedia of Animal Evolution. 160 pp.
351 Collins, London.

352

353 Cernea, C. R., Hojajj, F. C., De Carlucci, D., Gotoda, R., Plopper, C., Vanderlei, F., & Brandão,
354 L. G. (2009). Recurrent laryngeal nerve: a plexus rather than a nerve?. Archives of
355 Otolaryngology–Head & Neck Surgery, 135(11), 1098-1102.

356

357 Chan, W. F., Lang, B. H. H., & Lo, C. Y. (2006). The role of intraoperative neuromonitoring of
358 recurrent laryngeal nerve during thyroidectomy: a comparative study on 1000 nerves at risk.
359 Surgery, 140(6), 866-873.

360

361 Coyne, J.A. 2009. Why Evolution Is True. 304 pp. Penguin Group, New York.

362

363 Crile, G. (1932) Diagnosis and treatment of diseases of the thyroid gland. W.B. Saunders,
364 Philadelphia.

365

366 Darwen, P.J. and Yao, X. 1993. On evolving robust strategies for iterated prisoner's dilemma. In:
367 X. Yao (ed.), Proceedings of the AI'93 Workshop on Evolutionary Computation, 49–63.
368 Australian Defense Force Academy, Canberra.

369

370 Dawkins, R. 2009. The Greatest Show on Earth: The Evidence for Evolution. 496 pp. Free Press,
371 New York.

372

373 Dralle, H., Sekulla, C., Lorenz, K., Brauckhoff, M., & Machens, A. (2008). Intraoperative
374 monitoring of the recurrent laryngeal nerve in thyroid surgery. *World journal of surgery*, 32(7),
375 1358-1366.

376

377 Forsdyke, D.R. 1993. On giraffes and peer review. *FASEB Journal* 7: 619–621.

378

379 Goyal R, Sivarao D. Functional anatomy and physiology of swallowing and esophageal motility.
380 In: Catell OD, Richter JE, eds. *The Esophagus*, 3rd ed. Philadelphia: Lippincott Williams &
381 Wilkins, 1999:24–26.

382

383 Hooper, F.H. 1885. The respiratory function of the human larynx, from experimental studies in
384 the physiological laboratory of Harvard University. *Boston Medical and Surgical Journal* 113(2):
385 38-39.

386

387 Kaplan EL, Salti GI, Roncella M, Fulton N, and Kadowaki M. “History of the Recurrent
388 Laryngeal Nerve: From Galen to Lahey.” *World Journal of Surgery*. (2009); 33: 386-393.

389

390 King BT, Gregg RL. An Anatomical reason for various behaviors of paralysed vocal cords. *Ann*
391 *Otol Rhinol Laryngol* 1948; 57:925-44.

392

393 Kinsella, A.R. and Marcus, G.F. 2009. Evolution, perfection and theories of language.
394 *Biolinguistics* 3: 186–212.
395
396 Kuo B and Urma D. “Esophagus—anatomy and development.” *GI Motility Online* (2006).
397 <http://www.nature.com/gimo/contents/pt1/full/gimo6.html>. Retrieved November 20, 2014.
398
399 Lemere, F. 1932. Innervation of the larynx. I. Innervation of laryngeal muscles. *American*
400 *Journal of Anatomy* 51(2): 417-437.
401
402 Messenklinger, W., and Propst, A. 1953. [Recurrent pharyngeal nerve paralysis and
403 cricoarytenoid joint]. *Monatsschrift fur Ohrenheilkunde und Laryngo-Rhinologie* 87(3): 208-
404 213. [in German]
405
406 Mizeres, N. J. (1963). The cardiac plexus in man. *American Journal of Anatomy*, 112(2), 141-
407 151.
408
409 Nemiroff PM, and Katz AD. "Extralaryngeal Divisions of the Recurrent Laryngeal Nerve:
410 Surgical and Clinical Significance." *The American Journal of Surgery*. (1982); 144: 466-469.
411
412 O'Rahilly, R., & Tucker, J. A. (1973). The early development of the larynx in staged human
413 embryos. I. Embryos of the first five weeks (to stage 15). *The Annals of otology, rhinology, and*
414 *laryngology*, 82(1): 3-27..

