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Evaluation of the frequency of left renal vein variations in computed tomography and its relationship with cancer development

Frequency of left renal vein variations and cancer development

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### Abstract

**Background:** Left renal vein (LRV) variations occur in 0.8%–10.2% of the population. The most common LRV variations are retroacrtic left renal vein (RLRV) and circumacrtic left renal vein (CLRV). The purpose of this study is to determine the frequency of LRV variations in a large series on computed tomography (CT) and to investigate the association between LRV and malignancy development.

**Materials and methods:** Between January 2015 and January 2017, an abdominal CT examination of 12,341 (5,505 female, 6,836 male) patients was evaluated retrospectively in this study. Patients clinical and demographic data were recorded using the Hospital Information System.

**Results:** LRV variations were detected in 314 (2.54%) of the 12,341 patients within the study. Of the 314 cases found to have LRV variations, 227 (1.84%) had RLRV, and 87

(0.70%) had CLRV. There was no statistical difference in total LRV variations (p = .083) and CLRV variation (p = .96) groups in terms of gender. However, the RLRV variation was found to be 1.32 times higher in males than in females (p = .039). Of the 314 patients with LRV variations, 73 (23.2%) had any sort of concomitant malignancy.

**Conclusions:** A high incidence of malignancy was detected in patients with LRV variations. Of the LRV variations, RLRV variation is more common than CLRV variation. The presence of total LRV variations and CLRV variations is not associated with gender. Whereas the presence of RLRV variation is more common in the male gender.

Key words: renal vein, variation, abnormality, malignancy, computed tomography, circumaortic, retroaortic

### INTRODUCTION

Inferior vena cava (IVC) and renal vein variations originate from the anomalies in the anastomoses of the three paired veins, including subcardinal, supracardinal and postcardinal veins, during embryonic development, and they may have numerous forms (1,2). Normally, a single left renal vein (LRV) drains into the IVC by crossing anteriorly over the abdominal aorta in the majority of cases. However, the anomalies arising in the LRV during the complex embryogenesis lead to the formation of variations (3–5).

LRV variations occur in 0.8%–10.2% of the population, although it varies depending on the research method (Table 1). The most common LRV variations are retroaortic left renal vein (RLRV) and circumaortic left renal vein (CLRV) (Figure 1, 2). With recent developments in imaging techniques and frequent use of imaging modalities, these abdominal vein variations are identified more frequently during routine examinations. Although it is known that the detection of LRV variations is clinically important for both surgical and diagnostic reasons, the relationship between these variations and malignancy is not known (4–8).

In this study, we aim to determine the frequency of LRV variations in a large series on computed tomography (CT) and to investigate the association between LRV and malignancy development.

### MATERIALS AND METHODS

# **Patient characteristics**

Approval was obtained from the Marmara University Ethics Committee (No: 09.2018.347) before the study began. The clinical and demographic data of cases were obtained from the hospital information management system and patient files.

The patients abdominal CTs, which were taken for any reason during the two-year period between January 2015 and January 2017 in Marmara University Pendik Training and Research Hospital, were retrospectively reviewed. Patients whose vascular structures could not be evaluated due to previous abdominal surgery or intraabdominal mass were not included in the study. Only one abdominal CT examination per patient of those who has had multiple CT examinations was included in the study. In total, abdominal CTs of 12,341 patients, which were evaluated by a radiologist with 20 years of abdominal radiology experience, were included in the study.

# **CT** protocol and image interpretation

Examinations were carried out with 256-slice CT scanner or 16-slice CT scanner devices. The obtained axial images were loaded into a Picture Archiving Communication System (PACS) workstation and the imaging findings were evaluated using the axial slices and, if necessary, the reformatted multiplanar reconstruction (MPR) images. The image analysis was performed by the same radiologist using consecutive axial CT images. The course of the LRV was shown uninterruptedly from the kidney hilus to the IVC. An LRV that drains into the IVC by crossing anteriorly over the aorta is accepted to be normal, whereas an LRV draining into the IVC by passing posteriorly over the aorta is defined as RLRV. An LRV complex that drains into the IVC by passing anteriorly and posteriorly around the aorta and forming a venous circle was defined as CLRV.

