Transanal approach for rectal tumors: recent updates and future perspectives

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Abstract: Local excision of rectal tumors has long been performed. With the development of new technologies for endoluminal operation, there have been significant advances in the transanal approach in the last 30 years. It started with conventional transanal excision (TAE) and was developed by introduction of transanal total mesorectal excision (taTME). Appropriate patient selection for transanal approaches is the key for good outcomes. TAE of rectal tumors has been advocated for premalignant lesions and used as definitive treatment for early rectal cancers in select groups without adverse prognostic features. Regardless of procedures, the morbidity and mortality of transanal technique are lower than for radical surgery. According to the experience in the last decades, transanal endoscopic surgery has been accepted as effective treatments in selected patients with early rectal cancer, with similar oncologic outcomes to and better functional effects than those of radical surgery. The latest development in transanal approaches is taTME. Although taTME is another new technique with great promise, the supporting data are preliminary, and further studies with larger cohorts of patients are needed to evaluate long term functional and oncological outcomes. Transanal approaches have enabled mid and upper rectal lesion and sphincter salvage, leading to a better quality of life. Transanal approaches including taTME should be considered as good options for the treatment of rectal cancer because these techniques are definitely useful in selected patients.

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Introduction

Local excision of rectal tumors has long been performed. The transspincteric and transcoccygeal approaches had been used for local excision, especially for high-lying rectal tumors. The transcoccygeal (Kraske) approach requires mobilization of posterior pelvic floor muscles away from the coccyx to expose the rectum and the transspinhincteric (York-Mason) approach involves complete division of the anal sphincter. With the development of new technologies for endoluminal operation, the transspinhincteric and transcoccygeal approaches are rarely used today. Transanal excision (TAE) was first described by Parks as an alternative endoluminal treatment for certain rectal tumors in 1970 (1). After 10 years, anorectal surgical procedures with the use of different endoscopic devices into the anal canal were introduced. Transanal endoscopic microsurgery (TEM) was first described by Buess et al. in 1984 (2). A few years later, a newer and simpler alternative transanal endoscopic operation (TEO), was introduced and widely implemented. Maeda et al. (3,4) proposed a new transanal local excision procedure, minimally invasive transanal surgery (MITAS), for excising a proximal tumor at more distal sites. Recently, transanal minimally invasive surgery (TAMIS) has become increasingly more popular.
Selecting the appropriate surgical approach to rectal tumors is important. The approach must balance successful tumor eradication with functional implications for the patient. Owing to the significant morbidity and alterations in quality of life associated with rectal surgery (low anterior resection and abdominoperineal resection), a lot of time and research have been devoted to transanal approaches for rectal tumors. Several retrospective studies since the 1970s reported that TAE of early tumors with negative margins may provide similar outcomes those of radical resection (5). Since then, there had been many studies assessing the role of TAE of rectal cancer (6). Thereafter, the use of TAE had increased to 17.1% for T1 lesions and 11% for T2 lesions from 1989 to 2003 (7).

During the recent decades, total mesorectal excision (TME) has become the standard technique for the surgical treatment of rectal cancer (8). Nowadays, transanal TME (taTME) has been proposed as a new option in cases for which laparoscopic transabdominal TME (laTME) is difficult. TaTME is not a completely novel concept and it has benefited from previous experience of transabdominal-transanal (TATA) operations, TEM, TAMIS and natural orifice transluminal endoscopic surgery (NOTES) (9-11). Since the first taTME resection assisted with laparoscopy was reported in 2010 (12), taTME has shown promising results with regard to pathological quality and short- and mid-term outcomes (13-15).

**Indication for TAE to rectal tumors**

Local excision of rectal tumors has been advocated for premalignant lesions and used as definitive treatment for early rectal cancers in select groups without adverse prognostic features (16). Atypical rectal tumors such as neuroendocrine tumors and gastrointestinal stromal tumors are also usually approached through the transanal.