415 Owen, R. 1841. Notes on the anatomy of the Nubian giraffe (*Camelopardalis*). Transactions of
416 the Zoological Society of London 2: 217–248.

417

418 Reed, A.F. (1943) The relations of the inferior laryngeal nerve to the inferior thyroid artery.
419 Anat. Rec. 85:17-23.

420

421 Sanders I, Wu BL, Mu L, Li Y, and Biller H. "The Innervation of the Human Larynx." Arch
422 Otolaryngology Head Neck Surgery. (1993); 119: 934-939.

423

424 Standring, S. (ed.) 2008. Gray's Anatomy, 40th edition. Churchill Livingstone Elsevier.

425

426 Steinberg J., Khane GJ, Fernandes CMC, and Nel JP. "Anatomy of the Recurrent Laryngeal
427 Nerve: A Redescription. The Journal of Laryngology and Otology. (1986); 100: 919-927.

428

429 Sun SQ, Zhao J, Lu GO, He J, Ran JH, and Peng XH. "An Anatomical Study of the Recurrent
430 Laryngeal Nerve: Its Branching Patterns and Relationship to the Inferior Thyroid Artery."
431 Surgery Radiology Anatomy. (2001). 23: 363-369.

432

433 Toniato A, Mazzarotto R, Piotto A, Bernante P, Pagetta C, and Pelizzo MR. "Identification of
434 the Nonrecurrent Laryngeal Nerve during Thyroid Surgery: 20-Year Experience." World Journal
435 of Surgery. (2004); 28: 659-661.

436

437 Uchino, A., Saito, N., Takahashi, M., Okada, Y., Kozawa, E., Nishi, N., ... & Watanabe, Y.
438 (2013). Variations in the origin of the vertebral artery and its level of entry into the transverse
439 foramen diagnosed by CT angiography. *Neuroradiology*, 55(5), 585-594

440

441 Wedel, M.J. 2012. A monument of inefficiency: The presumed course of the recurrent laryngeal
442 nerve in sauropod dinosaurs. *Acta Palaeontologica Polonica* 57 (2): 251–256.