# Statistical analysis

Descriptive analysis was carried out for the case characteristics. The continuous variables were reported as mean  $\pm$  standard deviation. The rates were expressed in numbers and corresponding percentages. The percentages are indicated by confidence intervals (CIs). The data obtained were analyzed using the SPSS 17.0 statistical package program (SPSS Inc., Chicago, IL, USA), and the statistical analysis was assessed at a significance level of p < .05. The correlations of the categorical variables were determined using the chi-square test. The male and female cases were compared using probability rates and CIs.

### RESULTS

LRV variations were detected in 314 (2.54%) of the 12,341 cases within the study. Of these cases, 5,505 (44.6%) were female, 6,836 (55.4%) were male and the median age was 46.5 (distribution: 4–88 years). Of the 314 cases found to have LRV variations, 227 (1.84%) had RLRV, and 87 (0.70%) had CLRV.

In females, the total number of cases with LRV variations, RLRV variation and CLRV variation (with corresponding percentages) was 125 (2.27%), 86 (1.56%) and 39 (0.71%), respectively. In males, the total number of cases with LRV variations, RLRV variation and CLRV variation (with corresponding percentages) was 189 (2.76%), 141 (2.06%) and 48 (0.70%), respectively. The frequencies and distribution of LRV variations are given in Table 2, 3.

The correlation of LRV variations with gender was determined using the chi-square test. No statistically significant correlation was found between gender and total LRV variations (p = .083). CLRV variation was found to be approximately equally frequent in both males and females. Likewise, no statistically significant correlation was found between CLRV variation and gender (p = .96). However, the RLRV variation was found to be 1.32 times higher in males than in females (95% CI, 1.01–1.72), which was statistically significant (p = .039). Thus, the RLRV variation was determined to be related to gender.

Of the 314 patients with LRV variations, 73 (23.2%) had any sort of concomitant malignancy, the most common of which were gastrointestinal system (GIS) malignancy,

which appeared in 27 (37.0%) cases and hematological malignancy which appeared in 23 (31.5%) cases (Table 4). There was no statistical difference in the retroaortic LRV and circumaortic LRV groups in terms of gender (p = .269) and malignancy frequency (p = .59).

# **DISCUSSION**

The development of renal veins and IVC are highly complex processes associated with each other. This process covers the period between the fourth and eighth weeks of the embryogenesis, and consequently, IVC develops from a network of three parallel vascular pairs, including posterior cardinal, lower cardinal, and supracardinal vessels, in order of appearance. During this period, anastomotic communication occurs between the subcardinal and supracardinal canals with a vascular collar surrounding the anterior and posterior aorta. The dorsal arch of the circumcortic collar regresses and the progression of the ventral arch of the circumcortic collar forms a normal left renal vein. Conversely, the RLRV is caused by the persistence of the dorsal arch of the renal collar and the regression of the ventral arch, so that a single left renal vein passes through posterior to the aorta. CLRV results from the persistence of both the dorsal and ventral arches of the renal ring so that the left renal veins pass through both anterior and posterior to the aorta (1,3).

Although the clinical and imaging findings of the mesoaortic compression of the left renal vein (nutcracker phenomenon) are well described in the literature, there is limited information on RLRV and CLRV. With the increased use of imaging modalities, the identification of these variations has become clinically important for both surgical and diagnostic reasons. Recognition of vascular variations before renal venous sampling, preoperative evaluation of renal donors, splenorenal shunt placement, vena cava filter placement, nephrectomy, aortic surgery, and staging of renal cell carcinoma is crucial to preventing misdiagnosis and complications, such as retroperitoneal hemorrhage, nephrectomy, and even death. Recognition of these variations is also important in the differential diagnosis of retroperitoneal pathology (2,5,9).

Even though abdominal venous structures can be evaluated with different radiological modalities, such as CT, CT angiography, Doppler, MR and ultrasonography (USG), CT is the preferred imaging modality for the evaluation of abdominal venous vascular structures due to higher patient compliance, lower cost, fast and easy applicability and reliability (8,10–13).

