



Video-assisted thoracoscopic surgery for ectopic mediastinal parathyroid tumor: subxiphoid and lateral thoracic approach

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Background: This study aimed to investigate the initial results of an endoscopic surgical approach for the treatment of intramediastinal ectopic parathyroid adenoma and to evaluate the effectiveness of a single-incision resection using the subxiphoid approach.

Methods: Five cases of patients (1.89%) were diagnosed with ectopic mediastinal parathyroid tumor and underwent resection from 2008 to 2017 in Fujita Health University Hospital. They were retrospectively analyzed.

Results: Four patients underwent single-port mediastinal tumor resection using the subxiphoid approach and 1 patient underwent multi-port mediastinal tumor resection using the lateral thoracic approach. The operation time was 134 ± 83.52 min, and the amount of blood loss was 81.8 ± 173.41 mL. The rate of conversion to thoracotomy was 0%, and no intraoperative or postoperative complications were observed. The amount of postoperative oral analgesics was 112.83 ± 209.12 tablets, and their administration period was $561.6 \pm 1,229.5$ days. The length of hospital stay was 4 ± 2.35 days, and the duration of chest tube drainage was 1.33 ± 1.95 days. The patient who underwent multi-port mediastinal tumor resection using the lateral thoracic approach reported postoperative pain. Serum calcium levels decreased from 10.56 ± 1.52 mg/dL preoperatively to 8.96 ± 0.5 mg/dL postoperatively, and serum phosphorous levels increased from 2.84 ± 0.42 mg/dL preoperatively to 3.6 ± 0.51 mg/dL postoperatively. Intact-PTH hormone levels decreased from 221 ± 169.84 pg/dL preoperatively to 70.2 ± 44.28 pg/dL postoperatively. No recurrence of hyperparathyroidism has been observed in any patient.

Conclusions: The single-incision mediastinal tumor resection via the subxiphoid approach, without going through the intercostal space, is considered as a useful endoscopic surgical approach for the treatment of mediastinal ectopic parathyroid adenomas due to the limited occurrence of post-thoracotomy pain syndrome and the superior esthetic outcomes associated with the procedure as compared to thoracotomy and median sternotomy.

Keywords: Ectopic parathyroid tumor; mediastinal tumor; subxiphoid; video-assisted thoracoscopic surgery (VATS)

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Introduction

Ectopic mediastinal parathyroid tumor, which is one of the causes of primary hyperparathyroidism, accounts for 0.9% of all mediastinal tumors (1). Surgical approaches for mediastinal tumors include median sternotomy, lateral thoracotomy, mediastinoscopy, manubriotomy, transcervical incision, and thoracoscopic surgery using the lateral and subxiphoid approaches. Currently, thoracoscopic surgery using the lateral thoracic approach is the most common procedures (2-8). We have been performing multi-port mediastinal tumor resection using the lateral thoracic approach since 2008 and single-port mediastinal tumor resection using the subxiphoid approach since 2011 for mediastinal tumors (9). This study aimed to investigate the initial results of an endoscopic surgical approach for the treatment of intramediastinal ectopic parathyroid adenoma and to evaluate the effectiveness of a single-incision resection using the subxiphoid approach.

Methods

A total of 265 patients with mediastinal tumor underwent thoracoscopic surgery using the lateral or subxiphoid approach from 2008 to 2017 in Fujita Health University Hospital. Five (1.89%) of 265 patients who were diagnosed with ectopic mediastinal parathyroid tumor were included in the study and were retrospectively analyzed.

We examined the presence or absence of preoperative clinical symptoms, preoperative chest contrast-enhanced computed tomography findings, preoperative ^{99m}Tc -methoxyisobutylisonitrile scintigraphy (^{99m}Tc -MIBI scintigraphy) results, pre- and postoperative serum calcium, phosphorous, and intact-PTH hormone levels, operative time, the amount of intraoperative blood loss, the rate of conversion to thoracotomy, intra- and postoperative complications, the amount of postoperative oral analgesics, the administration period of postoperative oral analgesic, the duration of chest tube drainage, the length of hospital stay and pathological examination findings. Data were presented as mean values \pm standard deviation. This study was approved by the Institutional Review Board of Fujita Health University (No. HM18-194).

