Direct endoscopic full-thickness resection for submucosal tumors with an intraluminal growth pattern originating from the muscularis propria layer in the gastric fundus

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Abstract

Background and aims: Endoscopic full-thickness resection (EFTR) is difficult to perform in a retroflexed fashion in the gastric fundus. The present study aims at exploring whether direct EFTR can be a simple, effective and safe procedure to treat intraluminal-growth submucosal tumors originating from the muscularis propria.

Methods: The patients with intraluminal-growth submucosal tumors originating from the muscularis propria in gastric fundus treated by direct EFTR between 01 January 2017 and 01 September 2018 were retrospectively reviewed. In addition, we analyzed the patients with intraluminal-growth submucosal tumors originating from the muscularis propria in gastric fundus treated by traditional EFTR. The differences in tumor resection time, cost-effectiveness, and complication rate were evaluated.

Results: Forty patients were enrolled in the present study, 20 patients of which were in the direct EFTR group and 20 patients of which were in the traditional EFTR group. En-bloc resections of gastric tumors were successfully performed in all 40 cases. There was no significant difference in the average tumor size of the two groups (24.3 ± 2.9 mm in direct EFTR group verus 24.0 ± 2.6 mm in the traditional group, \( p = 0.731 \)), but significant difference existed in the operative time between two groups (35.0 ± 8.2 min in direct EFTR group verus 130.6 ± 51.9 min in the traditional group, \( p < 0.05 \)). No complications, such as postoperative bleeding and perforation, occurred in any groups.

Conclusions: Direct EFTR is a safe, simple and cost-effective procedure for SMTs with an intraluminal growth pattern originating from the muscularis propria layer in the gastric fundus.

Keywords: Gastric submucosal tumor, Endoscopic full thickness resection, Over the scope clip

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Background
Endoscopic full-thickness resection (EFTR) is applied to resect gastrointestinal submucosal tumors (SMTs) [1]. Compared with surgery and laparoscopy, EFTR could be a more minimally invasive choice for patients and provide the same therapeutic effect for gastric SMTs [2]. However, EFTR is a technical challenging procedure, especially when the lesion is located in the gastric fundus because of retroflexion of the endoscope and specific anatomical features. In order to simplify the procedure of EFTR, loop-mediated, rope-mediated or dental floss traction-assisted EFTR for SMTs in the gastric fundus has been reported [3, 4]. For intraluminal-growth SMTs originating from the muscularis propria in gastric fundus, it can be resected by a more simple method, direct EFTR.

The aim of this study was to evaluate the usefulness of direct EFTR for the resection of intraluminal-growth SMTs originating from the muscularis propria in gastric fundus.

Methods
Patients
The patients with intraluminal-growth SMTs originating from the muscularis propria in gastric fundus treated by direct EFTR between 2017/01/01 and 2018/09/01 were retrospectively reviewed; these patients were defined as the trial group. Based on the tumor size and growth pattern, the patients with similar gastric SMTs treated by traditional EFTR were defined as the control group. All the lesions were evaluated by preoperative endoscopic ultrasound (EUS) and computed tomography. Intraluminal growth pattern was defined as: the lesion was of intraluminal growth under endoscopic view and EUS and computed tomography showed that the whole lesion is in the side of gastric cavity.

All the patients signed the informed consent and this study was in accordance with Helsinki Declaration and approved by the ethics committee of Shengjing Hospital of China Medical University.

Endoscopic equipment and accessories
A standard single-channel gastroscope (EPK-i, Pentax) and transparent cap was used throughout the endoscopic procedure; A snare was used to resect the tumor in the trial group. A triangle-tipped knife and insulated-tip knife (both from Olympus Corporation, Tokyo, Japan) was used for the dissection and resection of tumors in the traditional EFTR group. During the procedure, A hot biopsy forceps (FD-410LR) was used for hemostasis. Metal clips (Olympus Corporation, Tokyo, Japan), and the OTSC system (Ovesco Endoscopy GmbH, Tuebingen, Germany) were used for defect closure.

