Risk factors for prolonged hospitalization in patients undergoing laparoscopic adrenalectomy

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Introduction: Even though laparoscopic adrenalectomy is currently a standard, there are important variations between different centres in short-term treatment results such as length of hospital stay (LOS) or morbidity.

Aim: To determine the factors affecting LOS in patients after laparoscopic transperitoneal lateral adrenalectomy (LTA).

Material and methods: The study enrolled 453 patients (173 men and 280 women, mean age 57 years) who underwent LTA between 2009 and 2017. Discharge from hospital after more than median hospital stay was considered as prolonged LOS. We evaluated factors that potentially may influence LOS (primary length of stay after surgery, excluding readmissions). Logistic regression models were used in univariate and corrected multivariate analyses, in order to identify the factors related to prolonged LOS.

Results: The median LOS after LTA in the studied group was 2 days. One hundred seventy-five (38.5%) patients required prolonged hospitalization. Univariate logistic regression showed that the following factors were related to prolonged LOS: presence of any comorbidity, cardiovascular disease, intraoperative complications, postoperative complications, day of the week of operation (surgery on Thursday or Friday), intraoperative blood loss, need for transfusion, hormonal activity, postoperative drainage, ASA (III–IV) and histological type – pheochromocytoma.

Multivariate logistic regression showed that only complications (OR = 3.86; 95% CI: 1.84–8.04), day of the week of operation (Thursday or Friday) (OR = 4.85; 95% CI: 3.04–7.73), need for drainage (OR = 3.63; 95% CI: 1.55–8.52), and histological type – pheochromocytoma (OR = 2.48; 95% CI: 1.35–4.54) prolonged LOS.

Conclusions: Prolonged length of hospital stay following laparoscopic transperitoneal lateral adrenalectomy is strongly associated with the presence of postoperative complications, day of the week of operation (Thursday or Friday), need for drainage, and histological type – pheochromocytoma.

Key words: laparoscopy, adrenalectomy, adrenal tumour, prolonged hospitalization.

Introduction

The gold standard for surgical treatment of adrenal tumours is laparoscopic adrenalectomy [1, 2]. Over the last two decades it has systematically replaced open procedures due to the multiple benefits of a minimally invasive approach. It has been proven that laparoscopic adrenalectomy hastens convalescence, reduces length of hospital stay (LOS) and, most importantly, lowers the morbidity rate [3, 4]. Even though laparoscopic adrenalectomy is currently a standard, there is considerable variation between different centres in short-term treatment results, such as LOS or morbidity [5]. Median LOS varies
from 2 to 8 days, whereas morbidity rates, according to some authors, range from 4% to 23% [6–8]. All of the above prompted us to identify which factors may potentially cause these differences.

Aim

The aim of this study was to determine the factors affecting LOS in patients after laparoscopic transperitoneal lateral adrenalectomy (LTA).

Material and methods

The prospective study included consecutive patients undergoing elective laparoscopic transperitoneal adrenalectomy from 2009 to 2017 in the 2nd Department of General Surgery. Patients who initially underwent open surgery or patients with an inoperable tumour with distant metastases were excluded from the study.

Our department is a tertiary referral university unit. Annually more than 700 laparoscopic procedures are performed – mainly gastric, bariatric, colorectal, pancreatic and hepatobiliary surgeries, with more than 60 cases of adrenalectomy [9, 10]. Since 2003 a laparoscopic approach has been the preferred access for adrenalectomy in our unit. However, in selected cases single access laparoscopy has been used [11, 12].

All patients had preoperative evaluation including imaging: most commonly computed tomography. In selected patients, magnetic resonance imaging (MRI) or positron emission tomography (PET) scans were performed. In all cases evaluation of the tumour’s hormonal activity was performed (plasma cortisol, urinary free cortisol, aldosterone, urinary aldosterone, plasma renin activity, metoxycatecholamines and vanillylmandelic acid, adrenocorticotropin, dexamethasone suppression test, dehydroepiandrosterenedione, 17-OH-progesterone, testosterone).

