Analysis of Predictors of Adherent Perinephric fat and its Impact on Perioperative Outcomes in Laparoscopic Partial Nephrectomy: A Retrospective Case-control Study

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Research Article

Keywords: Adherent perinephric fat, Laparoscopic partial nephrectomy, Renal cell carcinoma, Mayo Adhesive Probability score

DOI: https://doi.org/10.21203/rs.3.rs-144111/v1

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Abstract

**Background:** Adherent perinephric fat (APF), characterized by inflammatory fat surrounding the kidney, can limit the isolation of renal tumors and increase the operative difficulty in laparoscopic partial nephrectomy (LPN). The aim of this study was to investigate the predictors of APF and its impact on perioperative outcomes during LPN.

**Methods:** A total of 215 consecutive patients undergoing LPN for renal cell carcinoma (RCC) from January 2016 to June 2019 at our institute were included. We divided these patients into two groups according to the presence of APF. Radiographic data were retrospectively collected from preoperative cross-sectional imaging. The perioperative clinical parameters were compared between the two groups. Univariate and multivariate analyses were performed to evaluate the predictive factors of APF.

**Results:** APF was observed in 41 patients (19.1%) at the time of LPN. Univariate analysis demonstrated that APF was significantly correlated with male gender ($P = 0.001$), higher body mass index ($P = 0.002$), lower preoperative estimated glomerular filtration rate ($P = 0.004$), greater posterior perinephric fat thickness ($P < 0.001$), greater perinephric stranding ($P < 0.001$) and higher Mayo Adhesive Probability (MAP) score ($P < 0.001$). The MAP score ($P < 0.001$) was the only variable that remained an independent predictor for APF in multivariate analysis. We found that patients with APF had longer operative times ($P < 0.001$), warm ischemia times ($P = 0.001$), and greater estimated blood loss ($P = 0.003$) than those without APF. However, there were no significant differences in surgical approach, transfusion rate, length of postoperative stay, complication rate or surgical margin between the two groups.

**Conclusions:** Several specific clinical and radiographic factors including the MAP score can predict APF. The presence of APF is associated with an increased operative time, and warm ischemia time and greater estimated blood loss but has no impact on other perioperative outcomes in LPN.

**Background**

Renal cell carcinoma (RCC), a heterogeneous type of cancer originating from the renal parenchyma, accounts for 90% of all renal malignancies [1]. The annual incidence of RCC has increased by approximately 2% in the last two decades, with estimated 73,820 newly diagnosed kidney cancer cases and 14,770 cancer-related deaths in 2019 in the United States [2, 3]. Due to the widespread use of ultrasound and computed tomography (CT), more incidental small renal masses have been detected in recent years.

According to the European Association of Urology (EAU) Renal Cancer Guidelines, partial nephrectomy (PN) is the preferred option for clinical stage T1 tumors (defined as tumors of $\leq 7$ cm, confined to the renal parenchyma), when technically feasible [2]. With the increased availability and utilization of laparoscopic and robot assisted techniques, minimally invasive PN (MIPN) has been identified as a safe and reproducible surgical approach, combining the advantages of decreased blood loss and hospital stay with similar oncological outcomes, compared to open PN [4–6]. In clinical practice, the treatment strategy
of PN entails a complex decision process and is dependent on tumor and patient-specific factors [7]. Several image-based tumor anatomical classification systems such as the PADUA classification system, the centrality index (C-index), and the RENAL nephrometry score system, have been applied to evaluate the complexity and potential perioperative morbidity of PN [8–10]. Nevertheless, contemporary data assessing patient-specific factors that may also complicate the technical aspects of PN are limited.

Adherent perinephric fat (APF), a notable patient-specific factor, has attracted much attention over the years. APF is characterized by inflammatory fat surrounding the kidney can restrict the isolation of renal tumors and increase the operative difficulty in PN [11–13]. Davidiuk et al [14] proposed an image-based scoring algorithm called the Mayo Adhesive Probability (MAP) score to predict the presence of APF in robot-assisted PN (RAPN). However, the small cohort of patients enrolled and inadequate clinical predictors limit its extensive use. In this study, we sought to further investigate the predictive clinical and radiographic factors, including the MAP score, for APF, as well as to assess its impact on perioperative outcomes at the time of LPN.

