**Name of journal: *World Journal of Gastrointestinal Endoscopy***

**ESPS Manuscript NO: 26052**

**Manuscript type: ORIGINAL ARTICLE**

***Retrospective Study***

**Transanal endoscopic microsurgery as optimal option in treatment of rare rectal lesions: a single centre experience**

Ortenzi M *et al*. TEM for rare rectal lesions

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**Author contributions:** Ortenzi M and Ghiselli R designed the study; Ortenzi M, Cappelletti Trombettoni MM and Cardinali L contributed to acquisition of data and drafting the article; Guerrieri M approved the final version to be published.

**Institutional review board statement:** The study was reviewed and approved by the Ospedali Riuniti Institutional Review Board.

**Informed consent statement:** All study participants, or their legal guardian, provided informed written consent prior to study enrollment.

**Conflict-of-interest statement:** No financial support or incentive has been provided for this manuscript. All authors have no conflicts of interest or financial ties to disclose.

**Data sharing statement:** Technical appendix, original data, and statistical code of manuscript NO 26052 submitted to World Journal of Gastrointestinal Endoscopy are available from the corresponding author at monica.ortenzi@gmail.com. Partecipants gave informed consent for data sharing.

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**Manuscript source:** Invited manuscript

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**Received:** March 28, 2016

**Peer-review started:** March 29, 2016

**First decision:** May 17, 2016

**Revised:** June 2, 2016

**Accepted:** July 11, 2016

**Article in press:**

**Published online:**

**Abstract**

**AIM:** To analyze the outcomes of transanal endoscopic microsurgery (TEM) in the treatment of rare rectal condition like mesenchymal tumors, condylomas, endometriosis and melanoma.

**METHODS:** We retrospectively reviewed a twenty-three years database. Fifty-two patients were enrolled in this study. The lesions were considered suitable for TEM if they were within 20 cm from the anus. All of them underwent an accurate preoperative workup consisting in clinical examination, total colonoscopy with biopsies, endoscopic ultrasonography, and pelvic computerized tomography or pelvic magnetic resonance imaging. Operative time, intraoperative complications, rate of conversion, tumor size, postoperative morbidity, mortality, the length of hospital stay, local and distant recurrence were analyzed.

**RESULTS:** Among the 1328 patients treated by TEM in our Department, the 52 patients with rectal abnormalities other than adenoma or adenocarcinoma represented 4.4%. There were 30 males (57.7%) and 22 females (42.3%). Mean age was 55 years (median = 60, range = 24-78). This series included 14 (26.9%) GISTs, 21 NETs (40.4%), 1 ganglioneuroma (1.9%), 2 solitary ulcers in the rectum (3.8%), 6 cases of rectal endometriosis (11.5%), 6 cases of rectal condylomatosis (11.5%) and 2 rectal melanomas (3.8%). Mean lesion diameter was 2.7 cm (median: 4, range: 0.4-8). Mean distance from the anal verge was 9.5 cm (median: 10, range: 4-15). One patient operated for rectal melanoma developed distant metastases and died two years after the operation. We experienced 2 local recurrences (3.8%) with an overall survival equal to 97.6% (95%CI: 95%-99%) at the end of follow up and a disease free survival of 98% (95%CI: 96%-99%).

**CONCLUSION:** We could conclude that TEM is an important therapeutical option for rectal rare conditions.

**Key words:** Transanal endoscopic microsurgery; Rare rectal conditions; minimally invasive surgery; Full-thickness excision; Retrospective study

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**Core tip:** This paper is about the management of rare rectal lesions by transanal endoscopic microsurgery (TEM). The rarity of these conditions and the lack of big reports about this topic make this work important. We focused our attention on operative data and postoperative long-term outcomes. Our results suggested that TEM is a safe, minimally invasive procedure that can be adopted for the treatment of these conditions with excellent results.