Procedures
All direct EFTR procedures were performed by experienced experts, who can perform traditional EFTR. Direct EFTR was performed as follows (Figs. 1 and 2): First, we aspirated the gastric fluid to empty the stomach. Second, the snare was placed at the root of lesion and mild suction was applied to loosen the gastric wall and ensure the complete grasp of the tumor. After the snare was tightened, the tumor presented as a pseudo-polyp. If there was small amount of tissue in the snare and shaking snare could not continue to tighten the snare, it demonstrated complete grasp of the tumor. Then we resected the tumor and an iatrogenic gastric perforation was created. Third, after resection, electric cautery was used for hemostasis. Fourth, the iatrogenic gastric perforation was closed by metal clips or a metallic clip interrupted suture with an over the scope clip (OTSC). Traditional EFTR was performed as we previously reported [5].

Definition of outcomes
The primary outcome measure of this study was en-bloc resection rate. Secondary outcome measures included tumor resection time, cost-effectiveness and adverse events.

Fig. 1 a the lesion with intraluminal growth pattern was located in the gastric fundus. b the snare was placed at the root of lesion and mild suction was applied to loosen the gastric wall and ensure the complete grasp of the tumor. After the snare was tightened, the tumor presented as a pseudo-polyp.
Postoperative treatment and follow-up
The postoperative treatment included fasting, gastric tube decompression as well as routine administration of proton pump inhibitors and antibiotics. Patients were suggested for the endoscopic follow-up at 1, 3, 6, 12 months and annually thereafter to observe recurrence and wound healing.

Statistical analysis
Statistical analysis was performed with the SPSS software package version 19.0 (SPSS Inc., Chicago, IL, USA).

Results
In the direct EFTR group, there were 12 males and 8 females, with an average age of (52.6 ± 12.8) years. All the gastric SMTs were intraluminal growth confirmed by EUS, with an average size of 24.3 ± 2.9 mm. The en-bloc resection rate was 100%. Ten perforation wounds were sealed with over the scope clip (OTSC); ten were closed by OTSC and titanium clips. Operative time ranged from 12 to 47 mins, with mean of 35.0 ± 8.2 min. In the traditional EFTR group, there were 10 males and 10 females, with an average age of (54.6 ± 11.8) years. All the gastric SMTs were intraluminal growth confirmed by EUS, with an average size of 24.0 ± 2.6 mm. The en-bloc resection rate was 100%. 12 perforation wounds were sealed with over the scope clip (OTSC); Eight were closed by OTSC and titanium clips. Operative time ranged from 68 to 240 mins, with mean of 130.6 ± 51.9 min. There was no significant difference between the two groups about tumor size (p = 0.731), but the operative time of direct EFTR group was significantly less than that of traditional EFTR group (p < 0.05). In both groups, there were no postoperative complications, such as bleeding and perforation (Table 1). Of 40 tumors, there were 39 gastrointestinal stromal tumors and one leiomyoma. Complete resection of all the lesions was done.

Table 1 Demographic and clinical characteristics of the direct and traditional groups

| Item                        | Direct EFTR group(n=20) | Traditional EFTR group(n=20) | P value |
|-----------------------------|-------------------------|-----------------------------|---------|
| Mean age, year              | 52.6 ± 12.8             | 54.6 ± 11.8                 | P > 0.05|
| Gender (male: female)       | 12:8                    | 10:10                       |         |
| Tumor size, mm              | 24.3 ± 2.9              | 24.0 ± 2.6                  | P > 0.05|
| Operative time, minute      | 35.0 ± 8.2              | 130.6 ± 51.9                | P < 0.05|
| En-bloc resection rate      | 100%                    | 100%                        |         |
| Complications               |                         |                             |         |
| Postoperative perforation   | 0                       | 0                           |         |
| Postoperative bleeding      | 0                       | 0                           |         |

EFTR endoscopic full thickness resection
The costs between two groups were different. The cost comparison for the whole procedure was $23,352 \pm 512 \text{CNY}$ (traditional EFTR) vs $17,033 \pm 681 \text{CNY}$ (direct comparison for the whole procedure was $23,352 \pm 17,033 \text{CNY}$). There was a significant difference ($P < 0.05$).

