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## Impact of thermal denaturation on renal volume reduction after partial nephrectomy using soft coagulation hemostasis

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Partial nephrectomy has become the gold standard treatment for small renal masses. This study aimed to assess the impact of soft coagulation hemostasis on parenchymal volume reduction of the operated kidney after an open partial nephrectomy. We retrospectively reviewed 94 patients with small renal tumors who underwent open partial nephrectomy with soft-coagulation hemostasis at our institution. We measured the preoperative and postoperative renal volumes by computed tomography (CT) and calculated the renal volume reduction ratio as postoperative volume/preoperative volume. We performed multivariate analysis to identify the predictors of renal volume reduction. The median renal volume ratio was 0.75, and the median renal volume reduction rate was  $-2.49 \text{ cm}^3/\text{month}$  (IQR,  $-3.33$  to  $-1.59$ ). The RENAL score was inversely associated with ipsilateral renal volume reduction. In multivariate analysis, RENAL score, and tumor size were independent predictors of postoperative renal volume reduction. Soft coagulation hemostasis may influence the postoperative renal volume after partial nephrectomy, especially in patients with complex tumors.

**Keywords** Renal volume, Partial nephrectomy, Soft coagulation

Partial nephrectomy (PN) is the gold standard surgical treatment for renal tumors smaller than 7 cm in diameter<sup>1-3</sup>. One of the main objectives of PN is to preserve renal function, and the effect of different surgical techniques, including renal artery clamping and renorrhaphy, on postoperative renal function is still debated.

The common methods of hemostasis after tumor excision are soft-coagulation or parenchymal suture and renorrhaphy of the renal cortex<sup>4-7</sup>. Some studies have reported that soft coagulation is superior to renorrhaphy in preserving renal function, but this is inconclusive<sup>6</sup>. Our previous report investigated the preservation of the estimated glomerular filtration ratio (eGFR) one and three months after surgery using soft coagulation. The eGFR preservation rates were 91.0% and 90.7%, respectively<sup>5</sup>.

While soft coagulation is effective for achieving hemostasis, it can also lead to denaturing and necrosis of renal parenchyma<sup>8</sup>. However, the impact of soft coagulation on kidney volume reduction after partial nephrectomy remains poorly understood. In this retrospective study, we aimed to assess the chronological changes in kidney volume following surgery and identify predictive factors associated with kidney volume reduction.

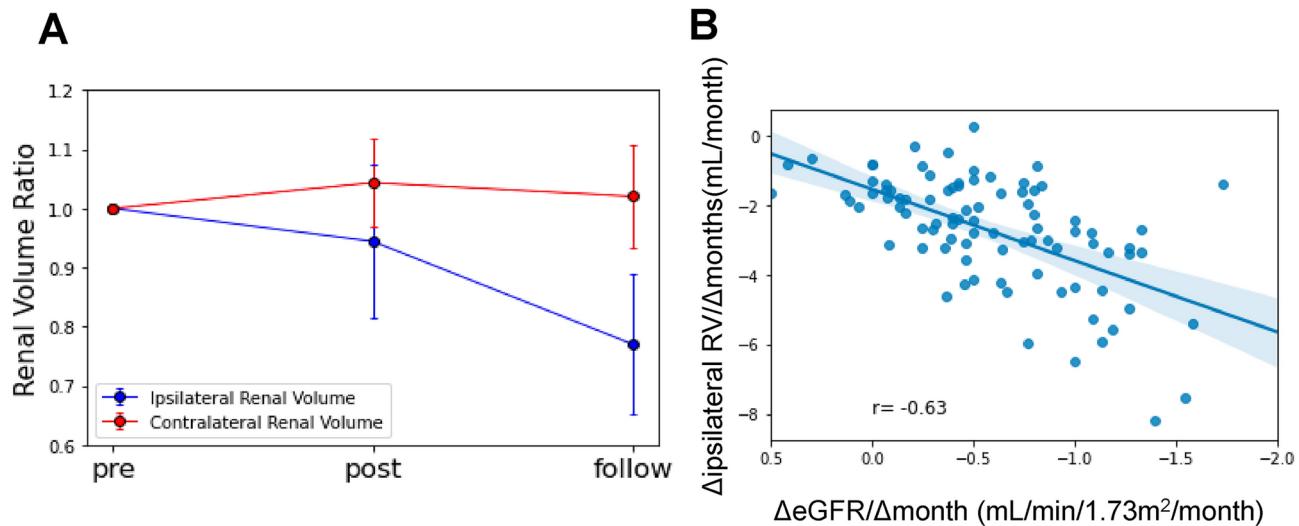
The decline in postoperative renal function after PN is attributed, in part, to the excision of functioning nephrons adjacent to the tumor and renorrhaphy, which results in focal devascularization<sup>9</sup>. Our institute performs PN using a soft coagulation system for hemostasis, omitting renal artery clamping and renorrhaphy. Despite this approach, we observed a trend toward volume decline in the operated kidney postoperatively. Various factors may contribute to this decline, including hemostasis methods. Our study aimed to shed light on the impact of soft coagulation hemostasis on parenchymal volume reduction in partial nephrectomy. We can optimize surgical techniques and improve patient outcomes by understanding these factors.

