ORIGINAL RESEARCH Safety and Intraoperative Results in Live Kidney Donors with Vascular Multiplicity After Hand-Assisted Laparoscopy Living Donor Nephrectomy

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Background and Purpose: Vascular multiplicity is the most frequent anatomic variation in kidney donors. Despite concerns about risks, these allografts are increasingly used to overcome the shortage of kidney donors. The safety and clinical outcomes in living kidney donors were evaluated with vascular multiplicity after hand-assisted laparoscopic living donor nephrectomy (HALDN).

Patients and Methods: Data from all living kidney donors who underwent HALDN from 2008 to 2021 was retrospectively reviewed. Patients were divided into two groups as single (SRV) and multiple renal vessels (MRV), and a comparative analysis was done. The primary outcomes include operating room time (ORT), days of hospital stay, estimated blood loss, complications, conversion, and re-operations.

Results: MRV were present in 166 out of 612 donors (27.1%). Among those, 10 (1.6%) donors had simultaneous multiple arteries and veins. Additionally, the prevalence of artery and vein multiplicity was 21.8% (n = 134) and 3.5% (n = 22), respectively. Warm ischemia time was significantly different among the two groups but not clinically important. The number of conversions to open technique, the mean ORT, the median blood loss, and days of hospital stay were similar between the SRV and MRV groups, without significant differences. According to the modified Clavien-classification system, no differences were found in the complication rates between the two groups (p = 0.29). Complication rates were 3.3% and 3.6% for the SRV and MRV groups, respectively.

Conclusion: HALDN is a procedure with safe intraoperative results, even with vascular multiplicity. The presence of multiple renal arteries or veins has no negative impact on the outcome of the donor after living donor nephrectomy.

Keywords: kidney transplant, vascular multiplicity, living kidney donors, laparoscopic nephrectomy, anatomic variation

Introduction

Living kidney donation is crucial to increase the donor pool in renal transplantation.¹ However, healthy or "ideal" living donors are not enough to accomplish a sufficient donation rate. Extended criteria for living kidney donors with obesity, hypertension, incompatibility blood group, vascular multiplicity, and older donors have been accepted in kidney transplant groups to expand opportunities for patients with chronic kidney disease on the waiting list.²

Research and Reports in Urology 2022:14 23-31

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Simple vascular anatomy is preferred in kidneys of living donors^{3–5} because renal vascular multiplicity could lead to a technical difficulty in anastomosis and potential secondary complications in the donor and the recipient. Contrarily, accepting these allografts to overcome a shortage of renal donors had trended worldwide. According to the British guidelines for living donor kidney transplantation, kidneys with vascular multiplicity are not an absolute contraindication for donation, and decisions should be made on an individual clinical evaluation.⁶ Nevertheless, other guidelines did not make any recommendation regarding vascular multiplicity in live kidney donation.²

Vascular multiplicity is due to the anatomical variations in the mesonephric vessels during the first trimester of fetal life.^{7–9} Kidneys with vascular multiplicity are common anatomical findings in kidney donors (18– 30%).^{10–12} A review showed an overall prevalence for two renal arteries, three renal arteries, and bilateral multiple renal arteries of 25%, 2.6%, and 10%, respectively.¹³ The prevalence of multiple renal veins varies between 3% and 14%.^{3,13,14} Diverse vascular renal anomalies can be classified by their distribution of origin.⁹

results Previous study vary. For instance, a retrospective study with 951 live kidney donors who underwent hand-assisted laparoscopic living donor nephrectomy (HALDN) found higher warm ischemia time with no clinical impact on donors with multiple renal arteries compared to donors with a single renal artery. Additionally, no significant differences were found in complications, conversion to open surgery, calculated blood loss, length of hospital stay, re-interventions, and readmissions as compared in both groups.¹⁵ Contrarily, a publication with 1350 live kidney donors found a higher complication rate and conversion rate to open surgery in patients with vascular multiplicity than patients without vascular anomalies.¹⁶ The mortality rate in patients with vascular anomalies is similar to those without any variation.¹²

Most published studies described only the clinical outcomes in recipients or donors with renal vascular anomalies that report only one type of vessel variation (arterial or venous).^{10,17–20} Vascular multiplicity, including both venous and arterial anomalies, was analyzed. This study aimed to evaluate the safety and clinical outcomes in live kidney donors with single renal vessels (SRV) compared to live kidney donors with multiple renal vessels (MRV) undergoing HALDN.