**Discussion**

Gastric SMTs can be exactly diagnosed by EUS [6, 7]. Considering the malignant potential of some SMTs, especially when the tumor size is more than 20 mm, resection is recommended for the lesion. With the development of endoscopic equipment, EFTR has been widely used for gastric SMTs [8, 9].

The SMT located in gastric fundus sometimes is difficult to access, even with the retroflexion of the endoscope, which makes the resection procedure difficult. The traction-assisted EFTR has been reported to improve EFTR procedure in gastric fundus [3, 4]. There were some advantages of traction-assisted EFTR. First, traction can help to expose the tumor boundaries and make the operation filed clear, which can simplify the operation process. In addition, traction can help to quickly locate bleeding point during the procedure, then we can do hemostasis and this method can prevent accidental damage of extravascular vessels. What’s more, traction can help prevent the tumors from falling into abdominal cavity and help to remove the tumor. Although this method has been used for EFTR, EFTR is also a difficult procedure and should be performed by experienced endoscopists.

Usually, the gastric SMTs present as a slightly-protruded lesion. With the growth of SMTs, Some gastric SMTs form an intraluminal growth pattern, like a pseudo-stalk polyp, due to the gravity. For this kind lesion, the tumor is totally in gastric cavity, which can be confirmed by EUS. In our study, we performed direct EFTR for these lesions, like using a mucosa resection for a polyp, which can simply the procedure.

Iatrogenic perforation after EFTR can be sutured by endoscopic equipment, such as over the scope clip (OTSC). The OTSC has shown clinical results over conventional methods. However, the OTSC system also has a limited function in regard to the perforation size. In generally, OTSC can fully suture perforations<20 mm [10]. For a perforation>20 mm, complete closure sometimes can not be achieved by one OTSC, but complete closure can be finished by combining OTSC with metal clips. In our study, all 40 cases were successfully closed. There were several advantages of direct EFTR as follows: first, it can make EFTR more easily to perform and compared with traditional EFTR, it can be done in short time. Second, direct EFTR is cost-effective and we just used a snare to resect the lesion. Third, even if there is a bleeding after resection, we can locate the bleeding site easily and do hemostasis. Fourth, this technique can make sure the tumor intact capsule and avoid the damage of the tumor during the dissection.

However, when we used direct EFTR, we should pay attention to some points as follows: first, before resection, we must use EUS to confirm the tumor totally in gastric cavity, otherwise we may cut the tumor and increase the risk. Second, the procedure should be performed by the endoscopic doctor with the ability of hemostasis and closure of iatrogenic perforation. Now this technique can just be used for the tumor with an intraluminal growth pattern. In the future, the retrievable anchor may be used for this technique and pull the tumor back the gastric cavity and resect it by a snare. It should be proved by further studies.

Some limitations are present in our study. First, complete grasp of the tumor was evaluated by the endoscopic doctor’s experience. A more reasonable way should be investigated to judge it by further studies. Second, this is a retrospective and single-center study and less cases were included in the present study. Therefore, a multi-center, prospective and randomized study should be conducted.

**Conclusions**

In conclusion, direct EFTR is a safe, simple and cost-effective procedure for SMTs, with an intraluminal growth pattern originating from the muscularis propria layer in the gastric fundus.

**Abbreviations**

EFTR: Endoscopic full-thickness resection; EUS: Endoscopic ultrasound; OTSC: Over the scope clip; SMTs: Submucosal tumors

**Authors’ contributions**

JLH: study concept and design; acquisition of data; analysis and interpretation of data; drafting of the manuscript; NG, SW, JTG, XL and GXW: study concept and design; critical revision of the manuscript for important intellectual content; administrative, technical, or material support; SYS: study concept and design; analysis and interpretation of data; critical revision of the manuscript for important intellectual content administrative, technical, or material support; study supervision. All authors approved the final version of the manuscript.

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**Availability of data and materials**

The dataset supporting the conclusions of this article is available from the corresponding author on reasonable request.

**Ethics approval and consent to participate**

This was a retrospective study. It was approved by our institution’s Institutional Review Board (Shengjing Hospital of China Medical University) and written informed consent was obtained from all subjects.

**Consent for publication**

Not Applicable.
Competing interests
The authors declare that they have no competing interests.

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