The prevalence of RLRV and CLRV was reported to be 0.5-7.4% and 0.3-6.3%, respectively, in a newer various studies conducted by using autopsy series, CT, CT angiography, magnetic resonance, and PET-CT (4,5,14–16,6–13). The reported incidence of LRV variations varies widely (Tables 3). The high difference between the study results may be due to the different modalities of the studies, patient characteristics included in studies or the number of patients. However, in general, the incidence of LRV variations is around 2%.

Our study included the CT images of 12341 patients which is one of the most comprehensive studies in the literature in terms of investigation of LRV variations. Similar to previous studies in the literature, the incidences of LRV, RLRV, and CLRV variations were also found to be 2.27%, 1.56%, and 0.71%, respectively, in our study. The results of our study show that the incidence of total LRV variations in the Turkish population is similar to that of other races (6,7,9,16,17).

Similarly to our results, most of the studies reported that the incidence of RLRV variations was higher than that of CLRV variations (4,8,10,13–16). However, some studies found that the incidence of CLRV variations higher that RLRV variations (5,7,9,11). RLRV variations are less complex than CLRV variations. This may be due to different embryological development processes of these renal vein variations.

While there was no gender correlation regarding LRV in some previous studies, Dilli et al. found that RLRV was more common in females than in males (6,13–15). In our study, there was no correlation between total LRV and CLRV; and gender. In contrast to Dilli et al., we found that the incidence of RLRV was statistically significantly higher in males than in females. This difference between the studies may be due to differences in the number, characteristics of patients included in the studies or due to differences in investigation modalities.

Interestingly, there was a concomitant malignancy in 73 (23.2%) of 314 cases with LRV variations in our study. The majority of concomitant malignancies were gastrointestinal system (37.0%) or hematologic (% 31.5) malignancies. Of the 27 (37.0%) concomitant gastrointestinal system malignancies, 11 (15.1%) were gastric cancer, 7 (9.6%) were colon cancer, 6 (8.2%) were pancreatic cancer, 2 (2.7%) were pancreatic cancer and 1 (1.4%) was esophageal cancer. Of the 25 (31.5%) accompanying hematologic malignancies, 9 (12.3%) were Hodgkin's lymphoma, 7 (9.6%) were Non-Hodgkin's lymphoma and 7 (9.6%) were other hematologic malignancies (Table 4). There is no comprehensive information in the literature other than a few cases with left renal variation accompanied by malignancy (18,19). More extensive studies are needed to demonstrate whether there is a correlation between the embryological development of the renal vein variation and malignancy development.

Our study has some limitations. Abdominal CT examinations of all the cases included in the study were performed for any reasons such as trauma scan, diagnosis of abdominal pathology, and cancer screening or follow-up, however, the results of the study cannot be generalized to the entire population since healthy people were not included in the study because of the radiation risk. Despite that, the size of the study population is the main strength of this study.

# **CONCLUSIONS**

In conclusion, CT is a fast and reliable method for detecting renal vein variations. Of the LRV variations, RLRV variation is more common than CLRV variation. The presence of total variations in LRV and CLRV is not associated with gender. Whereas the presence of RLRV variations is more common in the male gender.

The incidence of left renal vein variation is above 2% and incidentally identified vascular variations during routine CT scans should always be reported since they are of clinical and surgical importance. During routine abdominal CT reporting, it should be kept in mind that left renal vein variation will be detected in approximately one of every 50 patients, and if not detected, the renal vascular structures should be evaluated more carefully.

Because of the high rate of malignancy detected in patients with LRV variations in our study, we suggest that patients with LRV variations in imaging modalities should be carefully evaluated especially for gastrointestinal and hematologic malignancies.

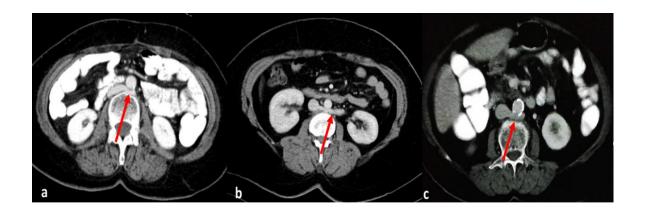
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**Fig 1.** Circumaortic left renal veins in a 66-year-old male patient (a and b) with colon cancer, a 20-year-old male patient (c and d) without concomitant malignancy and a 56-year-old female patient (e and f) with non-Hodgkin lymphoma. Contrast-enhanced CT scans show the left renal veins coursing anterior (red arrows in a, c, e) and posterior (red arrows in b, d, f) to the aorta.