**Methods**

**Patient selection and data collection**

With institutional review board approval, we retrospectively reviewed the database in our institute for patients with pathologically confirmed RCC who underwent LPN from January 2016 to June 2019. The exclusion criteria were patients with an ipsilateral renal surgery history, who received preoperative neoadjuvant therapy, who had multifocal tumors, and who had incomplete clinical information. Eventually, 215 patients were enrolled in this study. Data were obtained regarding patients’ baseline clinical characteristics (gender, age, body mass index [BMI], hypertension, diabetes mellitus, tobacco use, dyslipidemia, preoperative serum creatinine, estimated glomerular filtration rate [eGFR]) and pathological characteristics (pathological stage, histological subtype, Fuhrman grade, renal capsular invasion and perinephric fat invasion).

Radiographic data (tumor size, tumor location, RENAL nephrometry score, posterior perinephric fat thickness, perinephric fat stranding, and MAP score) were collected from preoperative CT imaging within one month before LPN by two authors (LH and WY) who were independently blinded to the results of the operative notes. Posterior perinephric fat thickness was measured at the level of the renal vein as the distance from the renal capsule to the posterior abdominal wall, following a previously described procedure [15]. Perinephric fat stranding was defined in accordance with prior study [16] as a line area of soft tissue attenuation in the perinephric space and was graded according to severity (shown in Fig. 1). The final MAP score was generated from the sum of the two parameters described above with a range from 0 to 5 [14].

LPN procedures were carried out similar to previously published methods [17] by two senior experienced urologist surgeons (ZT and YDX) divided briefly into the following three steps: Step 1, establishing the
laparoscopic approach and operating space; Step 2, dissecting the perinephric fat to expose the tumor and renal hilar vessels; and Step 3, resecting the tumor and closing the wound with hilar clamping. Through reviewing the operative notes and videos, APF was determined by these two surgeons according to the presence of inflammatory perirenal fat within the Gerota's fascia requiring subcapsular dissection.

The perioperative variables collected from medical records were surgical approach, operative time, warm ischemia time (WIT), estimated blood loss (EBL), transfusion, length of postoperative stay, postoperative complication, and surgical margin. Postoperative complications within 30 days of surgery were graded according to the Clavien-Dindo classification [18].

**Statistical analysis**

Normally and non-normally distributed continuous variables are summarized as means and standard deviations and medians (first quartile [Q1], third quartiles [Q3]), respectively. Categorical variables are reported as proportions with the number of patients. The distribution of continuous variables was checked by histograms and the Kolmogorov-Smirnov (K-S) test. Univariate and multivariate logistic regression analyses were performed to evaluate the predictive factors of APF. The differences in perioperative outcomes between patients with and without APF were compared using Student's t test or the Mann–Whitney U test for continuous variables. The Chi-squared test was used to compare categorical data. A P-value < 0.05 indicates a statistically significant difference. All statistical analyses were performed using SPSS 20.0 statistical software (IBM, Chicago, IL, USA).

**Results**

**Characteristics of patients**

Of the 215 consecutive patients enrolled in the present study, 41 (19.1%) had APF noted during LPN. Table 1 highlights the baseline characteristics of the cohort in detail. The mean age at the time of diagnosis was 57 years, the majority of patients were male (64.7%), hypertensive (54.0%), and had a mean BMI of 24.1 kg/m². The median preoperative serum creatinine was 70.0 µmol/l and the mean eGFR was 117.7 ml/min/1.73 m². The mean tumor size was 3.7 cm with a standard deviation of 1.5 cm. Pathological data revealed that most patients had a pT1 (91.1%) stage tumor and a clear cell RCC subtype (78.1%). The Fuhrman nuclear grade of RCC was marked on 174 patients, of whom 13 (7.5%) had grade I, 132 (75.9%) had grade II, and 29 (16.6%) had grade III. Fifty (23.3%) patients had renal capsular invasion, and 12 (5.6%) patients had perinephric fat invasion. Perinephric fat stranding was graded as none, mild/moderate, and severe in 51.2%, 33.5%, and 15.3% of patients, respectively. The mean posterior fat thickness was 1.1 cm, median nephrometry score was 6 (Q1, Q3: 6, 8) and median MAP score was 2 (Q1, Q3: 0, 3). The proportion of patients with APF for each level of the MAP score was as follows: 0 (n = 68), 0%; 1 (n = 38), 3%; 2 (n = 24), 17%; 3 (n = 52), 10%; 4 (n = 25), 92%; 5 (n = 8), 100% (Fig. 2).
Table 1
Clinicopathological and radiographic characteristics stratified by the presence of adherent perinephric fat (APF).