We evaluated factors that potentially may influence LOS (primary length of stay after surgery, excluding readmissions). Our analysis included: gender, age, body mass index (BMI), distance from patient’s home to hospital, risk of anaesthesia measured with American Society of Anaesthesiologists (ASA) score, diabetes, cardiovascular disease and other comorbidities, history of previous abdominal surgery, hormonal activity, radiographic tumour size (largest diameter measured on adrenal-CT or MRI), side, histopathology and character of the tumour, operative time, day of operation (day of the week), intraoperative blood loss, complications and conversions to open surgery.

All procedures were performed in accordance with the ethical standards laid down in the 1964 Declaration of Helsinki and its later amendments. Written informed consent for the proposed surgical treatment was obtained from all patients before surgery. The entire study group consisted of 453 patients (280 women and 173 men). The mean age was 57 years (18–87 years). Three hundred and ninety-eight patients were operated on for benign neoplasms, 55 for malignant tumours. The demographic analysis of the group is shown in Table I.

| Parameter                                      | Value                  |
|------------------------------------------------|------------------------|
| Number of patients                             | 453                    |
| Females, n (%)                                 | 280 (61.8)             |
| Males, n (%)                                   | 173 (38.2)             |
| Age, mean ± SD [years]                         | 56.68 ± 13.5           |
| ASA 1, n (%)                                   | 21 (4.6)               |
| ASA 2, n (%)                                   | 253 (55.8)             |
| ASA 3, n (%)                                   | 173 (38.2)             |
| ASA 4, n (%)                                   | 6 (1.3)                |
| Any comorbidity, n (%)                         | 355 (78.4)             |
| Cardiovascular, n (%)                          | 116 (25.6)             |
| Hypertension, n (%)                            | 284 (62.7)             |
| Diabetes, n (%)                                | 126 (27.8)             |
| Pulmonary disease, n (%)                       | 63 (13.9)              |
| Renal disease, n (%)                           | 20 (4.4)               |
| Liver disease, n (%)                           | 11 (2.4)               |
| Previous abdominal surgery, n (%)              | 209 (46.1)             |
| Right suprarenal tumour, n (%)                 | 223 (49.2)             |
| Left suprarenal tumour, n (%)                  | 230 (50.8)             |
| Benign tumour, n (%)                           | 398 (87.9)             |
| Malignant tumour, n (%)                        | 55 (12.1)              |
| Operative time, mean ± SD [min]                | 97.0 ± 39.0            |
| Operative time, median (IQR) [min]             | 90 (70–120)            |
| Intraoperative blood loss, mean ± SD [ml]      | 88.7 ± 194.4           |
| Intraoperative blood loss, median (IQR) [ml]   | 50 (20–70)             |
| Conversion, n (%)                              | 5 (1.1)                |
Statistical analysis

All data were analysed with StatSoft Statistica v.10. The results are presented as mean ± standard deviation (SD) or median with interquartile range (IQR) when appropriate. Prolonged LOS was defined as discharge from hospital after more than calculated median hospital stay. A univariate logistic regression analysis of individual demographic and perioperative parameters was undertaken to assess factors influencing prolonged LOS. Finally, the variables in the univariate logistic regression analysis that had a significant impact on the length of hospital stay were used to build a multivariate logistic regression model. Results were considered statistically significant when the $p$-value was found to be less than 0.05.

Results

The median LOS in the entire group was 2 days. Length of hospital stay was considered as prolonged if it was longer than 2 days. Only 175 (38.5%) patients required hospitalization longer than 2 days (Figure 1).

Mean operative time in the entire group was 97.0 ±39.0 min and mean intraoperative blood loss was 88.7 ±194.4 ml. Conversion was performed in 5 patients. The reasons for conversion were: adhesions after previous surgery, abnormal location of the tumour, infiltration to adjacent organs, damage to the tumour capsule and uncontrolled bleeding in 2 cases. Table II presents postoperative outcomes in the analysed group.