Materials and Methods Study Design and Study Population

This is a retrospective observational cohort of all living kidney donors who are operated in Colombiana de Trasplantes (including a network of two centers: Bogotá and Barranquilla) from August 2008 to May 2021. During the study period, 612 consecutive kidney donors were included. No exclusions were set, and the electronic medical records of the total donors were reviewed. During this period, two surgeons performed most of the procedures (92.4%). The feasibility of the procedure was determined after preoperative multidisciplinary donor evaluation that included appropriate psychological, medical, and surgical evaluations.

Medical evaluation of all living donors was made by a multidisciplinary team. Computed tomography angiography (CTA) was performed to identify the renal vascular anatomy and recognize renal vascular variants. The policy of our center indicates no absolute contraindication for vascular multiplicity. Kidneys with more than three arteries are not taken as no extensive literature is available on donors with more than three renal arteries and there are no definitive conclusions in this regard. Thus, the results of the safety of these donors are best applied to living donors with up to three renal arteries.

Perfusion areas of small accessory vessels were assessed before ligating and prioritizing reconstruction techniques for renal arteries or veins to not affect kidney perfusion especially if inferior polar arteries are found. Right nephrectomy was chosen in cases of confirmed lower differential renal function of the right versus the left kidney in diethylene triamine pentaacetic acid (DTPA) renogram, even with an anatomic disadvantage.

Asymmetric kidneys were defined as differences that are >1 cm that affects the volume of the organ documented in CTA. In such a case, a DTPA renogram was ordered and should not show a difference of >10% in kidney function between both kidneys to be accepted as a donor. The transplant team decided which kidney will be removed to avoid additional risk to the donor and take the kidney with less function.

Live kidney donors should have estimated the glomerular filtration rate (GFR) of \geq 90 mL/min/1.73 m², donors with GFR between 60 and 89 mL/min/1.73 m² will be assessed based on age, demographics, and risk factor characteristics.²¹ Live kidney donors with GFR of <60 mL/min/1.73 m² are not accepted for donation.

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Two groups were defined according to the number of vessels that are present in the renal pedicle. Kidneys with a single artery and vein were defined as SRV group and those with more than one artery or vein were classified as MRV group. The primary outcomes were warm ischemia time, total operating room time (ORT), days of hospital stay, blood loss, complications, conversion, and re-operations. Variables of interest were analyzed during the immediate postoperative care and after the first day when patients were discharged.

Donor complications were classified according to Kocak et al, which is a modification of the Clavienclassification system, to describe complications after live donor nephrectomy.²²

Operative Procedure

The HALDN was the technique of choice. Patients were positioned in a "flank-up" position and were secured with adhesive tapes. A hand port was placed through a transverse infraumbilical or midline incision. Two trocars (5 and 12 mm) and a 30° video endoscope were introduced. The pneumoperitoneum level was set by infusing CO2 with a flow rate of 400 cm/min up to a maximum of 15 mmHg of intra-abdominal pressure. The colonic splenic flexure mobilization was conducted using the ultracision (HARMONIC ® HD ultracision Johnson 1000i). Renal artery vessels and ureters were identified and dissected. Renal vessels were clamped using two large size non-absorbable polymer ligating clips (Weck[®] Hem-o-lok[®]) and transected.²³ Any secondary renal arteries were also carefully dissected until the aorta. The use of a 60 mm endovascular cutting stapler obtains a reasonable vessel length to mobilize and remove the right kidney. The kidney is immediately withdrawn to minimize the warm ischemia time. The kidney was delivered out through the hand port. A laparoscopic inspection was done to check for hemostasis. Urine output was maintained using intravenous hydration. Unfractionated heparin was not used before clamping the vessels. The postoperative protocol included urine catheter removal, regular diet after the first day, and analgesia on-demand.

Statistical Analysis

Descriptive and comparative analysis was carried out according to the two defined groups (SRV and MRV). Frequencies and percentages were used to describe categorical variables. Central tendency and dispersion measures were used to describe quantitative variables. Comparisons between the two groups for the main outcomes (days of hospital stay, ORT, blood loss, complications, conversion, and re-operations) were analyzed using the Chi-square and Mann–Whitney test. A *p*-value of <0.05 was accepted as statistically significant. Analysis was performed using the Software R version 4.0.3.