443

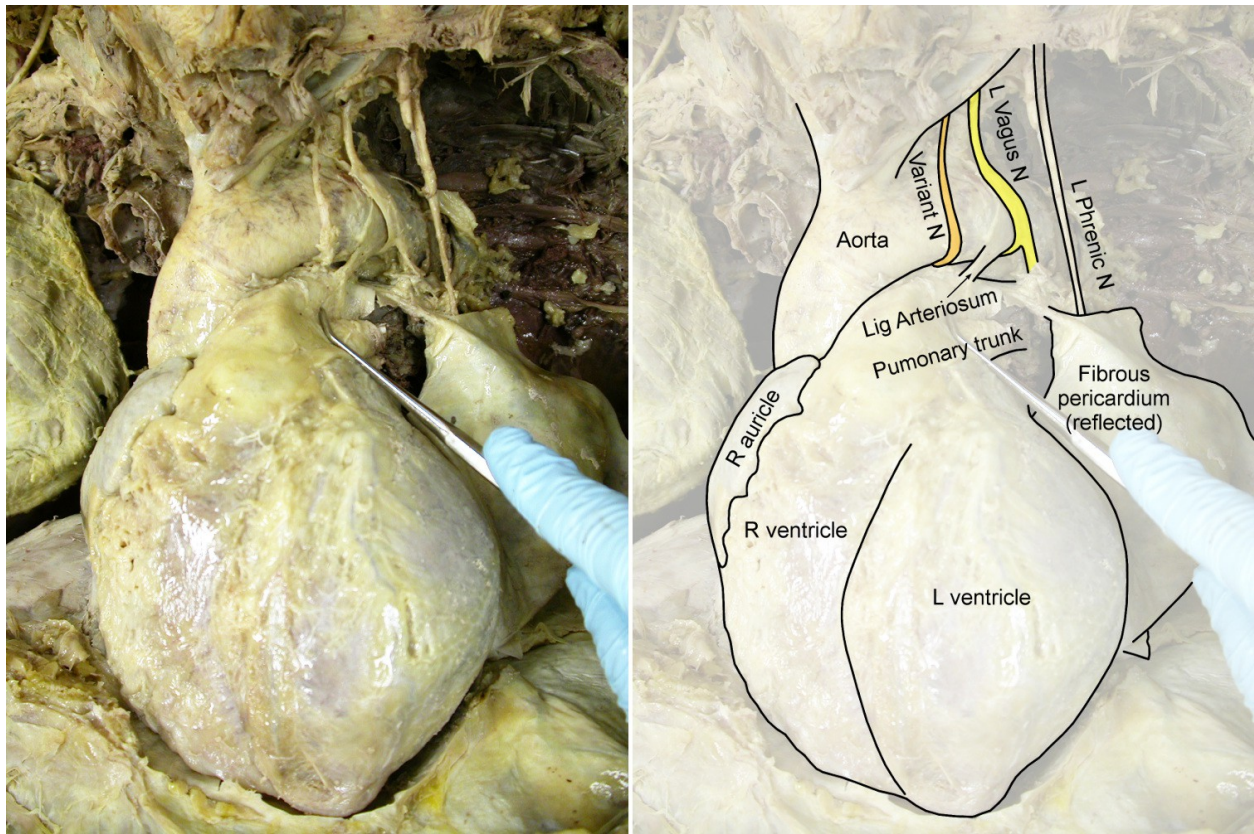
444 Yalcin B, Tugcu J, Canturk N, and Ozan H. "Laryngeal Branching Pattern of the Inferior
445 Laryngeal Nerve, Before Entering the Larynx." *Surgery Radiology Anatomy*. (2006); 28: 339-
446 342.

447

448 Yalcin B, Tunali S, and Ozan H. "Extralaryngeal Division of the Recurrent Laryngeal Nerve: A
449 New Description for the Inferior Laryngeal Nerve." *Surgery Radiology Anatomy*. (2008); 30:
450 215-220.

451 FIGURES AND FIGURE CAPTIONS

452



453

454 **Figure 1:** Discovery photo showing the variant nerve curving around the aorta.

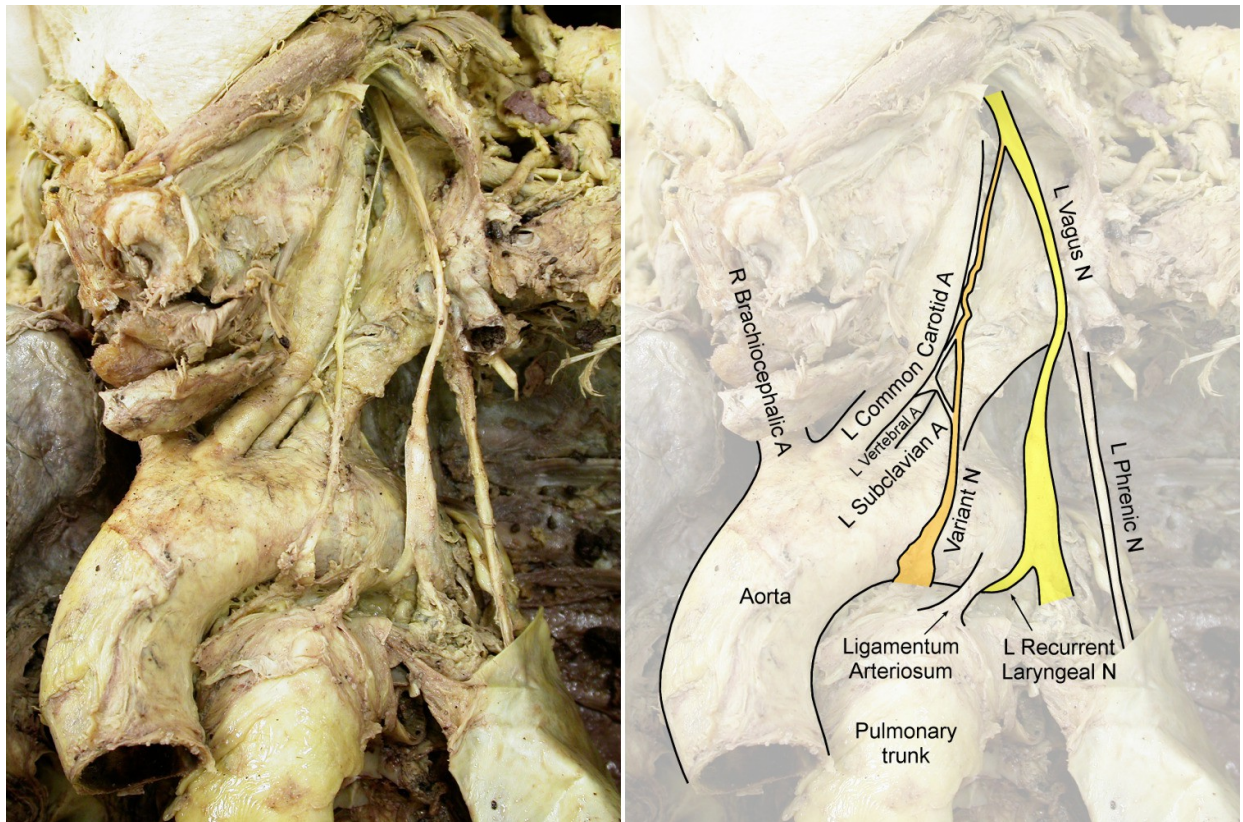
455 Photograph in ventral view showing the relationship of the variant nerve (orange) to the left