Table II. Postoperative outcomes in analysed groups

| Parameter | Value         |
|-----------|---------------|
| Patients with complications, n (%) | 35 (7.7)       |
| Clavien-Dindo 1, n (%)               | 19 (4.2)       |
| Clavien-Dindo 2, n (%)               | 10 (2.2)       |
| Clavien-Dindo 3, n (%)               | 2 (0.4)        |
| Clavien-Dindo 4, n (%)               | 3 (0.7)        |
| Clavien-Dindo 5, n (%)               | 1 (0.2)        |
| Length of hospital stay, mean ± SD [days] | 2.5 ±1.5     |
| Length of hospital stay [days], median (IQR) | 2 (2–3)       |
| Readmission, n (%)                   | 7 (1.5%)       |

Postoperative complications occurred in 35 (7.7%) patients, with 6 (1.3%) being severe (Clavien-Dindo 3-5) (Table III). Readmission within 30 days after discharge was necessary in 7 (1.5%) patients.

Univariate logistic regression showed that: presence of any comorbidity (OR = 1.65; 95% CI: 1.02–2.68), cardiovascular disease (OR = 1.79; 95% CI: 1.02–2.7) and surgical site infection (OR = 3.35; 95% CI: 1.05–10.61) were independent risk factors for prolonged hospitalization.
CI: 1.17–2.75), intraoperative complications (OR = 2.14; 95% CI: 1.06–4.32), postoperative complications (OR = 3.86; 95% CI: 1.84–8.04), day of the week of operation (surgery on Thursday or Friday) (OR = 3.75; 95% CI: 2.52–5.58), intraoperative blood loss (> 90 ml) (OR = 1.74; 95% CI: 1.12–2.71), need for transfusion (OR = 15.02; 95% CI: 1.89–119.61), hormonal activity (OR = 1.61; 95% CI: 1.10–2.35), need for drainage (OR = 3.52; 95% CI: 1.71–7.25), ASA (III–IV) (OR = 1.92; 95% CI: 1.31–2.83), and histological type – pheochromocytoma (OR = 1.95; 95% CI: 1.17–3.25) were related to prolonged LOS (Table IV).

Next, the multivariate logistic regression model showed that only the presence of postoperative complications (OR = 3.86; 95% CI: 1.84–8.04), day of the week (Thursday or Friday) (OR = 4.85; 95% CI: 3.04–7.73), need for drainage (OR = 3.63; 95% CI: 1.55–8.52) and histological type – pheochromocytoma (OR = 2.48; 95% CI: 1.35–4.54) were significant factors prolonging LOS. Gender, BMI, presence of previous abdominal surgery, presence of comorbidities, tumour side, and ASA had no effect on primary length of hospital stay (Table V).

**Discussion**

Our study showed that there are several unrelated factors affecting prolonged LOS in patients undergoing laparoscopic adrenalectomy. These factors include presence of postoperative complications, day of the week of the operation (Thursday or Fri-

