Second-stage treatment strategy for severely obese patient with primary aldosteronism—laparoscopic adrenalectomy after laparoscopic sleeve gastrectomy: a case report

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> **Background:** Primary aldosteronism (PA) is the major cause of curable hypertension. Obesity can affect the therapeutic efficacy of adrenalectomy for PA. Consequently, it is essential to consider weight loss when treating obese patients with PA. Herein, we report the case of a severely obese patient with PA who was successfully treated with laparoscopic sleeve gastrectomy (LSG) before laparoscopic adrenalectomy.

> Case Description: A 36-year-old man treated for hypertension over the previous nine years developed hypokalemia. Blood tests showed a high plasma aldosterone concentration (PAC)-to-plasma renin activity (PRA) ratio, and enhanced computed tomography (CT) revealed a low-density tumor in the left adrenal gland. Both furosemide upright and captopril challenge tests were positive, and adrenal venous sampling (AVS) after adrenocorticotropic hormone administration confirmed unilateral PAC elevation in the left adrenal vein. Adrenalectomy was considered; however, the patient was severely obese with a body mass index (BMI) of 45.9 kg/m² and type 2 diabetes (T2D). After considering the potential risks of perioperative complications and the therapeutic efficacy of adrenalectomy, we performed LSG. Six months after LSG, the patient's BMI had decreased to 34.1 kg/m² and the T2D was in remission. Consequently, we performed laparoscopic left adrenalectomy. Twelve months after the LSG and treatment with only one antihypertensive drug, the patient's blood pressure remained stable.

> **Conclusions:** Here, we report a two-stage treatment strategy for a patient with severe obesity complicated by PA. Performing LSG before adrenalectomy is a rational strategy to reduce the risk of perioperative complications and improve therapeutic efficacy after adrenal surgery.

> Keywords: Primary aldosteronism (PA); laparoscopic adrenalectomy; laparoscopic sleeve gastrectomy (LSG); obesity; case report

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Introduction

Primary aldosteronism (PA) accounts for approximately 10% of patients with hypertension who visit primary care centers, and is a major cause of curable hypertension (1). If a unilateral aldosterone-producing tumor can be confirmed by a detailed examination such as computed tomography (CT) or adrenal venous sampling (AVS), adrenalectomy is usually selected as a radical treatment (1). However, obesity can affect the PA treatment after adrenalectomy (2,3), and when treating obese patients with PA, it is prudent to consider weight loss.

Worldwide, the proportion of patients suffering from severe obesity has increased due to various factors including Westernized diets and a lack of exercise (4). Obesity not only contributes to the difficulty of surgical procedures, but also increases the risk of complications such as postoperative respiratory failure and deep vein thrombosis (5). In Japan, laparoscopic sleeve gastrectomy (LSG) for patients suffering from severe obesity has been covered by insurance since 2014; subsequently the number of LSGs has increased (6). Performing the procedure prior to radical surgery for benign or slowgrowth diseases under the management of skilled obesity treatment teams is a rational treatment strategy to reduce the risks associated with obesity described above. We have also encountered cases of LSG prior to radical surgery for pancreatic neuroendocrine neoplasm and abdominal aortic

Highlight box

Key findings

 We report the case of a severely obese patient with primary aldosteronism (PA) who was successfully treated with laparoscopic adrenalectomy 6 months after laparoscopic sleeve gastrectomy (LSG).

What is known and what is new?

- Obesity is related to perioperative complications and also affects therapeutic efficacy after adrenalectomy.
- Metabolic surgery, such as LSG, are powerful surgical procedures for weight loss in severely obese patients.
- Here, we report the usefulness of LSG before adrenalectomy in severely obese patients with PA.

What is the implication, and what should change now?

 Performing LSG prior to adrenalectomy in severely obese patients with PA should be considered a rational treatment strategy to reduce the risk of perioperative complications and improve therapeutic efficacy after adrenal surgery.

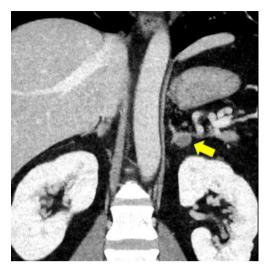


Figure 1 Enhanced computed tomography reveals a 15-mm well-defined tumor with low attenuation in the left adrenal gland (yellow arrow).

aneurysm, and reported on its efficacy and safety (5,7,8).

In the present study, we report the case of a severely obese patient with PA who was successfully treated with LSG and laparoscopic adrenalectomy in two phases. We present this case in accordance with the CARE reporting checklist (available at https://ales.amegroups.com/article/view/10.21037/ales-23-50/rc).