Surgical technique

The operations were performed under general anesthesia. A single-lumen endotracheal tube was used for single-

port mediastinal tumor resection using the subxiphoid approach and a double-lumen endotracheal tube for multi-port mediastinal tumor resection using the lateral thoracic approach. It has been reported that seeding of parathyroid adenomas can easily occur (10). Therefore, when performing thymectomy for tumors that had developed in the thymus, we resected the tumors en bloc with surrounding fatty tissues and ensured the actual tumor was not touched. The resected tumors were placed in a bag in the thoracic cavity and removed from the surgical wound.

The multi-port mediastinal tumor resection using the lateral thoracic approach was performed with the patient in lateral decubitus position. The operation was performed using three ports. Skin incisions of 1.5 cm were made in the third intercostal space along the middle axillary line, the fifth intercostal space along the anterior axillary line, and the seventh intercostal space along the posterior axillary line.

We performed single-port mediastinal tumor resection using the subxiphoid approach with the same techniques that we previously reported for thymectomy (9). The patient was placed in supine position with legs open. A 3-cm transverse incision was made 1 cm caudally from the xiphoid process. GelPOINT[®]Mini (Applied Medical Resources Co., Rancho Santa Margarita, CA, USA) was inserted at the incision site, and three subports were inserted into the GelPOINT[®]Mini platform. A camera scope and forceps were inserted into the three subports, and the operation was performed.

CO₂ insufflation of the mediastinum was performed at 8 mmHg. Both sides of the mediastinal pleura were transected to open both sides of the thoracic cavity. The left and right phrenic nerves were then identified and separated from the pericardium, including the anterior mediastinal pleura, thymus, and surrounding fatty tissue. The cervical region of the thymus was separated at the cranial side from the thyroid gland, on the right side from the right brachiocephalic vein, on the left side from the innominate vein, and on the dorsal side from the brachiocephalic artery and trachea. Finally, the thymus was pulled to either the left or right side. The thymic vein was transected, and the ectopic mediastinal parathyroid tumor with the thymus was removed (*Figure 1*).

Results

Table 1 summarizes the characteristics of the 5 patients (mean age: 50.6 \pm 14.79 years; 1 male and 4 females) with ectopic mediastinal parathyroid tumor who underwent

thoracoscopic surgery. Four patients presented with the following symptoms prior to surgery: weight loss, poor appetite, fatigue, knee pain, lower back pain, pain in the right abdominal region, hematuria, and facial edema. The tumor was located at two sites (superior and anterior mediastinum) in 1 patient, in the superior mediastinum in 1, in the anterior mediastinum in 2, and in the middle mediastinum in 1. Imaging revealed a mean tumor diameter of 1.83 ± 0.85 cm. Preoperative ^{99m}Tc -MIBI scintigraphy was performed in all patients. While uptake in the tumor was confirmed in 4 patients, only a slight uptake was noted in 1 patient. Serum calcium, phosphorous, and intact-

PTH hormone levels were measured via preoperative blood testing. Hypercalcemia was observed in 4 patients, with serum calcium levels measured at 10.56 ± 1.52 mg/dL. Hypophosphatemia was observed in 1 patient, with serum phosphorous levels measured at 2.84 ± 0.42 mg/dL. Intact-PTH hormone levels were elevated at 221 ± 169.84 pg/dL in all patients.

Table 2 summarizes the intra- and postoperative findings. Single-port mediastinal tumor resection using the subxiphoid approach was performed in 4 patients, whereas multi-port mediastinal tumor resection using the lateral thoracic approach was performed in 1 patient. The total operative time was 134 ± 83.52 min, and the amount of blood loss was 81.8 ± 173.41 mL. In multi-port mediastinal tumor resection using the lateral thoracic approach, the operative time was 255 min, and the amount of blood loss was 392 mL. Meanwhile, in single-port mediastinal tumor resection using the subxiphoid approach, the operative time was 103.75 ± 56.56 min, and the amount of blood loss was 4.25 ± 1.5 mL. In one patient who underwent multi-port mediastinal tumor resection using the lateral thoracic approach and a different operation at another hospital, left lung was completely adherent to the thoracic cavity. Therefore, the operative time was longer, and a higher amount of blood loss was observed. The conversion to thoracotomy intraoperatively and intraoperative complications was not observed in any patient.