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No. patients	94
Sex (%)	
Male, n (%)	66 (70.2)
Female, n (%)	28 (29.8)
Median age, year (IQR)	65 (56.0–69.75)
Median BMI (IQR)	23.1 (21.2–26.1)
Tumor size, mm (IQR)	31.36 (20.0–40.0)
RENAL score	7 (6–8)
Preoperative eGFR, mL/min/1.73m <sup>2</sup>	69.5 (60.25–78.75)
Comorbidity, n (%)	
Hypertension, n (%)	35 (36.8)
Diabetes mellitus, n (%)	13 (13.7)
Dyslipidemia, n (%)	10 (10.5)

**Table 1.** Patient characteristics. BMI, body mass index; IQR, interquartile range.

Median operative time, min (IQR)	124 (102–154)
Mean estimated blood loss, mL (IQR)	170 (40–407)
Mean CT tumor volume, cm <sup>3</sup> (IQR)	9.1 (3.4–19.6)

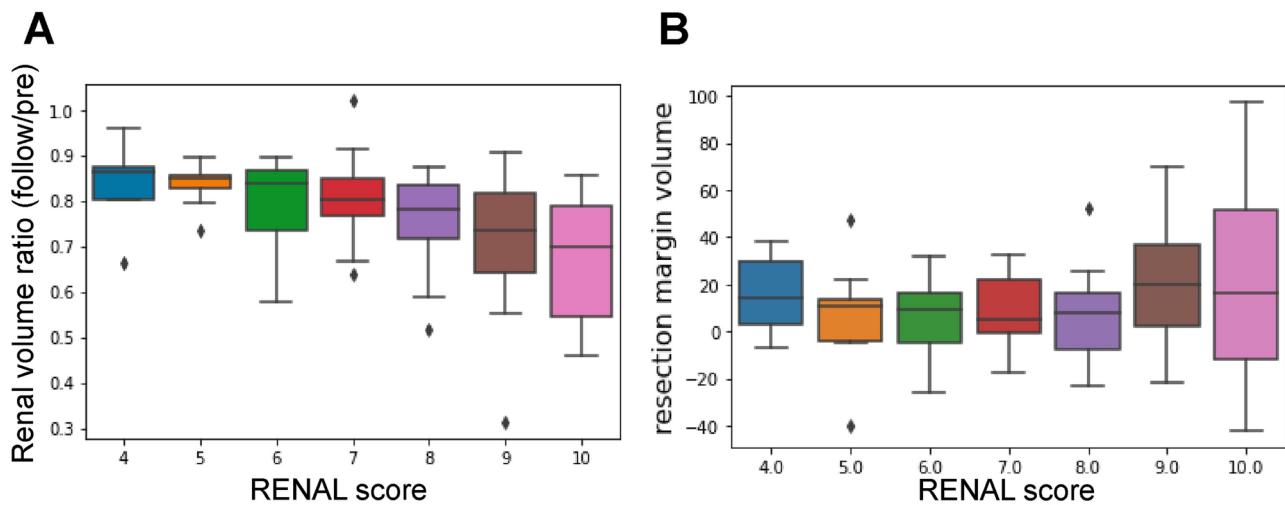
**Table 2.** Surgical outcome and CT volume results. IQR, interquartile range.**Fig. 1.** (A) Renal volume ratio of ipsilateral and contralateral kidney. Average  $\pm$  SD. (B) Correlation between ipsilateral renal volume reduction rate and the eGFR decline.

## Results

Patient characteristics are listed in Table 1. The median age and median tumor size were 65 years (Interquartile range (IQR), 56–69.75) and 28 mm (IQR, 20.0–40.0), respectively. The median RENAL nephrometry score was 7 (IQR, 6–8). The surgical results are shown in Table 2. The median operative time and estimated blood loss were 124 min (IQR, 102–152) and 170 ml (IQR, 70–407), respectively. The median postoperative follow-up period was 14 months (IQR, 12–15).

Volume reduction of the operated kidney was observed at a median ratio of 0.77, while the contralateral renal volume did not change at one-year follow-up (Fig. 1A). The median ipsilateral renal volume reduction rate was  $-2.49 \text{ cm}^3/\text{month}$  (IQR,  $-3.33$  to  $-1.59$ ), and the volume reduction correlated with estimated glomerular filtration rate (eGFR) decline ( $r = 0.63$ ) (Fig. 1B).