Case presentation

A 36-year-old man treated for hypertension over the previous 9 years developed hypokalemia (2.9 mmol/L). Blood tests revealed a high plasma aldosterone concentration (PAC) to plasma renin activity (PRA) ratio, and enhanced CT revealed a 15-mm low-density tumor in the left adrenal gland (*Figure 1*). The test results suggested PA; therefore, the patient was referred to our hospital for further examination.

The patient was severely obese, with a body weight of 136.0 kg and a body mass index (BMI) of 45.9 kg/m². He also had several obesity-related health issues including type 2 diabetes (T2D), obstructive sleep apnea, and hyperuricemia. He was being treated with 6 different antihypertensive drugs, but his systolic and diastolic blood pressures still measured 167 and 112 mmHg, respectively. Blood tests showed a high PAC (192 pg/mL) and PAC/PRA ratio [480]. The patient's hemoglobin A1C (HbA1c) level was also high (6.4%). Both the furosemide upright and captopril challenge

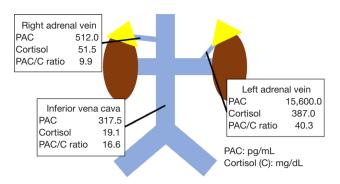


Figure 2 Schema demonstrating the results of adrenal vein sampling. Adrenal vein sampling after administration of adrenocorticotropic hormone demonstrates unilateral PAC elevation in the left adrenal vein. PAC, plasma aldosterone concentration; C, cortisol.



Figure 3 Intraoperative findings of laparoscopic sleeve gastrectomy. The gastric tube is created using linear staplers while an upper endoscopy enables observation.

tests were positive, but the dexamethasone suppression test was negative. AVS performed after adrenocorticotropic hormone administration showed unilateral PAC elevation in the left adrenal vein (*Figure 2*). Based on these results, the patient was diagnosed with PA caused by an adrenal adenoma localized in the left adrenal gland.

Laparoscopic adrenalectomy was considered, but we were concerned about the surgical procedure's difficulty and perioperative complications associated with severe obesity. To address these worries, we chose to perform LSG before laparoscopic adrenalectomy according to a surgical method we previously reported (9) (*Figure 3*). During the procedure, we used a liquid antiadhesive agent (AdSpray; Terumo Corporation, Tokyo, Japan) before closing the incision to prevent intra-abdominal adhesions.

Six months after the LSG, the patient's body weight and BMI decreased to 101.0 kg and 34.1 kg/m², respectively, and the number of antihypertensive drugs used decreased to four. In addition, the T2D symptoms resolved dramatically, with an HbA1c level of 5.4%, and improved insulin resistance. CT volumetry showed that the subcutaneous and visceral fat areas decreased significantly after LSG (*Figure 4*). As a result, we performed a laparoscopic adrenalectomy once substantial weight loss and metabolic improvement had been achieved.

Under general anesthesia, the patient was placed in the right lateral decubitus position, and three trocars were inserted. Laparoscopic adrenalectomy was performed via

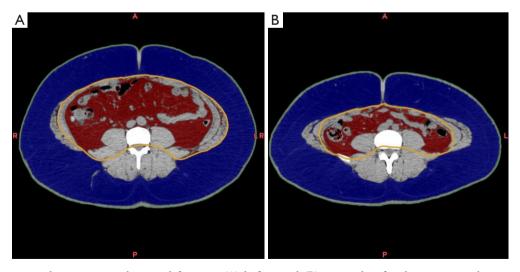


Figure 4 Changes in subcutaneous and visceral fat areas (A) before and (B) 6 months after laparoscopic sleeve gastrectomy. The subcutaneous fat area decreased from 690.1 to 581.5 cm², and the visceral fat area decreased from 223.2 to 82.6 cm².

the abdominal cavity, and only a small number of intraabdominal adhesions were observed (Figure 5). After mobilizing the descending colon and spleen, the left adrenal glands and surrounding tissue were identified and dissected. The central vein of the left adrenal gland was clipped and divided. Finally, the resected left adrenal gland was placed in a plastic bag and extracted via a 12-mm trocar wound on the left upper abdomen. Macroscopic findings revealed a 15-mm yellow tumor in the adrenal gland (Figure 6A), while, histopathological examination showed clear cell growth and no evidence of malignancy (Figure 6B). Therefore, adrenal adenoma, compatible with an aldosterone-producing tumor, was finally diagnosed. The patient's postoperative course was uneventful, and he was discharged on postoperative day 10. Twelve months after LSG (6 months after laparoscopic adrenalectomy), the patient's blood pressure was stabilized with only one antihypertensive drug, and



Figure 5 Intraoperative findings of laparoscopic adrenalectomy show only a small number of intra-abdominal adhesions.

further weight loss and improvement in insulin resistance were achieved. *Table 1* summarizes the treatment course.