456 vagus nerve (yellow) and the great vessels. The left RLN is just visible branching from the vagus

457 nerve and passing behind the ligamentum arteriosum. The lungs have been removed and the

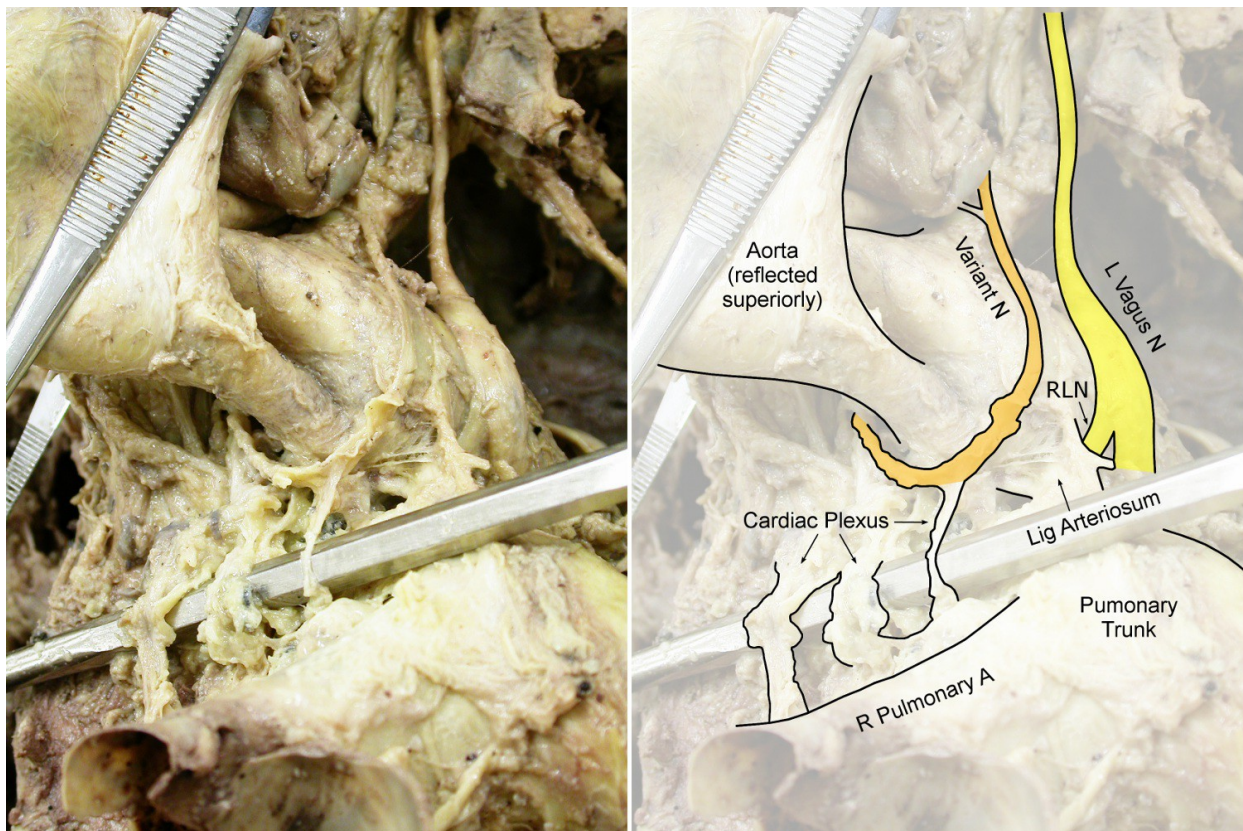
458 fibrous pericardium opened and laterally reflected.