The amount of postoperative oral analgesics was 135.4 ± 209.12 tablets, and their administration period



Figure 1 Single-port mediastinal tumor using the subxiphoid approach (11). Thymectomy was performed as the tumor was located near the innominate vein.

Available online: <http://http://www.asvide.com/watch/32887>

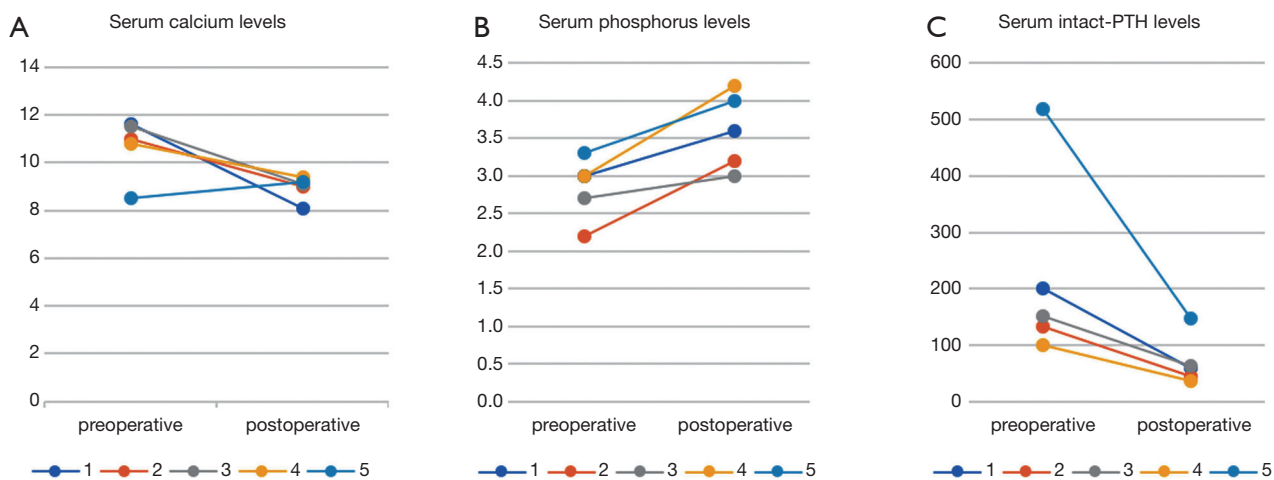
Table 1 Preoperative data of the patients

| Clinical date | Case 1 | Case 2 | Case 3 | Case 4 | Case 5 |
|--------------------------------------|--------------------|----------------------|----------------------|----------------------|--|
| Age (years) | 57 | 63 | 25 | 53 | 55 |
| Gender | Female | Male | Female | Female | Female |
| Symptom | + | - | + | + | + |
| Tumor location | Middle mediastinum | Anterior mediastinum | Anterior mediastinum | Superior mediastinum | Superior mediastinum, anterior mediastinum |
| Tumor size (cm) | 1.5 | 1.8 | 1.2 | 0.8 | Superior: 2.8, anterior: 2.9 |
| ^{99m}Tc -MIBI scintigraphy | Positive | Positive | Positive | Positive | Negative |
| Serum calcium level (mg/dL) | 11.6 | 11 | 11.5 | 10.8 | 7.9 |
| Serum phosphorus level (mg/dL) | 3 | 2.2 | 2.7 | 3 | 3.3 |
| Serum intact-PTH level (pg/dL) | 200 | 134 | 152 | 101 | 518 |

^{99m}Tc -MIBI, ^{99m}Tc -methoxyisobutylisonitrile; PTH, parathyroid hormone.