All procedures performed in this study were in accordance with the ethical standards of the institutional and/or national research committee(s) and with the Helsinki Declaration (as revised in 2013). Written informed consent was obtained from the patient for the publication of this case report and accompanying images. A copy of the written consent is available for review by the editorial office of this journal.

Discussion

In the present study, we report a rational two-stage surgical strategy for patients suffering from severe obesity complicated by PA. Shinoda *et al.* reported the efficacy of LSG before laparoscopic adrenalectomy in patients with severe obesity complicated by PA (10) and *Table 2* summarizes the comparison with our case. All three patients were male, and their BMIs before LSG were between 35.9 and 45.9 kg/m². Subsequently, all cases underwent laparoscopic adrenalectomy within 1 year after LSG, and the BMIs at laparoscopic adrenalectomy were between 27.7 and 34.1 kg/m². Furthermore, treatment with antihypertensive drugs was reduced, and T2D symptoms improved after LSG, which contributed dramatically to the reduction in perioperative risk in severely obese patients with PA awaiting laparoscopic adrenalectomy.

In PA caused by a unilateral aldosterone-producing tumor, adrenalectomy is usually performed as a radical treatment (1). However, obesity affects the therapeutic

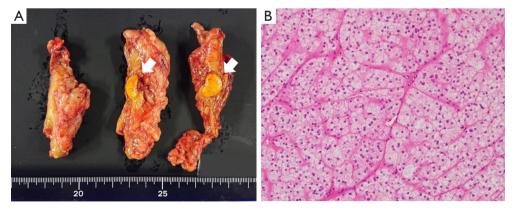


Figure 6 Macroscopic and histopathological findings. (A) Macroscopic findings reveal a well-defined yellow tumor of 15 mm in the adrenal gland (white arrows). (B) Histopathological findings show the growth of clear cells and no atypical cells suggestive of malignancy (hematoxylin and eosin, ×200).

Table 1 Treatment course of the patient from before LSG to 12 months after LSG

Variables	Before LSG	6 months after LSG	12 months after LSG	
Body weight (kg)	136.0	101.0	97.8	
BMI (kg/m²)	45.9	34.1	32.7	
SBP (mmHg)	167	128	130	
DBP (mmHg)	112	80	93	
Antihypertensive drug (mg/day)				
Azilsartan	40	20	-	
Amlodipine	10	10	10	
Carvedilol	20	10	-	
Doxazosin	4	-	_	
Trichlormethiazide	2	-	-	
Methyldopa	500	-	-	
Spironolactone	-	50	_	
PAC (pg/mL)	192	200	41.4	
PRA (ng/mL/h)	0.4	0.3	0.6	
BS (mg/dL)	142	97	101	
HbA1c (%)	6.4	5.4	5.0	
HOMA-IR (no unit)	5.0	3.2	2.4	
Insulinogenic index (no unit)	0.5	1.0	1.0	
Matsuda index (no unit)	1.8	2.4	3.4	
Disposition index (no unit)	0.9	2.4	3.2	

LSG, laparoscopic sleeve gastrectomy; BMI, body mass index; SBP, systolic blood pressure; DBP, diastolic blood pressure; PAC, plasma aldosterone concentration; PRA, plasma renin activity; BS, blood sugar; HbA1c, hemoglobin A1C; HOMA-IR, homeostatic model assessment for insulin resistance.

Table 2 Comparison between our patient and two severely obese patients with primary aldosteronism reported by Shinoda et al. and our patient

Case	First author	Age (year)	Gender	LSG		Interval between	LA			
				BMI (kg/m²)	Number of AD	HbA1c (%)	LSG and LA (months)	BMI (kg/m²)	Number of AD	HbA1c (%)
1	Shinoda M (10)	43	Male	35.9	_	6.6	8	27.7	0	5.4
2	Shinoda M (10)	51	Male	39.0	-	8.9	11	32.1	-	6.0
3	Kumagai H	36	Male	45.9	6	6.4	6	34.1	4	5.4

BMI, body mass index; LSG, laparoscopic sleeve gastrectomy; AD, antihypertensive drug; HbA1c, hemoglobin A1C; LA, laparoscopic adrenalectomy.

efficacy of adrenalectomy (2,3). BiLiGe *et al.* reported that the odds ratio in patients with BMIs from 25 to 29.9 kg/m² in uncured hypertension after adrenalectomy was 2.97, and 6.42 in patients with a BMI \geq 30 kg/m² (3). Thus,

we should recognize the importance of surgical procedures and weight loss in treating obese patients with PA. In addition, these retrospective observational studies reported that the number of antihypertensive drugs administered before adrenalectomy affected the postoperative course of PA treatment (2,3). In our case, and that reported by Shinoda *et al.* (10), a decrease in antihypertensive drugs was achieved after LSG. For severely obese patients with PA, performing LSG before adrenalectomy is considered an effective treatment strategy to optimize therapeutic efficacy after adrenal surgery.