Ortenzi M, Ghiselli R, Cappelletti Trombettoni MM, Cardinali L, Guerrieri M. Transanal endoscopic microsurgery as optimal option in treatment of rare rectal lesions: a single centre experience. *World J Gastrointest Endosc* 2016; in press

**INTRODUCTION**

Adenocarcinoma is the most frequent malignancy of the rectum, but the distal part of the bowel can host several other rare lesions which together represent an important part of rectal tumors[1]. This heterogeneous group comprehends mesenchymal tumors like gastrointestinal stromal tumors (GISTs), Neuroendocrine tumors (NETs) and ganglioneuromas. Other abnormalities can involve the rectal wall, and surgery is the only curative option, as is also the case for condyloma, endometriosis and melanoma. The aim of this study was to analyze the results of TEM in the treatment of these rare rectal conditions.

**Materials and methods**

A retrospective accurate analysis of a twenty-two-year old database built from 1992 to 2015 identified 52 patients eligible for the study. Indications for TEM were determined on the basis of the anatomical criteria assessed by rigid preoperative rectoscopy in order to locate the lesions and to measure its distance from the anal verge.

All patients were properly informed about the operation and give their consensus to surgery. The lesions were considered suitable for TEM if they were within 20 cm from the anus. Preoperative workup included clinical examination, total colonoscopy with biopsies, endoscopic ultrasonography, and pelvic computerized tomography or pelvic magnetic resonance imaging. Patients’ characteristics such as age and gender were considered. All patients received similar pre-operative management with an oral intake of an osmotic solution the day before surgery and a short term intravenous antibiotics prophylaxis to provide coverage for the normal bowel flora, aerobic and anaerobic species.

Procedures were performed by the Wolf TEM equipment (Knittlingen, Germany) consisting of a rigid 12 or 20 cm long rectoscope, an endosurgical unit steadily controlling rectal endoluminal pressure, and curved instruments. In all cases, a full-thickness excision was performed, and the rectal defect was closed by a running suture secured with silver clips at the extremities.

The operative data examined included operative time, intraoperative complications and conversion to abdominal surgery. Tumor size was measured macroscopically and reported as the maximum diameter. Pathological examination included histopathological definition, degree of differentiation, macroscopical measurement, and the examination of radial margins of excision. A urinary catheter was placed in all the patients at the time of surgery, which was removed 24 h after the operation. In the post-operative period, we analyzed postoperative morbidity, mortality and the length of hospital stay. Long-term outcomes included local and distant recurrence. We considered as local recurrence any recurrence diagnosed endoscopically and confirmed by biopsy. Follow- up included digital examination, rigid rectoscopy and endorectal ultrasound every 6 mo for the first year from the time of operation and subsequently every year.

Quantitative variables are shown as the mean value with median and range in brackets. Recurrence-free survival was considered as a continuous variable. The probability of overall survival at the end of follow up and the probability of disease-free survival were estimated using the Kaplan-Meier method. All analyses were performed using the R statistical package.

**RESULTS**

Among the 1328 patients treated by TEM in our Department, the 52 patients with rectal abnormalities other than adenoma or adenocarcinoma represented 4.4%. There were 30 males (57.7%) and 22 females (42.3%). Mean age was 55 years (median = 60, range = 24-78). We excised, by TEM, 14 (26.9%) GISTs, 21 NETs (40.4%), 1 ganglioneuroma (1.9%) and 2 solitary ulcers in the rectum (3.8%). We used TEM to treat 6 cases of rectal endometriosis (11.5%), 6 cases of rectal condilomatosis (11.5%) and 2 rectal melanomas (3.8%).

Preoperative symptoms ranged from rectal bleeding (9/52, 17.3%), urgency (3/52, 5.8%) and alteration in bowel habit (7/52, 13.5%). Thirty-two (61.5%) patients were asymptomatic and the lesions were discovered incidentally. Mean lesion diameter was 2.7 cm (median: 4, range: 0.4-8). Mean distance from the anal verge was 9.5 cm (median: 10, range: 4-15) (Table 1).

GISTs had a mean diameter of 1.4 cm (median = 1, range = 0.4-5). Two of them received neoadjuvant Imatinib resulting in reduction in tumor size. Six GISTs were defined as medium risk GISTs and 4 as high risk.