**Fig 2.** Retroortic left renal veins in a 62-year-old female patient (a) with colon cancer, a 57-year-old male patient (b) without concomitant malignancy and a 79-year-old female patient (c) with non-Hodgkin lymphoma. Contrast-enhanced CT scans show the left renal veins (red arrows in a, b, c) coursing posterior to the aorta.

**Table 1.** Frequency of left renal vein variations in studies using different methods

Study (Year)	Method	Sample size	RLRV	CLRV	Total LRV
1. (2008)(9)	CT	153	0.7%	1.3%	2%
2. (1982)(7)	CT	433	1.8%	4.4%	6.2%
3. (2016)(11)	PET-CT	222	2.70%	3.15%	5.85%
4. (2013)(14)	CT	1204	3.1%	2.1%	5.2%
5. (2007)(5)	CT	1120	4.7%	5.5%	10.2%
6. (2012)(13)	MRI	2644	1.66%	1.02%	2.68%
7. (2017)(8)	MRI	3000	2.1%	0.5%	2.6%
8. (2017)(16)	Autopsy	550	4.2%	3.8%	8.0%
9. (2014)(10)	CT	746	7.4%	2.4%	9.8%
10. (2015)(6)	CT	1452	2.1%	2.1%	4.2%
11. (2004)(15)	CT	984	2.3%	0.9%	3.2%
12. (1999)(4)	Autopsy and in-vivo	1008	0.5%	0.3%	0.8%
13. (2016)(12)	CT Ang	504	4.2%	5.2%	9.7%

CT: Computed Tomography; CT Ang: Computed Tomography angiography; MR: Magnetic Resonance Imaging; PET-CT: Positron emission tomography—computed tomography

**Table 2.** The frequencies and distribution of left renal vein variations in cases

	Total (n = 12341), n of patients (%)	Females (n = 5505), n of patients (%)	Males (n = 6836), n of patients (%)
Retroaortic left renal vein variations	227 (1.84%)	86 (1.56%)	141 (2.06%)
Circumaortic left renal vein variations	87 (0.70%)	39 (0.71%)	48 (0.70%)
Total Left renal vein variations	314 (2.54%)	125 (2.27%)	189 (2.76%)

**Table 3.** The frequencies and distribution of retroaortic and circumaortic left renal vein variations

Variation		Ger	nder	
		Female	Male	Total
Circumaortic left renal vein	Count	39	48	87
	% within CLRV	44.8%	55.2%	100.0%
	% within gender	31.2%	25.4%	27.7%
	% of Total	12.4%	15.3%	27.7%
Retroaortic left renal vein	Count	86	141	227
	% within RLRV	37.9%	62.1%	100.0%
	% within gender	68.8%	74.6%	72.3%
	% of Total	27.4%	44.9%	72.3%
Total left renal vein	Count	125	189	314
	% within LRV	39.8%	60.2%	100.0%
	% within gender	100.0%	100.0%	100.0%
	% of Total	39.8%	60.2%	100.0%

CLRV: Circumaortic left renal vein, RLRV: Retroaortic left renal vein, LRV: left renal vein

Table 4. Cancers distribution according to location/system in patients with left renal vein variation

	Number	% of Total
Digestive/Gastrointestinal	27	37.0
Stomach Cancer	11	15.1
Colon Cancer	7	9.6
Pancreatic Cancer	6	8.2
Rectal Cancer	2	2.7
Esophageal Cancer	1	1.4
Hematologic/Blood	25	31.5
Non-Hodgkin lymphoma	9	12.3

Hodgkin lymphoma	7	9.6
Hematology malignancy	7	9.6
Others		
Lung Cancer	4	5.5
Genitourinary Cancer	4	5.5
Gynecologic Cancer	3	4.1
Breast Cancer	3	4.1
Miscellaneous	9	12.3
Total	73	100.0