| Variable                      | Total (N = 215) | APF group (N = 41) | Non APF group (N = 174) |
|-------------------------------|-----------------|--------------------|------------------------|
| Gender*                       |                 |                    |                        |
| Male                          | 139 (64.7%)     | 36 (87.8%)         | 103 (59.2%)            |
| Age (years)+                  | 57.1 ± 13.4     | 59.9 ± 14.5        | 56.5 ± 13.1            |
| BMI (kg/m^2)+                 | 24.1 ± 3.7      | 25.7 ± 3.7         | 23.7 ± 3.6             |
| Hypertension*                 | 116 (54.0%)     | 27 (65.9%)         | 89 (51.1%)             |
| Diabetes mellitus*            | 52 (24.2%)      | 13 (31.7%)         | 39 (22.4%)             |
| Tobacco use*                  | 70 (32.6%)      | 17 (41.5%)         | 53 (30.5%)             |
| Dyslipidemia*                 | 88 (41.0%)      | 16 (39.0%)         | 72 (41.4%)             |
| Preoperative creatinine (µmol/l)# | 70.0 (58.0, 84.0) | 80.0 (70.0, 97.5) | 68.0 (54.0, 81.3)      |
| Preoperative eGFR (ml/min/1.73 m^2)+ | 117.7 ± 34.5   | 103.6 ± 30.1       | 121.1 ± 34.7           |
| Tumor size (cm)+              | 3.7 ± 1.5       | 4.0 ± 1.4          | 3.6 ± 1.5              |
| Tumor location*               |                 |                    |                        |
| Left side                     | 105 (48.8%)     | 22 (53.7%)         | 83 (47.7%)             |
| Pathological stage*           |                 |                    |                        |
| pT1a                          | 131 (60.9%)     | 22 (53.7%)         | 109 (62.6%)            |
| pT1b                          | 65 (30.2%)      | 16 (39.0%)         | 49 (28.2%)             |
| pT2a                          | 6 (2.8%)        | 1 (2.4%)           | 5 (2.9%)               |
| ≥pT3                          | 13 (6.1%)       | 2 (4.9%)           | 11 (6.3%)              |
| Histological subtype*         |                 |                    |                        |
| Clear cell                    | 168 (78.1%)     | 36 (87.8%)         | 132 (75.9%)            |
| papillary cell                | 6 (2.8%)        | 1 (2.4%)           | 5 (2.9%)               |

*N (%) ; +Mean ± SD; #Median (Q1, Q3); &Fuhrman grade, 174/215 had recorded Fuhrman grade.

N, number; SD, standard deviation; Q, quartile; BMI, body mass index; eGFR, estimated glomerular filtration rate; MAP, Mayo Adhesive Probability.
Variable | Total (N = 215) | APF group (N = 41) | Non APF group (N = 174)
--- | --- | --- | ---
chromophobe | 12 (5.6%) | 1 (2.4%) | 11 (6.3%)
other subtype | 29 (13.5%) | 3 (7.4%) | 26 (14.9%)
& Fuhrman grade*
I | 13 (7.5%) | 1 (2.7%) | 12 (8.8%)
II | 132 (75.9%) | 27 (73.0%) | 105 (76.6%)
III | 29 (16.6%) | 9 (24.3%) | 20 (14.6%)
IV | 0 (0.0%) | 0 (0.0%) | 0 (0.0%)
Renal capsular invasion* | 50 (23.3%) | 11 (26.8%) | 39 (22.4%)
Perinephric fat invasion* | 12 (5.6%) | 2 (4.9%) | 10 (5.7%)
RENAL nephrometry score# | 6.0 (6.0, 8.0) | 7.0 (6.0, 8.0) | 6.0 (5.0, 8.0)
Posterior fat thickness (cm)+ | 1.1 ± 0.6 | 2.0 ± 0.6 | 0.9 ± 0.5
Perinephric stranding*
None | 110 (51.2%) | 4 (9.8%) | 106 (60.9%)
Mild/Moderate | 72 (33.5%) | 13 (31.7%) | 59 (33.9%)
Severe | 33 (15.3%) | 24 (58.5%) | 9 (5.2%)
MAP score# | 2.0 (0.0, 3.0) | 4.0 (4.0, 4.0) | 1.0 (0.0, 3.0)

*N (%); + Mean ± SD; # Median (Q1, Q3); & Fuhrman grade, 174/215 had recorded Fuhrman grade.

N, number; SD, standard deviation; Q, quartile; BMI, body mass index; eGFR, estimated glomerular filtration rate; MAP, Mayo Adhesive Probability.