The RENAL score negatively correlated with volume reduction of the ipsilateral kidney (Fig. 2A). Notably, the RENAL score did not correlate with the resected renal parenchymal volume (Fig. 2B). Univariate analysis revealed tumor size, RENAL score, operation time, and estimated blood loss correlated with renal volume reduction (Table 3). Further, multivariate analysis revealed that preoperative eGFR, RENAL score, and the estimated blood



**Fig. 2.** (A) Correlation between renal volume ratio (follow/pre) and the RENAL nephrometry score. (B) Correlation between resection renal volume and the RENAL nephrometry score.

	Univariate analysis		Multivariate analysis	
	Coefficient (95% CI)	P-value	Coefficient (95% CI)	P-value
Age	0.00 (-0.002, 0.002)	0.924		
BMI	-0.0011 (-0.005, 0.07)	0.728		
HTN	-0.0087 (-0.059, 0.042)	0.733		
DM	0.0083 (-0.062, 0.079)	0.907		
DL	-0.7167 (-1.846, 0.413)	0.817		
Tumor size <sup>†</sup>	-0.027 (-0.0041, -0.001)	<0.001	-0.0015 (-0.003, 0)	0.035
RENAL <sup>†</sup>	-0.0292 (-0.044, -0.015)	<0.001	-0.0186 (-0.034, -0.004)	0.016
Operative time <sup>†</sup>	-0.0178 (-0.028, -0.007)	<0.001	-0.0001 (-0.001, -0.001)	0.741
eBL <sup>†</sup>	-0.0002 (-0.000, -0.000)	0.001	-0.000 (-0.000, 0.000)	0.072
Preoperative eGFR <sup>†</sup>	-0.0009 (-0.002, 0.001)	0.223	-0.0009 (-0.002, 0.000)	0.204

**Table 3.** Predictors of renal volume reduction, univariate and multivariate analysis. BMI, body mass index; DL, dyslipidemia; DM, diabetes mellitus; eBL, estimated blood loss; eGFR, estimated glomerular filtration ratio; HTN, hypertension; RENAL, sum of RENAL score. <sup>†</sup>indicate factors used in the multivariate analysis.

loss were the independent predictors of the postoperative volume reduction with an unstandardized regression coefficient B (95% CI) of -0.026 (-0.046, -0.007), -0.24 (-0.45, -0.032), and -0.0017 (-0.003, -0.000), respectively (p-value, 0.007, 0.024 and 0.022, respectively) (Table 3).

## Discussion

Renal function in the operated kidney occasionally experiences a decline of approximately 20% due to incomplete recovery from ischemia or nephron loss following partial nephrectomy. Factors such as excision of renal parenchyma and damage from reconstruction also play a critical role in postoperative renal function impairment<sup>9</sup>.

In our study, we performed partial nephrectomies using soft coagulation, avoiding renal hilar clamping and renal reconstruction, as reported previously<sup>4,5</sup>. Remarkably, renal function at one-year follow-up remained similar to that at three months postoperatively. The eGFR preservation rate was 89.0% at three months postoperatively and 86.9% at 12 months postoperatively<sup>4,5</sup>. However, the benefit of the off-clamp technique in preserving renal function after PN remains controversial. While some studies have reported less decline in renal function with off-clamp surgery than on-clamp surgery<sup>10</sup>, several other studies have failed to demonstrate the advantage of off-clamp surgery in eGFR preservation over on-clamp surgery in the setting of a pneumoperitoneum<sup>11–14</sup>.

The impact of renorrhaphy on postoperative renal function also remains a topic of debate. Omitting cortical renorrhaphy may contribute to eGFR preservation in the short postoperative period<sup>14–17</sup>. However, renorrhaphy can damage intraparenchymal vessels, which may lead to renal artery pseudoaneurysms. Notably, a propensity score-matched analysis of open partial nephrectomy found that renal artery pseudoaneurysms were more frequent in the renorrhaphy group<sup>18</sup>. Conversely, the non-renorrhaphy technique has not demonstrated benefits in preserving the vascularized parenchymal mass of the operated kidney and global renal function, especially

for T1b renal tumors<sup>18</sup>. It is worth noting that in the study mentioned, soft coagulation hemostasis was applied exclusively to the non-renorrhaphy group<sup>19</sup>.