| Parameter                          | OR     | 95% CI          | P-value |
|------------------------------------|--------|-----------------|---------|
| Sex (female vs. male)              | 0.88   | 0.60–1.30       | 0.5293  |
| Age [years]                        | 1.00   | 0.99–1.02       | 0.6215  |
| BMI [kg/m²]                        | 0.99   | 0.96–1.02       | 0.4877  |
| Previous abdominal surgery (yes vs. no) | 1.11 | 0.75–1.62       | 0.6118  |
| Any comorbidity (yes vs. no)       | 1.65   | 1.02–2.68       | 0.0418  |
| Diabetes (yes vs. no)              | 1.40   | 0.92–2.12       | 0.1154  |
| Cardiovascular disease (yes vs. no) | 1.79 | 1.17–2.75       | 0.0074  |
| Tumour site (right vs. left)       | 0.86   | 0.59–1.25       | 0.4235  |
| Intraoperative complications (yes vs. no) | 2.14 | 1.06–4.32       | 0.0350  |
| Complications (yes vs. no)         | 3.86   | 1.84–8.04       | 0.0004  |
| Conversion (yes vs. no)            | 6.48   | 0.72–58.46      | 0.0959  |
| Day of the week                    | 3.75   | 2.52–5.58       | < 0.0001|
| Operative time (> 120 vs. ≤ 120 min) | 1.49 | 0.98–2.24       | 0.0596  |
| Need for transfusion               | 15.02  | 1.89–119.61     | 0.0105  |
| Blood loss (> 90 vs. ≤ 90 ml)      | 1.74   | 1.12–2.71       | 0.0141  |
| Drainage                           | 3.52   | 1.71–7.25       | 0.0006  |
| ASA (III–IV vs. I–II)              | 1.92   | 1.31–2.83       | 0.0009  |
| Hormonal activity (yes vs. no)     | 1.61   | 1.10–2.35       | 0.0152  |
| Distance from home                 | 1.00   | 1.00–1.00       | 0.7952  |
| Pheochromocytoma                   | 1.95   | 1.17–3.25       | 0.0101  |
| Tumour character (malignant vs. benign) | 1.16 | 0.65–2.05       | 0.6146  |
| Size of the tumour [cm]            | 1.05   | 0.97–1.14       | 0.2150  |
drenectomy. The study featured 38.5% of patients requiring hospitalization longer than 2 days, with most patients discharged on postoperative day 1 or day 2. As compared to other studies, the median hospital stay was reported to be 5–6 days. The median hospital stay in our study group was 2 days, with 38.5% of patients requiring hospitalization longer than 2 days. The low hospital stay is attributed to the experience and standards of care in the tertiary hospital.

The median hospital stay in our study was 2 days. Only 38.5% of patients required hospitalization longer than 2 days, which we set as the cut-off point for stating prolonged LOS. More than 61.3% of patients were discharged on postoperative day 1 or day 2. Similar LOS was reported by Karabulut et al. and Pineda-Solis et al., whereas Shi et al. and Wang et al. report their LOS to be 5–6 days [7, 8, 13, 14]. Such short hospital stays in our unit are the result of several factors. First of all, our department is part of a tertiary referral university hospital with a high annual volume of laparoscopic adrenalectomies (around 60 per year). In a previously published study, based on 500 adrenalectomies performed in our unit, it was proven that both surgical experience and perioperative care lead to reduced LOS [15]. Secondly, over the years the perioperative care has improved, which also affected the outcomes [16–19].

Length of hospital stay is inextricably related to the extent of the surgery and the patient’s general condition in the postoperative period. The classical approach and the occurrence of complications are undoubtedly reasons for prolonged hospital stay. Most cases of open adrenalectomy require a much longer hospital stay, and discharges after 24 to 48 h are rare. Due to the fact that since 2003 almost all adrenalectomies have been performed with minimally invasive access, open procedures are performed mainly in cases of conversion. The conversion rate in our group was relatively low (1.1%). In addition, there was a small proportion of patients operated on using single access. We did not observe any shortening of LOS despite the reduction of the number of trocars. It seems that the only benefit of the approach is a minor cosmetic improvement, but at the cost of a longer operative time [20, 21]. Therefore this technique still remains questionable.