Table 2 Intra- and postoperative findings

| Intra- and postoperative findings | Case 1 | Case 2 | Case 3 | Case 4 | Case 5 |
|--|---------------------|---------------------|---------------------|---------------------|------------------|
| Operation | Multiport | Subxiphoid | Subxiphoid | Subxiphoid | Subxiphoid |
| Operative time (min) | 255 | 57 | 83 | 89 | 186 |
| Amount of blood loss (mL) | 392 | 5 | 2 | 5 | 5 |
| Conversion to thoracotomy | - | - | - | - | - |
| Intraoperative complication | - | - | - | - | - |
| Amount of postoperative oral analgesics (tablet) | 509 | 42 | 27 | 42 | 57 |
| Administration period (day) | 2,761 | 13 | 8 | 16 | 10 |
| Duration of chest tube drainage (day) | 5 | 1 | 0 | 1 | 1 |
| Postoperative complication | + | - | - | - | - |
| Length of hospital stay (day) | 8 | 2 | 4 | 3 | 3 |
| Pathological examination findings | Parathyroid adenoma | Parathyroid adenoma | Parathyroid adenoma | Parathyroid adenoma | Parathyroid cyst |

**Figure 2** Pre- and postoperative blood examination results. (A) Serum calcium levels, (B) serum phosphorus levels, (C) serum intact-PTH levels.

was $561.6 \pm 1,229.5$ days. In multi-port mediastinal tumor resection using the lateral thoracic approach, 509 tablets were used, and the administration period was 2,761 days. Conversely, in single-port mediastinal tumor resection using the subxiphoid approach, 42 ± 12.25 tablets were used, and the administration period was 11.75 ± 3.5 days. While no postoperative pain (12) was observed in cases 2 through 5, 1 patient who underwent surgery using the lateral thoracic approach reported postoperative pain. After 7.5 years from operation, the patient still experiences pain.

The duration of chest tube drainage was 1.6 ± 1.95 days,

and the length of hospital stay was 4 ± 2.35 days. In multi-port mediastinal tumor resection using the lateral thoracic approach, the duration of chest tube drainage was 5 days, and the length of hospital stay was 8 days. Conversely, in single-port mediastinal tumor resection using the subxiphoid approach, the duration of chest tube drainage was 5 days, and the length of hospital stay was 0.75 ± 0.5 and 3 ± 0.82 days. Pathological examination findings indicated parathyroid adenoma in 4 patients and parathyroid cyst in 1 patient.

Figure 2 summarizes the pre- and postoperative serum

calcium, phosphorous, and intact-PTH hormone levels in all patients. These levels improved postoperatively in all patients. Serum calcium levels decreased from 10.56 ± 1.52 mg/dL preoperatively to 8.96 ± 0.5 mg/dL postoperatively, and serum phosphorous levels increased from 2.84 ± 0.42 mg/dL preoperatively to 3.6 ± 0.51 mg/dL postoperatively. Moreover, intact-PTH hormone levels decreased from 221 ± 169.84 pg/dL preoperatively to 70.2 ± 44.28 pg/dL postoperatively. Recurrence was not observed in any of the cases.

Discussion

Parathyroid adenomas are the most common causes of primary hyperparathyroidism. While most of these adenomas develop near the thyroid gland, 16% are ectopic and develop in sites such as the mediastinum, pharynx, back surface of the esophagus, near the aortic arch, or around the pericardium (13,14). The frequency of parathyroid adenoma arising from a mediastinal tumor is 0.9%.