As for NETs, the mean lesion diameter was 2.7 cm (median = 2, range = 0.5-5). Except for one, all of them were G1 well differentiated NETs. There was only one ganglioneuroma which extended circumferentially on the rectal wall and had a diameter of 10 cm. The condyloma had a mean diameter of 2.7 cm (median: 3, range: 2-3). The 2 solitary ulcers had a diameter of 3 and 4 cm respectively and were completely excised.

Complete resection with disease-free margins was achieved in all the cases except for one case in which the pathologist was unable to assess the margin due to thermal damage. Mean operative time was 41 min (median: 45, range: 20–55). There was no conversion to abdominal surgery. We observed one intraoperative minor complication (1.9%) consisting in rectal bleeding controlled by TEM.

We observed a postoperative morbidity rate of 3.8 % (2/50), consisting of one case of acute urinary retention and one case of mild incontinence to gas resolved within two months from the operation by means of physiotherapy. Mean hospital stay was 3 d (median: 4, range: 2-7).

All the patients completed the follow-up protocol, including clinical and instrumental assessment. Two patients (3.8%) died from unrelated causes. One patient with rectal NET showed local recurrence within a year after operation. One patient operated for rectal melanoma developed distant metastases and died two years after the operation (Table 2). We observed an overall survival equal to 97.6% (95%CI: 95%-99%) at the end of follow up and a disease free survival of 98% (95%CI: 96%-99%) (Figure 1).

**DISCUSSION**

Rectal lesions different from adenomas-carcinomas represent a small but important group in terms of oncological and functional implications. Surgery is the main choice in the treatment of these conditions, but debate regarding the best method for their management exists[1-3]. Their localization in the rectum may represent a therapeutical challenge. Most authors opt for anterior resection or even abdominal perineal resection, but traditional surgery may represent an overtreatment[1,2,4].

NETs represent the largest group in our series. This kind of tumors are being diagnosed increasingly frequently, and current European Neuroendocrine Tumor Society (ENETS) guidelines recommend endoscopic resection for G1 rectal NET < 10 mm with a low risk of metastatic disease[5]. The current methods of endoscopic removal are polypectomy, endoscopic mucosal resection (EMR), endoscopic submucosal dissection (ESD) and TEM. Since complete surgical resection for a localized lesion was demonstrated as the only effective option, several studies have proved the superiority of TEM over the other endoscopic techniques in the treatment of rectal NET. EMR and ESD achieve a complete microscopic resection in 46.3% to 65.5% and in 75% to 82.6% of cases, respectively[6-11]. TEM allows us to achieve a 100% rate of free resection margins, as observed in other reports[9,10]. We did not observe cases of incomplete resection nor recurrence in our experience. Most tumors (80%) were ≤ 10 mm in diameter, and the risk of metastases has been estimated at less than 3% for rectal NETs within 1 cm in diameter[9]. In our series, all the lesions were G1 well-differentiated rectal NET without lymphovascular invasion except for one patient with a G3 poorly differentiated NET with lymphatic and vascular invasion, who relapsed within a year from operation and was treated by means of an abdominal perineal resection.

As for GISTs, according to Miettinen et colleagues, the rectum is the third most common site of onset, comprising approximately 5%-10% of all GISTs[12]. Neither radiation therapy nor chemotherapy has any proven efficacy as adjuvant therapy. Rectal GIST exhibits two specific features which may significantly affect surgical management: metastases are extremely rare in loco-regional lymphnodes, and GISTs typically show a tendency to grow away from the intestinal lumen. These characteristics may make these tumors eligible for TEM[13-16]. In our series, all GISTs were completely resected by TEM. TEM excision is considered to be an interesting alternative for small GISTs located within the rectal wall, which are usually incidental findings during endoscopy. This approach, however, is considered not indicated for larger (> 5 cm) tumors growing away from the rectal lumen. In our series, only one GIST had a 5 cm diameter, but it was completely excised, and we did not observe recurrence.

Condyloma mainly affects the anorectal region, and rare reports have described condylomata involving the rectal wall which have often been incidentally discovered by endoscopy[17-19].

Standard therapy such as laser, fulguration, freezing or microwaves can be difficult to apply inside the rectum[18]. Surgical resection by TEM can offer a good local disease control, and none of the patients treated by TEM experienced recurrence.