Predictors of APF

The clinical and radiographic variables predicting the presence of APF were evaluated by logistic regression model. According to univariate analysis, APF significantly correlated with male gender (OR 4.963, P = 0.001), higher body mass index (OR 1.171, P = 0.002), lower preoperative estimated glomerular filtration rate (OR 0.983, P = 0.004), greater posterior perinephric fat thickness (OR 38.141, P < 0.001), greater perinephric stranding (OR 5.839; OR 70.667, P < 0.001), and higher MAP score (OR 8.945, P < 0.001) (Table 2). Based on these factors, multivariate analysis demonstrated that the MAP score (OR 8.870, P < 0.001) was the only variable that remained an independent predictor of APF (Table 3).
Table 2
Univariate logistic regression analysis for association of index variables and adherent perinephric fat (APF).

| Variable                  | Univariate Analysis | P value |
|---------------------------|---------------------|---------|
| Gender                    |                     | 0.001   |
| Male                      | 4.963 (1.857–13.264)|         |
| Female                    | Reference           |         |
| Age (years)               | 1.020 (0.993–1.047) | 0.145   |
| BMI (kg/m$^2$)            | 1.171 (1.058–1.296) | 0.002   |
| Hypertension              | 0.092               | 0.092   |
| Yes                       | 1.842 (0.905–3.749) |         |
| No                        | Reference           |         |
| Diabetes mellitus         | 0.214               | 0.214   |
| Yes                       | 1.607 (0.761–3.396) |         |
| No                        | Reference           |         |
| Tobacco use               | 0.178               |         |
| Yes                       | 1.617 (0.803–3.257) |         |
| No                        | Reference           |         |
| Dyslipidemia              | 0.783               |         |
| Yes                       | 0.907 (0.452–1.819) |         |
| No                        | Reference           |         |
| Preoperative creatinine (µmol/l) | 1.010 (0.998–1.023) | 0.101   |
| Preoperative eGFR (ml/min/1.73 m$^2$) | 0.983 (0.972–0.995) | 0.004   |
| Tumor size (cm)           | 1.214 (0.970–1.521) | 0.090   |
| Tumor location            | 0.493               |         |
| Left side                 | 1.269 (0.642–2.511) |         |
| Right side                | Reference           |         |

OR, odds ratio; CI, confidence interval; BMI, body mass index; eGFR, estimated glomerular filtration rate; ccRCC, clear cell renal cell carcinoma; MAP, Mayo Adhesive Probability.
| Variable                              | Univariate Analysis |          |
|--------------------------------------|---------------------|----------|
|                                      | OR (95% CI)         | P value  |
| Pathological stage                   |                     | 0.290    |
| pT1a                                 | Reference           |          |
| >pT1a                                | 1.448 (0.729–2.877) |          |
| Histological subtype                 |                     | 0.103    |
| ccRCC                                | Reference           |          |
| non ccRCC                            | 0.437 (0.161–1.184) |          |
| Fuhrman grade                        |                     | 0.234    |
| I                                    | Reference           |          |
| II                                   | 3.086 (0.384–24.783)|          |
| III                                  | 5.400 (0.607–48.078)|          |
| Renal capsular invasion              |                     | 0.548    |
| Yes                                  | 1.269 (0.583–2.761) |          |
| No                                   | Reference           |          |
| Perinephric fat invasion             |                     | 0.828    |
| Yes                                  | 0.841 (0.177–3.994) |          |
| No                                   | Reference           |          |
| RENAL nephrometry score              | 1.065 (0.885–1.282) | 0.506    |
| Posterior fat thickness (cm)         | 38.141 (12.524-116.156)| < 0.001 |
| Perinephric stranding                | < 0.001             |          |
| None                                 | Reference           |          |
| Mild/Moderate                        | 5.839 (1.821–18.719)|          |
| Severe                               | 70.667 (20.078-248.724)|          |
| MAP score                            | 8.945 (4.160-19.236)| < 0.001 |

OR, odds ratio; CI, confidence interval; BMI, body mass index; eGFR, estimated glomerular filtration rate; ccRCC, clear cell renal cell carcinoma; MAP, Mayo Adhesive Probability.
Table 3
Multivariate logistic regression analysis of adherent perinephric fat (APF).

| Variable                          | Multivariate Analysis |            |
|----------------------------------|-----------------------|------------|
|                                  | **OR (95% CI)**       | **P value**|
| Gender (Male vs Female)          | 2.238 (0.611-8.200)   | 0.224      |
| BMI (kg/m²)                      | 0.957 (0.826–1.108)   | 0.555      |
| Preoperative eGFR (ml/min/1.73 m²)| 1.000 (0.983–1.016)   | 0.969      |
| MAP score                        | 8.870 (3.875–20.306)  | <0.001     |

OR, odds ratio; CI, confidence interval; BMI, body mass index; eGFR, estimated glomerular filtration rate; MAP, Mayo Adhesive Probability.