459



460 **Figure 2:** The arch of the aorta, the variant nerve, and the normal RLN. Photograph in ventral
 461 view showing the proximal connection of the variant nerve (orange) to the left vagus nerve
 462 (yellow). In addition to the variant nerve, a minor vascular anomaly is also visible: the left
 463 vertebral artery branches directly from the arch of the aorta, between the left common carotid
 464 and left subclavian arteries. The heart has been removed.

465

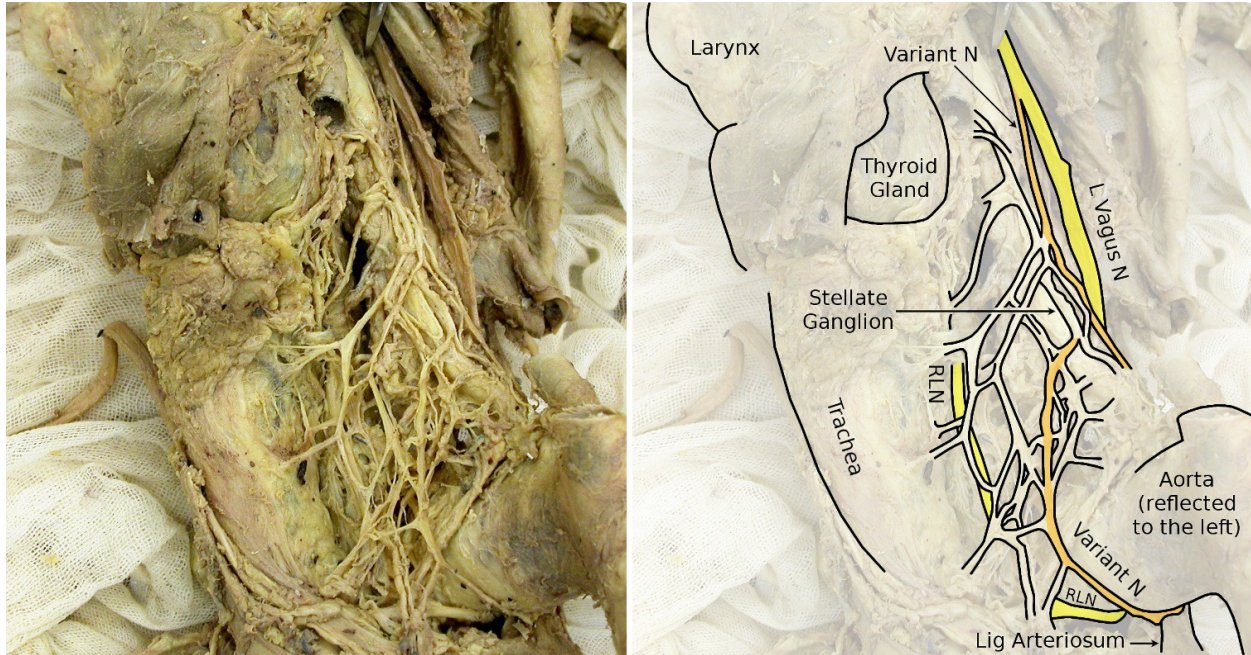


466

467 **Figure 3:** The recurrent course of the variant nerve around the aorta.