Rectum can also be the site of extapelvic endometriosis[2,4,8,20]. Open or laparoscopic surgery is the primary mode of treatment in most of the infiltrating diseases. Surgical treatment is effective in relieving painful defecation, pelvic pain and dyspareunia[20]. We registered a positive resection margin in one patient affected by endometriosis, but no recurrence was observed in this case. Probably, the margin presented to the pathologist as elettrocoagulated. Primary anorectal malignant melanoma is an extremely rare malignancy that is believed to arise from melanocytes in the mucosa around the anorectal junction. Surgery resection is the only curative option, but this malignancy is associated with poor prognosis[21,22]. We treated only two patients with rectal melanoma by TEM who were incidentally diagnosed during endoscopy. Both cases had an early stage of melanoma confirmed by the pathologist. Both patients received adjuvant chemotherapy. One of them developed local recurrence at 1 year from surgery and was treated with laparoscopic anterior resection.

TEM has demonstrated to be feasible in the treatment of different conditions different from adenomas and carcinomas which may affect the rectum. TEM allows us to reach lesions located up to 20 cm from the anal verge. The magnified tridimensional vision offered by TEM is crucial to reach the complete rate of complete resection. The possibility to perform a full thickness excision of the rectal wall makes TEM appropriate for tumors like GISTs arising from submucosal layers.

In this series, we did not experience long term morbidity. We registered only one patient with mild gas incontinence which was resolved within two months from surgery by means of physiotherapy.

We could conclude that TEM is an important therapeutic option for rectal rare diseases. Other studies with more numerous series will be necessary to understand the real role of minimally invasive transanal techniques in the treatment of these lesions.

**COMMENTS**

***Background***

The rectum can be the site of origin of different lesion far more rare than adenocarcinoma but that have surgery as the only curative option. The full thickness excision reached by transanal endoscopic microsurgery (TEM) offers the possibility to achieve a complete resection with very low morbidity.

***Research frontiers***

TEM has several advantages compared with traditional approach. It allows to perform a complete transanal full thickness excision of the lesions, with an accurate individuation of free margins due to a magnified stereoscopic view. The morbidity related to this technique is low and

***Innovations and breakthrough***

The exact role of TEM in the treatment of rare rectal lesions is hard to define manly due to the lack of large series. The retrospective analysis of our experience allowed us to built one of the largest series now available on this topic.

***Applications***

This retrospective analysis of the authors’ experience suggest TEM can be considered safe and feasible in the treatment of these lesions.

***Peer-review***

this is a large retrospective analysis on the treatment of rare rectal lesions by TEM. The paper is overall well written. The results are well reported.

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**P-Reviewer:** McSorley ST, Samardzic S, Santoro GA, Sterpetti AV

**S-Editor:** Gong ZM **L-Editor:** **E-Editor:**



**Figure 1 Disease free survival after transanal endoscopic microsurgery for rare rectal lesions.**

**Table 1 Population characteristics** **n (%)**

|  |  |
| --- | --- |
| **Variables** |  |
| Sex |  |
| Male | 30 (67.7) |
| Female | 22 (42.3) |
| NETs | 21 (40.4) |
| GISTs | 14 (26.9) |
| Ganglioneuroma | 1 (1.9) |
| Solitary ulcers | 2 (3.8) |
| Endometriosiosis | 6 (11.5) |
| Condylomas | 6 (11.5) |
| Melanomas | 2 (3.8) |
| Diameter (cm), [mean(median, range)] | 2.7 (4, 0.4-8) |

**Table 2 Operative and post-operative data n (%)**

|  |  |
| --- | --- |
| **Variables** |  |
| Operative time-min [mean (median, range)] | 41(45, 20-55) |
| Intraoperative complications | 1 (1.9) |
| Hospital stay (d) [mean (median, range)] | 3 (4, 2-7) |
| Post-operative complications | 2 (3.8) |
| Recurrence | 1 (1.9) |
| Follow up (yr) [mean (median, range)] | 11 (13, 23-1) |
| Death at the end of follow up | 2 (3.8) |