Impact of APF on perioperative outcomes in LPN

As shown in Table 4, most of the patients received a retroperitoneal approach (82.8%) for LPN.
Impact of adherent perinephric fat (APF) on perioperative outcomes in laparoscopic partial nephrectomy (LPN).

| Variable                        | Total (N = 215) | APF group (N = 41) | Non APF group (N = 174) | P value |
|---------------------------------|-----------------|--------------------|-------------------------|---------|
| Surgical approach*             |                 |                    |                         | 0.344   |
| Retroperitoneal                 | 178 (82.8%)     | 36 (87.8%)         | 142 (81.6%)             |         |
| Transperitoneal                 | 37 (17.2%)      | 5 (12.2%)          | 32 (18.4%)              |         |
| Operative time (min)*          | 130.7 ± 41.0    | 158.0 ± 38.3       | 124.2 ± 39.0            | < 0.001 |
| Warm ischemia time (min)*      | 14.3 ± 7.3      | 17.9 ± 7.2         | 13.5 ± 7.2              | 0.001   |
| Estimated blood loss (ml)#     | 50.0 (30.0, 100.0) | 80.0 (50.0, 150.0) | 50.0 (30.0, 80.0)       | 0.003   |
| Transfusion*                   | 7 (3.3%)        | 3 (7.3%)           | 4 (2.3%)                | 0.254   |
| Length of postoperative stay (days)# | 8.0 (7.0, 9.0) | 8.0 (7.0, 9.0)     | 8.0 (7.0, 9.0)          | 0.191   |
| Postoperative complication*    | 68 (31.6%)      | 12 (29.2%)         | 56 (32.2%)              | 0.746   |
| Clavien-Dindo I-II             | 64 (29.8%)      | 11 (26.8%)         | 53 (30.5%)              |         |
| Clavien-Dindo III-IV           | 4 (1.8%)        | 1 (2.4%)           | 3 (1.7%)                |         |
| Surgical margin*               |                 |                    |                         | 0.957   |
| Positive                       | 5 (2.3%)        | 1 (2.4%)           | 4 (2.3%)                |         |
| Negative                       | 210 (97.7%)     | 40 (97.6%)         | 170 (97.7%)             |         |

* N (%); † Mean ± SD; # Median (Q1, Q3).

Compared with the non-APF group, the APF group was associated with a significantly longer operative time (158.0 vs. 124.2 min, P < 0.001), warm ischemia time (17.9 vs. 13.5 min, P = 0.001), and greater estimated blood loss (80 vs. 50 ml, P = 0.003). The rate of transfusion in this study population was relatively low (3.3%), and there was no difference in the length of postoperative stay. Overall, 30-day complications and positive surgical margins occurred in 31.6% and 2.3% of patients, respectively, with no difference between the two groups.

Discussion
Since LPN was first reported by Winfield et al in 1993 [19], it has increasingly become a preferred approach for the surgical management of cT1 renal masses, given evidence supporting similar oncologic efficacy and better perioperative outcomes compared to open PN [4–6]. However, LPN is technically challenging because it requires not only a negative surgical margin resection but time-dependent renal reconstruction [20]. The implementation of LPN is affected by a variety of factors, including tumor size, location, depth, and its relationship to renal hilar vessels and the urinary collecting system. Several scoring systems that quantify renal tumor anatomical factors have been developed to evaluate the surgical complexity and perioperative outcomes. Among them, the PADUA classification system, C-index, and RENAL nephrometry score system are the most widely used algorithms [8–10]. Nevertheless, these algorithms focus entirely on tumor-specific factors and ignore patient-specific factors that may also play an essential role in the LPN procedure.