Ethical Considerations

The study was approved by the Ethics Committee Dexa Diab. This retrospective research does not present any risk according to the ethical considerations that were established in the National Regulations, such as resolution number 8430 of 1993, published by the Ministry of Health in Colombia²⁴ and International Regulations, such as the Declaration of Helsinki²⁵ and the Declaration of Istanbul.²⁶ We confirmed that all kidneys were voluntarily donated with written informed consent and that this was conducted according to the Declaration of Istanbul.²⁶

Results

Donor Characteristics and Clinical Outcomes

During the study period, 612 patients had HALDN with a median age of 37.6 years, wherein 324 (52.9%) were females and 544 (88.8%) had left nephrectomy. The mean body mass index was 24.9 m²/kg. Previous abdominal surgery was determined in 262 (42.8%) patients (Table 1). The mean warm

Table I Dem	nographic Data	of Donors	with and	without	Multiple	Renal \	/essels
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	All (n = 612)	Donors with SRV (n = 446)	Donors with MRV (n = 166)	p-value
Age (years)	37.6 (11.1)	37.6 (11.1)	37.6 (11.2)	0.9543
Gender (%)				0.6996
Female	324 (52.9)	234 (52.4)	90 (54.2)	
Male	288 (47.1)	212 (47.6)	76 (45.8)	
Right/left kidney	68/544	44/402	24/142	0.1080
Body mass index (SD)	24.9 (3.4)	24.9 (3.4)	25.0 (3.5)	0.7422
Previous abdominal surgery (%)	262 (43.4)	187 (42.8)	75 (45.2)	0.5970

Abbreviations: SRV, single renal vessels; MRV, multiple renal vessels; SD, standard deviation; IQR, interquartile range.

Table 2 Clinical Outcomes of Donor	s with and without Multiple Renal Vessels
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	All (n = 612)	Donors with SRV (n = 446)	Donors with MRV (n = 166)	p-value
Mean warm ischemia time, min (SD)	3.3 (2.4)	3.1 (1.2)	3.8 (4.0)	0.0269
Mean operative time, hour (SD)	1.9 (0.9)	1.9 (0.8)	2.0 (1.1)	0.0751
Median blood loss, cc (IQR)	50 (50)	50 (30)	50 (50)	0.1505
Number of conversion (%)	5 (0.8)	2 (0.45)	3 (1.7)	0.0968
Hospital stay, days (SD)	2.2 (1.7)	2.2 (1.9)	2.1 (0.6)	0.1775

Abbreviations: SRV, single renal vessels; MRV, multiple renal vessels; SD, standard deviation; IQR, interquartile range.

	Single Renal Artery (n = 468)	Double Renal Arteries (n = 128)	Triple Renal Arteries (n = 16)	p-value
Mean warm ischemia time, min (SD)	3.1 (1.2)	3.8 (4.5)	3.9 (2.1)	0.0069
Mean operative time, hour (SD)	1.9 (0.8)	2.1 (1.2)	2.1 (0.5)	0.0962
Median blood loss, cc (IQR)	50 (30)	50 (50)	67.5 (50)	0.0919
Number of conversion (%)	2 (0.04)	2 (1.5)	I (6.25)	0.0225
Hospital stay, days (SD)	2.2 (1.9)	2.0 (0.4)	2.1 (0.5)	0.509

Abbreviations: SRV, single renal vessels; MRV, multiple renal vessels; SD, standard deviation; IQR, interquartile range.

ischemia time was significantly different among the two groups but not clinically important. The number of conversions to open technique, the mean ORT, the median blood loss, and days of hospital stay were similar between the SRV and MRV groups, without significant differences (Table 2).

Renal Artery and Vein Evaluation

Vascular multiplicity was present in 166 (27.1%) donors, of whom 10 (1.6%) had simultaneous multiple arteries and veins. Additionally, the prevalence of artery and vein multiplicity was 21.8% (n = 134) and 3.5% (n = 22), respectively. Among the donors with simultaneous multiple renal arteries and veins, 9 (5.4%) patients had two arteries plus two veins and one patient (0.06%) had two arteries plus three veins (Table 2). Among the remaining 156 patients with vascular anomalies, 118 (71.0%) had two arteries, 16 (9.6%) had three arteries, 19 (11.4%) had two veins, and 3 (1.8%) had

Table 4 The Prevalence of Single and Multiple Renal Artery/Arteries in 612 Living Donors

	No. of Pa	No. of Participants		
No of Renal Arteries	Right Kidney (n = 68)	Left Kidney (n = 544)		
1	51 (75.0)	417 (76.7)		
2	16 (23.5)	112 (20.6)		
3	I (0.15)	15 (0.28)		

three veins. The comparison of clinical outcomes between single, double, and triple renal arteries revealed that warm ischemia time was significantly higher in the kidneys with triple renal arteries (Table 3).

The number of kidneys with single and multiple renal arteries and veins are described in Tables 4 and 5, respectively.