468 Photograph in ventral view with the aorta reflected upward to show the variant nerve wrapping
 469 around it. The probe is passing under nerve fibers of the cardiac plexus, at least one of which is
 470 originating from the variant nerve (in this and other figures, only the main trunk of the variant
 471 nerve is highlighted in orange in the interpretive diagram on the right).

472



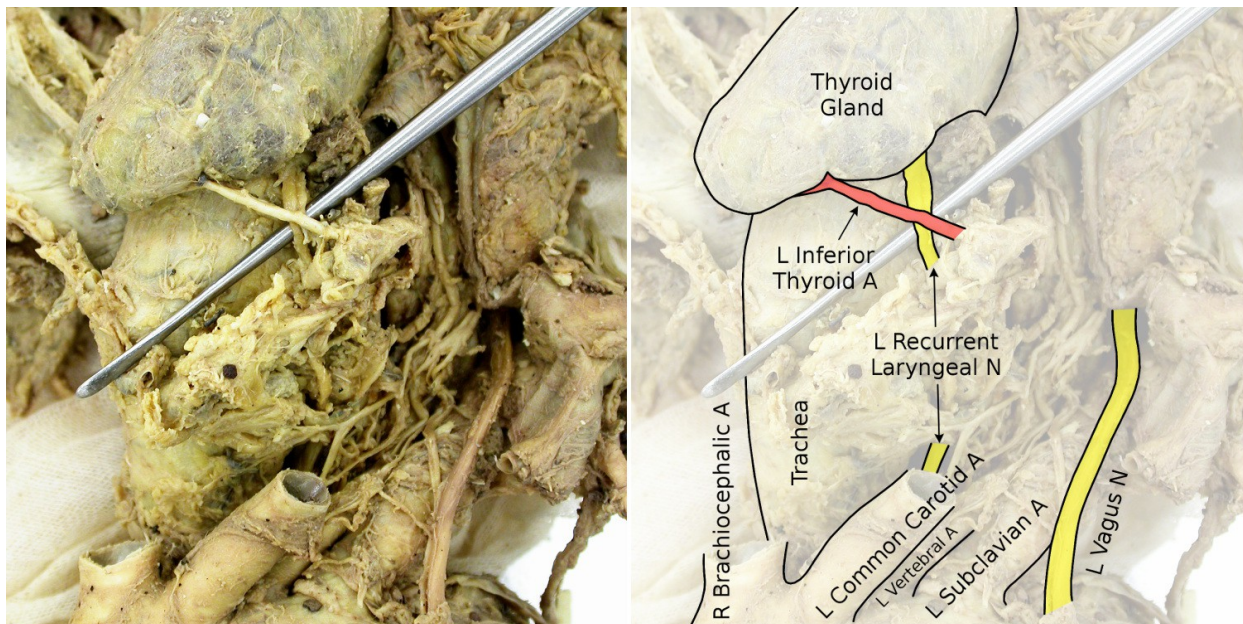
473

474 **Figure 4:** Connections of the variant nerve to the esophageal plexus.

475 Photograph of the root of the neck in left ventrolateral view showing the esophageal plexus. The
 476 variant nerve appears twice here: on the far right, descending from the left vagus nerve, and on
 477 the middle right, wrapping around the aorta to join the esophageal plexus. Also shown here is the
 478 connection of the variant nerve to the stellate ganglion (this pathway is highlighted in orange),
 479 and the normal left RLN (yellow) coursing behind the esophageal plexus on its ascending path to
 480 the larynx.