It is not an uncommon occurrence when performing PN that thick and adherent perinephric adipose tissues within the Gerota's fascia complicate the exposure of the renal parenchyma and tumor. As a notable patient-specific factor, APF has attracted much attention in the last decade. Prior studies have demonstrated that the presence of APF can result in adverse perioperative outcomes during MIPN. Kocher and colleagues revealed a statistically significant association among APF, longer operative time, and higher estimated blood loss [12]. Additionally, Khene et al emphasized an elevated risk of conversion to open surgery or radical nephrectomy in patients with APF [13]. Similarly, in a large cohort of patients with RCC that underwent LPN, our data also identified APF as significantly correlated with an increased estimated blood loss (P = 0.003) and operative time (P < 0.001). We observed that APF had no impact on the surgical margins and postoperative complications. Additionally, under comparable surgeons’ experience and tumor complexity, the warm ischemia time in cases with APF was 4 min longer than in those without APF (P = 0.001), which agreed with the finding from Borregales et al [21]. The possible explanation for these results is as follows, adherent perinephric adipose tissues are more brittle and prone to bleeding, and when exposing and resecting the renal tumor, a blurred boundary caused by APF usually requires subcapsular dissection and an expanded scope of resection to ensure a negative surgical margin (Fig. 1), which further increases bleeding and suture difficulty and prolongs the warm ischemia time and operative time.

In view of the adverse perioperative outcomes associated with APF, a series of studies have been performed to investigate its physiologic mechanism and predictive factors. While the underlying pathogenesis of APF remains unclear, studies suggest that inflammation, idiopathic fibrosis, and the autoimmune response may account for APF [22]. Previous basic research has indicated the contributions of inflammation and fibrosis to abnormal adipose tissue expansion in obesity. Inflammation can lead to hypoxia and fibrosis in adipocytes, which can, in turn, promote the migration of immune cells into adipose depots [23]. As an index of obesity, the role of BMI in predicting APF is contentious. According to our univariate analysis, BMI was found to be closely associated with APF (P = 0.002), and similar findings were confirmed in other studies [13, 14]. However, it has also been argued that there is no significant correlation between BMI and APF [12], probably because BMI does not accurately reflect the variation in fat distribution, especially visceral fat (obesity), which is strongly related to metabolic syndrome [24]. This
variation manifests in gender as well, as women have more subcutaneous fat than men, while men have more perirenal fat than women [15]. As a result, most studies, including ours, indicate that males have a higher incidence of APF (P = 0.001). Furthermore, other clinical factors predicting the presence of APF, such as age, cardiovascular disease, and diabetes mellitus, have been reported in a few studies [12–14, 21]. Notably, in the present study, we found that APF correlated with a decreased preoperative level of eGFR (P = 0.004), which may suggest that a chronic inflammatory response participates in the formation of APF [25].

To further investigate the predictors of APF, the radiographic parameters were analyzed at the same time. Posterior perinephric fat thickness, as a measurement of intra-abdominal fat, has a significant relationship with APF and complications of MIPN [11, 14]. Perinephric fat stranding was initially observed in cross-sectional imaging under inflammatory conditions, such as pyelonephritis and ureteral obstruction [16], and has also been identified in cases of APF recently. Based on these two radiographic factors, a semiquantitative scoring system called the MAP score has been proposed to predict APF during RAPN [14]. Our multivariate analysis revealed that the MAP score was an independent predictor of APF (P < 0.001), providing concomitant external validation in a large cohort of LPN.

As mentioned above, the pathogenesis of APF may correlate with inflammation, while cancer-related inflammation is known to be involved in tumor development and progression, including RCC [26]. Kocher et al showed that APF was associated with malignant renal histology (versus benign disease) [12], and Thiel and colleagues revealed that high MAP scores were related to decreased progression-free survival of RCC [27]. Interestingly, our study failed to elucidate the association between APF and tumor aggressive behaviors.

There are several limitations in this study. First, although we performed our investigation on the same patient population (RCC) with the same surgical approach (LPN), the retrospective nature of this study might introduce selection and recall bias. Second, the limited number of single-center patients with APF and the relatively strong correlations among previously mentioned clinical factors made the application of multivariate model analysis challenging. Third, our subjective definition of APF that follows that of most studies could still be debated.

**Conclusions**

APF can be preoperatively predicted with the comprehensive assessment of several specific clinical and radiographic factors, including male gender, higher BMI and the MAP score. The presence of APF is associated with an increased operative time, and warm ischemia time and greater estimated blood loss but has no impact on other perioperative outcomes in LPN. Consequently, the accurate evaluation and adequate understanding of APF will be helpful to counsel patient selection and improve outcomes.

**Abbreviations**
Declarations

Ethics approval and consent to participate

This study was approved by the Ethics Committee of The Second Hospital of Anhui Medical University (No. YX2020-015-F2). The study protocol is performed in accordance with the Declaration of Helsinki. The written informed consent was not required for this retrospective study. The Ethics Committee of The Second Hospital of Anhui Medical University permitted waiving the need of informed consent, and we guaranteed that data used in this study was de-identified.