Perioperative and Postoperative Complications

No differences were found between the two groups according to the modified Clavien-classification system (p = 0.29) that was proposed by Kocak et al²² (Table 6). Complication rates were 3.3% and 3.6% for the SRV and MRV groups, respectively. Figure 1 shows the frequency of complications and conversions among donors. Renovascular complications were present in seven donors, including anatomical variant injuries (n = 3), renal artery injury (n = 1), renal vein injury

Table 5 The Prevalence of Single and Multiple Renal Vein/Veinsin 612 Living Donors

	No. of Participants		
No of Renal Veins	Right Kidney (n = 68)	Left Kidney (n = 544)	
1	57 (83.8)	523 (96.1)	
2	9 (13.2)	19 (0.3)	
3	2 (0.3)	2 (0.04)	

Grade ^a	Description	Donors with SRV (n = 446)	Donors with MRV (n = 166)
I	Epigastric vessel injury	I (0.22)	None
2a	Anatomical variant injury	2 (0.44)	(0.6)
	Renal vein injury	I (0.22)	None
2b	Colon injury	2 (0.44)	I (0.6)
	Duodenal injury + suture	I (0.22)	None
	Renal vessels injury	None	I (0.6)
	Serosal colon injury	2 (0.44)	I (0.6)
	Splenic capsule injury	3 (0.67)	None
2c	Renal artery injury + conversion	None	I (0.6)
	Splenic injury + conversion	None	I (0.6)
	Renal vein injury + conversion	I (0.22)	None
3	Splenectomy	I (0.22)	None
Percentage of total series (p = 0.1421)		3.1%	3.6%

Notes: ^aGrades according to Kocak et al.²²

(n = 1), and renal vessel injuries (n = 1). Among those, four occurred in donors with SRV, and three occurred in donors with MRV. The overall conversion to open surgery rate was 0.8% (5 cases). One of those conversions was due to anatomical identification and dissection difficulty. Two patients had a conversion to open surgery in donors with SRV and three patients had a conversion to open surgery in donors with MRV. No deaths or re-operations were revealed among the donors (Table 6).

has been adopted by several institutions since 1995 when Ratner et al²⁷ reported their first experience, thus multiple advantages, such as lower hospital stay, shorter time of recovery, and better aesthetic results,²⁸ have been documented. Our preferred operation employs a hand-assisted approach with some potential advantages over LDN, including shorter operative time, shorter learning curve, and the ability to manually assist in dissection.^{29,30}

Discussion

As the demand for kidney transplantation continues to rise, live kidney donation becomes crucial for expanding the donor pool. The laparoscopy donor nephrectomy (LDN) Potential live donors with multiple vessels are common; however, no guidelines regarding living kidney donation with renal vascular multiplicity are available, and its exclusion should be made by individual transplant centers. Renal vascular multiplicity presents a significant technical challenge as it may affect both donor safety and



Year of HALDN

Figure I Frequency of complications and conversions among donors.

recipient outcome.² Most studies that report vascular multiplicity focus on recipient results, and evaluating these results in the donor seems important since it exposes an otherwise healthy patient to the risks of surgery entirely for the benefit of another person. Therefore, very little literature is found regarding donor outcomes.

This study provides an analysis and overview of data of a large cohort of live kidney donors over 12 years. This study revealed that kidneys with multiple vessels are common. Moreover, no significant differences were found in variables that would affect donor outcomes such as length of stay, operative time, the likelihood of reoperation, conversion, blood loss, and complication rate.

The incidence of supernumerary renal veins varies but is frequently seen more on the right side.³¹ Our study revealed 16.1% of donors on the right side and 3.8% on the left side. During nephrectomy, the vein is sacrificed when venous drainage of the smaller vein is <20%. Moreover, side-to-side anastomosis is done if two veins are of equal caliber. The incidence of accessory renal arteries fluctuates from 4% to 61.5% in a Malaysian population and Brazilian population, respectively.³² Our cohort revealed that 23.5% of kidneys have supernumerary renal arteries. Clinically, the vertical trajectory of the upper or lower polar supernumerary renal artery can lead to polar infarction and can be injured during mobilization. Therefore, the anatomical knowledge of supernumerary renal arteries is essential before performing the surgery.³³

Previous studies that compared the outcomes in patients with SRV and MRV have shown variable results. A systematic review of 24 comparative studies that were published between 1970 and 2016 that was conducted by Afriansyah et al³⁴ showed no difference in donor outcomes with multiple renal arteries compared to with single renal artery kidney.

Regarding the operative time, some centers have seen longer operating times. Troppmann et al³⁵ reported longer operative times; however, this difference was not statistically significant. Contrarily, a higher rate of kidneys with multiple renal arteries was reported. This high rate of multiple renal artery grafts may theoretically exert an adverse effect on the outcome.