Approximately 67–80% of ectopic mediastinal parathyroid tumors are located in the thymus (15,16). In 4 patients in our study, the tumor was located in the thymus. Approaches to the thymus include median sternotomy, lateral thoracic thoracotomy, mediastinoscopy, manubriotomy, cervical collar incision, and thoracoscopic surgeries using the lateral and subxiphoid approaches. Recently, thoracoscopic surgery using the lateral thoracic approach has become the primary option for minimally invasive surgery (2–8). However, intercostal neuralgia is a common complication associated with thoracoscopic surgery performed using the lateral thoracic approach (12). While we performed multi-port mediastinal tumor resection using the lateral thoracic approach for mediastinal tumors from 2008 to 2010, we started using the single-port mediastinal tumor resection using the subxiphoid approach for such cases in 2011 (9). Because single-port mediastinal tumor resection using the subxiphoid approach does not require an intercostal approach, procedures such as thoracoscopic surgery do not cause intercostal neuralgia from the lateral thoracic region, which is considered its advantage (17). In fact, a study comparing postoperative pain in an intercostal thoracoscopic surgery group and a subxiphoid surgery group found that visual analog scale scores were significantly lower in the subxiphoid surgery group than in the intercostal thoracoscopic surgery group (18). In our study, while the postoperative oral analgesics were not necessary on postoperative day 16 for patients who

underwent single-port mediastinal tumor resection using the subxiphoid approach, the patient who underwent multi-port mediastinal tumor resection using the lateral thoracic approach presented with postoperative pain and required long-term administration of oral analgesics. The case of the lateral thoracic approach in our study can not be compared with the subxiphoid approach for postoperative pain, because there was a history of ipsilateral surgery in another hospital, and the left lung was completely adherent to the thoracic cavity. However, the single-incision mediastinal tumor resection via the subxiphoid approach, without going through the intercostal space, is a very useful approach for patients in terms of postoperative pain (17). Due to the fact that it is a single-port approach, it has advantages in terms of cosmetics (19). A study comparing single-incision thoracoscopic surgery and three port video-assisted thoracic surgery in terms of patient satisfaction with the wound site has reported that patient satisfaction with single-incision thoracoscopic surgery was significantly higher (20).

No significant differences were noted between the procedures in terms of the duration of chest tube drainage or length of hospital stay. Amer *et al.* have reported that the duration of chest tube drainage and length of hospital stay in three port video-assisted thoracoscopic surgery for ectopic mediastinal parathyroid tumor were 1 ± 0.57 and 2 ± 2.36 days, respectively (21). Thus, no significant differences were noted in terms of the duration of chest tube drainage and length of hospital stay in single-port mediastinal tumor resection using the subxiphoid approach.

When treating parathyroid adenomas, postoperative metastasis and subsequent recurrence can easily occur as a result of intraoperative damage to the tumor capsule (10,22). Rattner *et al.* have reported that extreme caution is required during surgery to prevent such metastasis and damage to the capsule or touching the parathyroid tissue (10). Moreover, Akerström *et al.* have reported that in addition to performing intraoperative procedures with caution, surgeons must not divide the parathyroid gland in the surgical field (22). Thymectomy is not required for patients with ectopic mediastinal parathyroid tumor. However, if sufficient tumor margins cannot be secured or if the tumor is close to the innominate vein and the site of this vein must be confirmed to safely perform surgery, thymectomy may be performed. In all the 5 patients in our study, the thymus was completely resected without grasping the tumor or area around it intraoperatively, thereby preventing damage in the tumor capsule. Taking this into consideration, we believe that any recurrence of hyperparathyroidism was

postoperatively prevented.

Conclusions

We performed thoracoscopic surgery in patients with ectopic mediastinal parathyroid tumor and obtained satisfactory results in terms of postoperative pain and cosmetic outcomes. Single-port mediastinal tumor resection using the subxiphoid approach did not result in postoperative pain in 4 patients. However, 1 patient who underwent multi-port mediastinal tumor resection using the lateral thoracic approach reported postoperative pain. The single-incision mediastinal tumor resection via the subxiphoid approach, without going through the intercostal space, is considered as a useful endoscopic surgical approach for the treatment of mediastinal ectopic parathyroid adenomas due to the limited occurrence of post-thoracotomy pain syndrome and the superior esthetic outcomes associated with the procedure as compared to thoracotomy and median sternotomy.

Acknowledgments

None.

Footnote

Conflicts of Interest: 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. This study was approved by the Institutional Review Board of Fujita Health University (No. HM18-194).

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