However, using soft coagulation for hemostasis can adversely affect renal function. Aggressively applying soft coagulation to control bleeding may damage the renal parenchyma, impairing renal function. For instance, when soft coagulation was employed until hemostasis was achieved during partial nephrectomy, denaturation of renal parenchyma was observed up to 4.6 mm from the surface, regardless of the cauterization time *in vivo*<sup>8,20</sup>. Fujisaki et al. reported in their pig experiments that the renal parenchymal temperature increased by 15.6 °C at a depth of 5 mm and 8.8 °C at 10 mm when the surface of the kidney was cauterized with soft-coag over a period of 2, 5 and 10 s<sup>8</sup>. Despite these concerns, off-clamp partial nephrectomy using a soft coagulation system is gaining recognition as a promising surgical technique due to its positive impact on postoperative renal function<sup>11,21</sup>.

Our study identified tumor size and the RENAL score as independent predictors for postoperative parenchymal volume reduction. Additionally, reports suggest that lower parenchymal mass preservation is associated with larger tumor size, greater tumor complexity, and longer ischemia time<sup>22,23</sup>. Complex tumors often necessitate aggressive soft coagulation to control bleeding, but this approach must be carefully balanced to avoid compromising renal parenchyma and causing renal dysfunction and volume loss.

This study did not assess the split renal function using 99 m Tc dimercaptosuccinic acid (DMSA) renography. Although previous reports have suggested that CT volumetry can serve as an alternative to DMSA renography for calculating split renal function, incorporating DMSA could have provided additional insights into our study<sup>24–26</sup>. Other limitations of our study include a small patient cohort and a relatively short follow-up period. Future investigation into the long-term effects of soft coagulation on renal volume and function is anticipated. Hemostasis, in partial nephrectomy, using a soft coagulation system that may impact postoperative renal volume, particularly in patients with complex tumors.

## Methods

### Patients

Of the 220 patients with renal tumors who underwent off-clamp non-nephrectomy partial nephrectomy at our institution from 2013 to 2020, 95 were included in this retrospective cohort study. Patients without computed tomography (CT) volumetry data, with multiple partial nephrectomies, bilateral tumors, and a horseshoe kidney were excluded. Patients who underwent a single unilateral partial nephrectomy and were followed up by CT scans for at least one year were included. We finally analyzed the clinical records of 94 patients.

### Ethics approval and consent to participate

This study was approved by the Ethics Committee of NTT Medical Center, Tokyo (ID: 21-3). Informed consent was obtained in the form of opt-out on the website. Those who were rejected were excluded. This study was conducted in accordance with the Declaration of Helsinki (revised in 2013).

### Surgical techniques

All patients underwent retroperitoneal open partial nephrectomy as described elsewhere<sup>4</sup>. Briefly, partial nephrectomy was performed by blunt separation and sharp cutting followed by hemostasis with monopolar SOFT COAG (VIO300D, ERBE, Germany). To minimize blood loss, tumor resection was advanced by millimeters at a time. The renal pedicle was not secured or clamped, and cortical renorrhaphy was omitted. Resection beds were sutured with 4-0 VICRYL® only when the collecting system was opened. Urine leakage was ruled out by intravenous injection of indigo carmine solution. TachoSil® was placed on the resection surface to ensure hemostasis.

### Assessment of renal function and perioperative reduction in renal function

The eGFR was calculated using the equation:

$$186 \times (\text{Creatinine}/88.4)^{-1.154} \times (\text{Age})^{-0.203} \times (0.742 \text{ if female})$$

Perioperative eGFR preservation at five days, one month, three months, and 12 months after surgery was calculated as:

$$\text{Postoperative eGFR/preoperative eGFR} \times 100(\%)$$

### Assessment of renal volume

Renal volume (RV) measurements were performed with the SYNAPSE SAI viewer (FUJIFILM, Tokyo, Japan). The images reconstructed from non-contrast CT (5 mm slice) were analyzed, and a radiologist supervised renal volumetry. The preoperative ipsilateral renal volume was calculated as (preoperative renal volume – tumor volume). The renal volume reduction rate was calculated as follows:

$$(\text{RV}_{\text{post}} - \text{RV}_{\text{pre}}) / \text{period months}$$

and the renal volume ratio was calculated as:

$$\text{RV}_{\text{post}} / \text{RV}_{\text{pre}}$$

## Statistical analysis

Friedmann's test was performed for multi-group comparison. For post-hoc two-group analysis, the Mann-Whitney test was performed. Univariate and multivariate regression analyses were performed to identify postoperative renal volume reduction predictors. Statistical significance was set at  $p < 0.05$ . All statistical analyses were performed using the SPSS version 24 ((IBM co. ltd., Tokyo, Japan).

## Data availability

The datasets generated during the current study are available from the corresponding author on reasonable request.

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## Author contributions

IT, MK, and MN wrote the original draft. IT, MK, TI, AO, YM, TM, and MN analyzed the data. HK, SK, YS, and MN supervised the research. All authors reviewed the manuscript.

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## Declarations

### Competing interests

The authors declare no competing interests.

### Additional information

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