Consent for publication

Not applicable.

Availability of data and materials

The datasets generated and analysed during the current study are not publicly available because we are conducting further investigations but are available from the corresponding author on reasonable request.

Competing interests

The authors have no conflicts of interest to disclose.

Funding

This study was supported by the Scientific Research Foundation of Anhui Medical University (No. 2019xkj037).

Authors’ contributions

Lu Fang, Yi Wang, and Dexin Yu designed the study. Huan Li, Rui Liu, Taotao Zhang, Tao Zhang, Liangkuan Bi and Dongdong Xie collected and analyzed data. Lu Fang and Huan Li wrote the paper. Yi Wang and Dexin Yu revised the paper.
Acknowledgement

The authors gratefully thank all of the participants in this study for supporting this study.

References

1. Guo S, Yao K, He X, Wu S, Ye Y, Chen J, Wu CL: Prognostic significance of laterality in renal cell carcinoma: A population-based study from the surveillance, epidemiology, and end results (SEER) database. Cancer medicine 2019, 8(12):5629–5637.

2. Ljungberg B, Bensalah K, Canfield S, Dabestani S, Hofmann F, Hora M, Kuczyk MA, Lam T, Marconi L, Merseburger AS et al: EAU guidelines on renal cell carcinoma: 2014 update. European urology 2015, 67(5):913–924.

3. Siegel RL, Miller KD, Jemal A: Cancer statistics, 2020. CA: a cancer journal for clinicians 2020, 70(1):7–30.

4. Bravi CA, Larcher A, Capitanio U, Mari A, Antonelli A, Artibani W, Barale M, Bertini R, Bove P, Brunocilla E et al: Perioperative Outcomes of Open, Laparoscopic, and Robotic Partial Nephrectomy: A Prospective Multicenter Observational Study (The RECORd 2 Project). European urology focus 2019.

5. Chang KD, Abdel Raheem A, Kim KH, Oh CK, Park SY, Kim YS, Ham WS, Han WK, Choi YD, Chung BH et al: Functional and oncological outcomes of open, laparoscopic and robot-assisted partial nephrectomy: a multicentre comparative matched-pair analyses with a median of 5 years’ follow-up. BJU international 2018, 122(4):618–626.

6. Pereira J, Renzulli J, 2nd, Pareek G, Moreira D, Guo R, Zhang Z, Amin A, Mega A, Golijanin D, Gershman B: Perioperative Morbidity of Open Versus Minimally Invasive Partial Nephrectomy: A Contemporary Analysis of the National Surgical Quality Improvement Program. Journal of endourology 2018, 32(2):116–123.

7. Tran MGB, Aben KKH, Werkhoven E, Neves JB, Fowler S, Sullivan M, Stewart GD, Challacombe B, Mahrous A, Patki P et al: Guideline adherence for the surgical treatment of T1 renal tumours correlates with hospital volume: an analysis from the British Association of Urological Surgeons Nephrectomy Audit. BJU international 2020, 125(1):73–81.

8. Ficarra V, Novara G, Secco S, Macchi V, Porzionato A, De Caro R, Artibani W: Preoperative aspects and dimensions used for an anatomical (PADUA) classification of renal tumours in patients who are candidates for nephron-sparing surgery. European urology 2009, 56(5):786–793.

9. Simmons MN, Ching CB, Samplaski MK, Park CH, Gill IS: Kidney tumor location measurement using the C index method. The Journal of urology 2010, 183(5):1708–1713.

10. Kutikov A, Uzzo RG: The R.E.N.A.L. nephrometry score: a comprehensive standardized system for quantitating renal tumor size, location and depth. The Journal of urology 2009, 182(3):844–853.

11. Gorin MA, Mullins JK, Pierozazio PM, Jayram G, Allaf ME: Increased intra-abdominal fat predicts perioperative complications following minimally invasive partial nephrectomy. Urology 2013,
12. Kocher NJ, Kunchala S, Reynolds C, Lehman E, Nie S, Raman JD: Adherent perinephric fat at minimally invasive partial nephrectomy is associated with adverse peri-operative outcomes and malignant renal histology. BJU international 2016, 117(4):636–641.

13. Khene ZE, Peyronnet B, Mathieu R, Fardoun T, Verhoest G, Bensalah K: Analysis of the impact of adherent perirenal fat on peri-operative outcomes of robotic partial nephrectomy. World journal of urology 2015, 33(11):1801–1806.