481

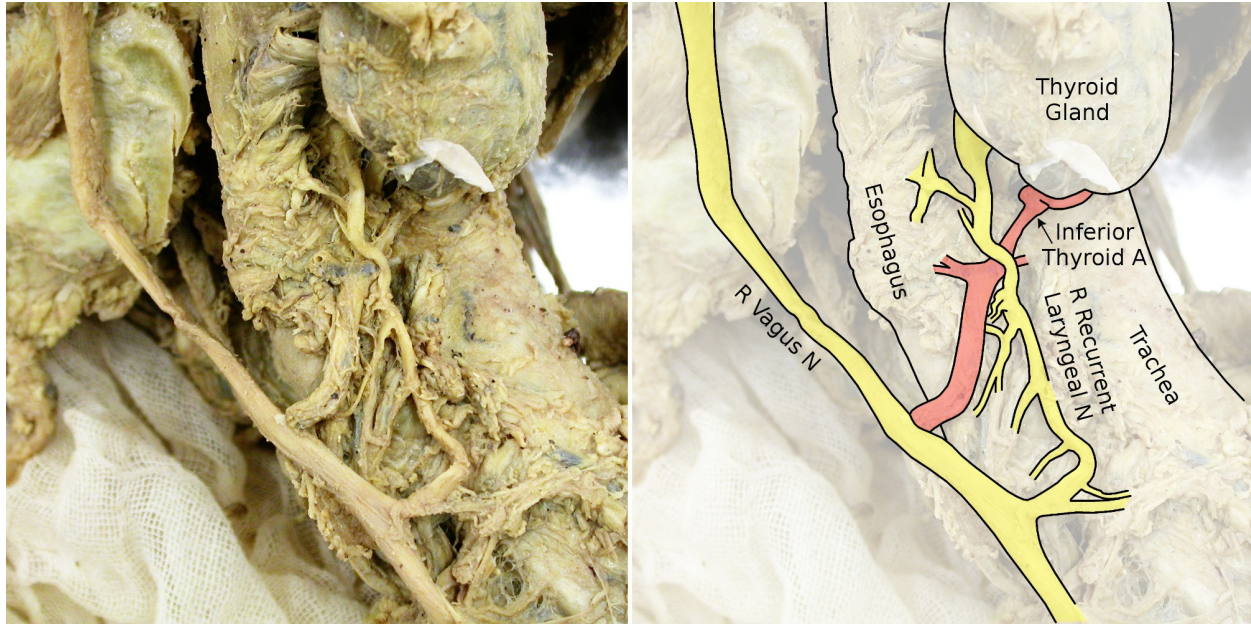
482



483 **Figure 5:** The left RLN passing behind the inferior thyroid artery.

484 Close-up photo of the root of the neck in left ventrolateral view. Compare the relationship of the
485 artery and the nerve on the left, as shown here, with their reversed relationship on the right,
486 shown in Fig. 6.

487



488

489 **Figure 6:** The course of the right RLN.

490 Close-up photo of the root of the neck in right ventrolateral view, showing the course of the right
491 RLN from the right vagus nerve to the point where it passes behind the thyroid gland. The
492 subclavian artery (which the right RLN wraps around before coursing superiorly) has been
493 removed; originally it sat at the bottom of the U-shape formed by the right vagus nerve and right
494 RLN. Note the numerous branches from the right RLN to the esophagus. Also note that on this
495 side the RLN passes in front of the inferior thyroid artery, in contrast to their relationship on the
496 left, shown in Fig. 5.

497



498

499 **Figure 7:** The anastomosis of Galen between the superior and recurrent laryngeal nerves on the
 500 right.

501 Close-up photo of the piriform recess inside the larynx in posterior view; superior is to the right.

502 On the far right, the internal laryngeal nerve pierces the muscular wall of the larynx and enters
 503 the piriform recess, where its fibers become intertwined (middle) with those of the internal
 504 branch of the recurrent laryngeal nerve (left). This connection is known as the ramus

505 anastomoticus or Galen's anastomosis. In this individual it was only present on the right side of
 506 the larynx; the left SLN and RLN did not share any visible connections.