Another factor quite obviously related to prolonged LOS is complications. On the other hand, the rate of severe complications in our group was relatively low. Therefore, we tried to determine which other factors (demographic and perioperative) may allow prediction of the necessity for prolonged hospitalization. We observed that the histological type of the tumour being pheochromocytoma affects the length of hospital stay, which was proven in the multivariate logistic regression model. These patients require special care in the perioperative period. The morbidity rate is greater and ranges from 5% to 23% [22]. In addition, surgery for pheochromocytoma is considered more difficult compared to other tumours [23]. The most important complication in this study group is haemodynamic instability in the perioperative period [24]. It may often require catecholamine infusion, even up to several days after the surgery. Similar conclusions, regarding LOS in patients with pheochromocytoma, were drawn by other authors. Conzo et al. reported LOS of 4 days, Kim et al. almost 6 days and Gagner et al. over 8 days [6, 25, 26]. On the other hand, Kercher et al. reported a short time

### Table V. Multivariate logistic regression affecting prolonged hospitalization (> 2 days)

| Parameter                        | OR   | 95% CI       | P-value |
|----------------------------------|------|--------------|---------|
| Any comorbidity (yes vs. no)     | 1.15 | 0.63–2.11    | 0.6519  |
| Cardiovascular disease (yes vs. no) | 1.31 | 0.73–2.36    | 0.3667  |
| Day of the week                  | 4.85 | 3.04–7.73    | < 0.0001|
| Hormonal activity (yes vs. no)   | 1.34 | 0.85–2.13    | 0.2117  |
| Intraoperative complications (yes vs. no) | 0.95 | 0.27–3.31    | 0.9329  |
| Complications (yes vs. no)       | 3.86 | 1.84–8.04    | < 0.0001|
| Need for transfusion             | 7.55 | 0.58–98.76   | 0.1233  |
| Blood loss (> 90 vs. ≤ 90 ml)    | 1.26 | 0.62–2.55    | 0.5180  |
| Need of drainage                 | 3.63 | 1.55–8.52    | 0.0031  |
| ASA (III–IV vs. I–II)            | 1.42 | 0.82–2.46    | 0.2060  |
| Pheochromocytoma                 | 2.48 | 1.35–4.54    | 0.0133  |
of hospitalization, 2 to 3 days, and a relatively low morbidity rate of 4% [27]. Similar short LOS were reported by Cheah et al. and Jaroszewski et al. [28, 29]. Such vast diversity in LOS is mainly caused by the differences in reported complication rate, including the most important one – haemodynamic instability. Some authors have reported that patients with different hormonal activity of the tumours such as patients with Cushing syndrome, who require steroid supplementation in the perioperative period, may have prolonged hospitalization. In most cases, early introduction of an oral diet, early mobilization and optimal analgesic protocol allow hospitalization to be shortened, while steroid supplementation is continued after discharge orally.

Another factor significantly prolonging hospitalization is postoperative drainage. It is not routinely used in our centre, which supports the idea of modern perioperative care. In one study, Major et al. showed that drainage after laparoscopic adrenalectomy is not only unnecessary, but may also be associated with an increased risk of complications [30]. In our unit the use of drains was justified only when there was an increased risk of postoperative bleeding, and they were removed as soon as possible.

The last factor significantly extending LOS was the day of the week on which the surgery was performed. The operation being on Thursday and Friday was related to a longer stay as a result of the reluctance to discharge patients during the weekend.

Intraoperative blood loss and duration of surgery were the next possible factors prolonging LOS. However, the multivariate regression model revealed that these factors had no significant impact. It may seem that these parameters are associated with intraoperative difficulties, which in turn may increase the risk of complications. Because of that, the multivariate regression model shows that the morbidity has a significant impact on the length of hospital stay.

Our study has limitations associated with the single centre design. Another limiting factor is the fact that throughout this period, the perioperative care protocol in our unit has been modified and the proficiency in laparoscopic surgery has increased. These factors may bias our results.

Conclusions

Our study has proven that the tumour histological type being pheochromocytoma, postoperative complications, operation at the end of the week (on Thursday or Friday), and the need for drainage may cause prolonged LOS in patients undergoing laparoscopic adrenalectomy.

Conflict of interest

The authors declare no conflict of interest.

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Received: 23.10.2017, accepted: 3.12.2017.