Patients suffering from obesity are at risk of experiencing difficulty during surgical procedures due to the increase in visceral fat associated with excess weight. Additionally, obesity can increase postoperative complications such as respiratory failure, deep vein thrombosis, and surgical site infection (5). Therefore, weight reduction before radical surgery for the primary disease is important for reducing perioperative complications. LSG is an effective treatment option for these complications. We have previously reported the weight loss and metabolic improvement effects of LSG in severely obese patients (4,9,11,12). Weight loss occurred rapidly from 6 to 12 months after LSG (excess weight loss: 51.4% at 6 months, and 55.8% at 12 months) (11). The T2D remission rate was 70% to 80% in severely obese patients within 1 year after surgery (11,12), and obstructive sleep apnea also improved dramatically within 1 year after surgery (apnea-hypopnea index: 51.1 events/h at baseline, 25.4 events/h at 6 months, and 23.0 events/h at 12 months, respectively) (11). In addition, we encountered severely obese patients who underwent LSG before radical surgery for pancreatic neuroendocrine neoplasm and abdominal aortic aneurysm, and reported its efficacy (5,7,8). In our previously reported cases, radical surgery for the primary disease was performed 6 to 15 months after LSG, and the postoperative course was uneventful. These results suggest that the appropriate time for radical surgery for the primary disease is 6 months after LSG. However, strict perioperative blood pressure management is essential before LSG in patients suffering from severe obesity complicated by PA. In our hospital, a multidisciplinary obesity treatment team, including surgeons, endocrinologists, diabetologists, registered dietitians, and physical therapists treats severely obese patients. In this study, the obesity treatment team chose, after repeated discussions, to perform LSG before adrenalectomy. Under the management of a skilled obesity treatment team, a two-stage treatment strategy that indicates LSG before adrenalectomy may be a safe treatment option for severely obese patients with PA.

Laparoscopic minimally invasive surgery is the recognized

standard of care for unilateral aldosterone-producing tumors. We have previously reported on the technical safety, cosmetic benefits, and potential for postoperative pain reduction of laparoscopic single-site surgery for unilateral adrenal tumors (13,14). In this case series, an additional 5 mm port was required in two patients with BMIs >27 kg/m²; thus, we chose conventional multiport laparoscopic adrenalectomy for the present patient. Recently, the safety and efficacy of partial adrenalectomy and robotic-assisted surgery for unilateral aldosterone-producing tumors have been reported (15-17). Compared to total adrenalectomy, partial adrenalectomy, which can be expected to preserve adrenal function, has been suggested to shorten the operative time and length of hospital stay and reduce the overall postoperative complication rate (15,16). Prospective validation by case series or randomized control studies is needed in the future.

In summary, performing LSG before adrenalectomy is a rational strategy for severely obese patients with PA to reduce the risk of perioperative complications and improve therapeutic efficacy after adrenal surgery. However, strict blood pressure control is required during the LSG-to-adrenalectomy perioperative period. In addition, it is clear that patients with severe obesity typically suffer from many obesity-related diseases, and that comprehensive disease management by skilled obesity treatment teams, including surgeons and endocrinologists, is essential.

Conclusions

A two-stage treatment strategy that indicates LSG before adrenalectomy can be a rational treatment strategy for severely obese patients with PA. This approach may reduce the risk of perioperative complications and enhance therapeutic efficacy after adrenalectomy. Severely obese patients typically have hypertension associated with PA, as well as other obesity-related diseases, and comprehensive disease management by a skilled obesity treatment team is essential for achieving treatment success.

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Footnote

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Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at https://ales.amegroups.com/article/view/10.21037/ales-23-50/coif). The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. All procedures performed in this study were in accordance with the ethical standards of the institutional and/or national research committee(s) and with the Helsinki Declaration (as revised in 2013). Written informed consent was obtained from the patient for the publication of this case report and accompanying images. A copy of the written consent is available for review by the editorial office of this journal.

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