14. Davidiuk AJ, Parker AS, Thomas CS, Leibovich BC, Castle EP, Heckman MG, Custer K, Thiel DD: Mayo adhesive probability score: an accurate image-based scoring system to predict adherent perinephric fat in partial nephrectomy. European urology 2014, 66(6):1165–1171.

15. Eisner BH, Zargooshi J, Berger AD, Cooperberg MR, Doyle SM, Sheth S, Stoller ML: Gender differences in subcutaneous and perirenal fat distribution. Surgical and radiologic anatomy: SRA 2010, 32(9):879–882.

16. Kim S, Choi SK, Lee SM, Choi T, Lee DG, Min GE, Jeon SH, Lee HL, Chung JY, Joh JH et al: Predictive Value of Preoperative Unenhanced Computed Tomography During Ureteroscopic Lithotripsy: A Single Institute's Experience. Korean journal of urology 2013, 54(11):772–777.

17. Gill IS, Desai MM, Kaouk JH, Meraney AM, Murphy DP, Sung GT, Novick AC: Laparoscopic partial nephrectomy for renal tumor: duplicating open surgical techniques. The Journal of urology 2002, 167(2 Pt 1):469–467; discussion 475 – 466.

18. Clavien PA, Barkun J, de Oliveira ML, Vauthey JN, Dindo D, Schulick RD, de Santibañes E, Pekolj J, Slankamenac K, Bassi C et al: The Clavien-Dindo classification of surgical complications: five-year experience. Annals of surgery 2009, 250(2):187–196.

19. Winfield HN, Donovan JF, Godet AS, Clayman RV: Laparoscopic partial nephrectomy: initial case report for benign disease. Journal of endourology 1993, 7(6):521–526.

20. Aboumarzouk OM, Stein RJ, Eyraud R, Haber GP, Chlosta PL, Somani BK, Kaouk JH: Robotic versus laparoscopic partial nephrectomy: a systematic review and meta-analysis. European urology 2012, 62(6):1023–1033.

21. Borregales LD, Adibi M, Thomas AZ, Reis RB, Chery LJ, Devine CE, Wang X, Potretzke AM, Potretzke T, Figenshau RS et al: Predicting Adherent Perinephric Fat Using Preoperative Clinical and Radiological Factors in Patients Undergoing Partial Nephrectomy. European urology focus 2019.

22. Bylund JR, Qiong H, Crispen PL, Venkatesh R, Strup SE: Association of clinical and radiographic features with perinephric "sticky" fat. Journal of endourology 2013, 27(3):370–373.

23. Crewe C, An YA, Scherer PE: The ominous triad of adipose tissue dysfunction: inflammation, fibrosis, and impaired angiogenesis. The Journal of clinical investigation 2017, 127(1):74–82.

24. Després JP, Lemieux I: Abdominal obesity and metabolic syndrome. Nature 2006, 444(7121):881–887.

25. Gupta J, Mitra N, Kanetsky PA, Devaney J, Wing MR, Reilly M, Shah VO, Balakrishnan VS, Guzman NJ, Girndt M et al: Association between albuminuria, kidney function, and inflammatory biomarker
profile in CKD in CRIC. Clinical journal of the American Society of Nephrology: CJASN 2012, 7(12):1938–1946.

26. Chang Y, An H, Xu L, Zhu Y, Yang Y, Lin Z, Xu J: Systemic inflammation score predicts postoperative prognosis of patients with clear-cell renal cell carcinoma. British journal of cancer 2015, 113(4):626–633.

27. Thiel DD, Davidiuk AJ, Meschia C, Serie D, Custer K, Petrou SP, Parker AS: Mayo Adhesive Probability Score Is Associated With Localized Renal Cell Carcinoma Progression-free Survival. Urology 2016, 89:54–60.

Figures

Figure 1

Grading of perinephric stranding and corresponding intraoperative images. (A1) None: 0 points. The fat around the kidney demonstrates no stranding. (A2) The difficulty of perinephric fat dissection was quite simple with clear boundary and rare bleeding. (B1) Mild/moderate: 2 points. The fat around the kidney has some image-dense stranding present but no thick bars of inflammation. (B2) The difficulty of perinephric fat dissection was moderate with still clear boundary and minor bleeding. (C1) Severe: 3 points. Image shows severe stranding around the kidney with thick image-dense bars of inflammation. (C2) The difficulty of perinephric fat dissection was hard with blurred boundary and obvious bleeding.