Concerning the warm ischemia time, some centers had reported higher warm ischemia time in cases with arterial multiplicity.^{35,36} Carter et al³⁶ reported longer warm ischemia time in cases where multiple vessels required revascularization and could not be managed by simple ligation (pole accessory arteries of <2 mm and supplying <10% of the renal cortex).

The mean first warm ischemia time was significantly longer in donors with MRV; however, the impact on clinical outcome is minor. According to the literature, a duration of 4 min is considered to obtain renal grafts without vascular variants and between 4.5 and 5 min in kidneys with multiple arteries or veins.^{15,34} In comparison, some publications do not include the analysis for the donor warm ischemia time since a timeframe limit was not established, and it is not a relevant clinical factor in the donor outcomes.^{19,37} Our study revealed a short (<4 min) warm ischemia time; therefore, we hypothesized no negative impact on allograft function from a clinical perspective.

Our study revealed that live kidney donors who underwent previously abdominal surgery did not show worse outcomes than live kidney donors without abdominal surgery history. Previous abdominal surgery is not a contraindication to be a kidney live donor.³⁸

The modified Clavien gradient system classified nephrectomy complications in the live kidney donor with SRV of grade 1 until 10%, grade 2 until 2.8%, grade 3 until 7%, and grade 4 until 0.3%. A live kidney donor with MRV has a slight classification change in proportions (grade 1 until 6%, grade until 2 11%, grade until 3 9%, and grade 4 until 0.8%).¹⁸ Our classification of the perioperative complications by the modified Clavien system is similar to the results reported in the literature with HALDN and different donor nephrectomy techniques.^{22,39,40} Compared to our results, a retrospective study with 700 kidney live donors with HALDN calculated the modified Clavien gradient system with a higher incidence of grade 1 complications than our study, but the lower incidence in the other categories.¹⁹

Arpali et al¹⁶ revealed that the overall conversion to open surgery rate and renovascular complication rate were significantly higher in patients with multiple renal arteries. The total complication rate in our series is comparable with experiences from other centers that range between 3% and 13.5%.^{41–44}

Our conversion rate was lower than what is reported by other authors.^{29,44} Our study revealed that live kidney donors with MRV and >3 vessels had a higher conversion rate, and cases of conversion mainly cause renovascular injuries, which is comparable with other large series.²⁹

Finally, Hsu et al⁴⁵ revealed that the use of renal allografts with multiple renal arteries does not adversely affect the parameters such as blood loss, operative time, complication rate, or hospital stay. Additionally, most

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studies showed no differences in donor outcomes with arterial multiplicity compared to living kidney donors without vascular anomalies.^{12,46–49}

Previous studies included donors with three or fewer renal arteries.^{5,47,50,51} A systematic review that evaluated extended donor criterion, including vascular multiplicity, concluded that the presence of multiple renal arteries should not be a contraindication for live kidney donation. Nevertheless, as no literature was available on donors with four or more renal arteries conclusions were not made definitive in this regard.² Additionally, our study described the presence of multiple veins, whereas the majority of studies analyzed the impact of multiple arteries. Only included donors а few studies with venous anomalies.^{3,4,15,52} showing that laparoscopic donor nephrectomy is safe and feasible in patients with vein anomalies. Our cohort had only ten kidneys with simultaneous multiple arteries and veins and no negative impact on donor outcomes. However, to the best of our knowledge, no study has been conducted on the clinical outcomes of donors with simultaneous multiple arteries and veins.

This study has some limitations. First, information bias intrinsic to the retrospective design is possible. Second, a low number of renal allografts with rare anatomical variations (simultaneous multiple arteries and veins) was found, and no solid conclusions may be provided about the feasibility of kidney donation in these circumstances. Moreover, the potential donors with vascular multiplicity that were excluded early in the evaluation process were not analyzed. Only intraoperative and early complications after the first day when patients were discharged were accounted and other minor complications may be possibly missed, such as wound infections, hematomas, readmission, and constipation, among others. As only early outcomes were measured, alteration of operative technique, renal function and proteinuria were not documented, and those variables are relevant to determine the safety of the procedure.

Conclusion

In conclusion, HALDN is a procedure with safe intraoperative results, even with vascular multiplicity. The presence of multiple renal veins or arteries has not significantly affected the rate of adverse intraoperative donor outcomes. Thus, renal vascular multiplicity should not be a contraindication to becoming a kidney donor. Furthermore, accepting extended criteria live kidney donors has great potential to reduce the kidney donor scarcity and improve the opportunity for transplantation and potentially reduce the mortality of those on the waiting list.

Disclosure

The authors report no conflicts of interest in this work.

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