Local excision is suitable for Tis (carcinoma in situ) or T1 cancers with a favorable histology. The criteria for local treatment include T0 or Tis lesions; low-risk differentiated (well-to-moderately) T1 cancer; absence of lymphatic, vascular, or perineural invasion; and tumors ≤3 cm in diameter occupying ≤40% of the circumference of the rectal lumen. These principles apply to all local excision techniques. However, with the development of the technique and increased experience of surgeons, the indications have expanded. These technical approaches have extended beyond local excision, with the development of taTME being the most important in recent years. Additionally, for accurate patient selection, routine preoperative cardiopulmonary assessment, physical examination with digital rectal examination, fecal incontinence test, endoscopy, trans-rectal ultrasound (TRUS), and in case of malignancy; complete staging workup with pelvic magnetic resonance imaging and abdominal computed tomography, are recommended. For determining local excision, local T staging should be accurate. TRUS allows predicting early T1 lesions that might be suitable for local excision. Hildebrandt et al. (17) proposed a preoperative tumor staging system based on ultrasonic determination of the infiltrative depth of tumors, so-called uTNM, and this technique contribute to a more accurate determination of the depth of invasion with classification of the rectal wall layer.

**Techniques for the transanal approach**

- Conventional TAE.
- MITAS.
- Transanal endoscopic surgery (TES).
  - TEM;
  - TEO;
  - TAMIS;
- taTME.

**TAE**

TAE has been the mainstay of treatment for many years. TAE is a simple method that can be easily performed if the tumor is located in the anal canal and easily accessible under adequate exposure of the anal canal. There is no need for additional equipment, and it can be performed in the outpatient department. An additional benefit to this approach is the minimal, if any, compromise of anorectal and urogenital function. These factors make TAE the most common method of local excision (18).

Conventional TAE is often limited to tumors ≤4 cm in diameter that lie within 6–8 cm of the anal verge (16). Lesions in the middle and upper rectum are usually inaccessible with TAE because of their distance from the anal verge, and attempted excisions are hampered by inadequate surgical exposure, confinement of the operating field, and uncertainty of a clear surgical resection margin (19).

Patients should undergo preoperative assessment including digital rectal examination and rigid sigmoidoscopy to confirm location and mobility. Patients receive a cleansing enema the day before the operation; prophylaxis with
antibiotics and antithrombotics is usually recommended. After the induction of local, regional, or general anesthesia, the patient is placed in position. Some authors additionally propose pudendal nerve block for sphincter relaxation. The positioning of the patient is dependent on the preference of the surgeon; however, the orientation of the lesion is usually the deciding factor, with preference taken to operating downward. Most operations are performed with the patient in the prone jackknife position; however, some posteriorly located lesions may be better approached with the lithotomy position. The perianal area is exposed by taping the buttocks apart; the lesion is exposed with direct vision by using a Hill Ferguson, Park, or Barr retractor. Traction sutures can be placed distal to the lesion to improve visualization. The calculated excision margin (10 mm) is typically marked by using electrocautery in a circumferential pattern around the lesion. The specimen must be carefully taken so as not to manipulate the lesion or handle it with instruments. After a full thickness excision of the lesion, the specimen is oriented on a needle board and sent to the pathology laboratory. After irrigation, the defect can be either left open or closed transversely with absorbable sutures. If the lesion is posterior to the rectum and above the puborectalis muscle, the defect is closed with absorbable sutures. In case of anteriorly located lesions, repair should be conducted to avoid injury of adjacent structures, such as the prostate, urethra, or vagina. At the end of the procedure, a proctoscopic examination is essential to confirm that the rectal lumen was not inadvertently closed or narrowed.

Complications associated with TAE include urinary retention, urinary tract infection, infections of the perirectal and ischiorectal space, fecal impactions, and delayed hemorrhage. Nevertheless, the incidences of these complications and mortality are very low. TAE has several limitations. In general, this approach is technically difficult with higher lesions owing to poor visualization. More important, visualization during TAE can be suboptimal, which can affect the quality of an oncologic resection margin. Concerns have also been raised about the high rates of tumor fragmentation and recurrence with TAE. The high rates of recurrence are likely due to rates of margin positivity that exceed 10% in even the most experienced and expert case series (6,18). Patients treated with TAE compared with those treated with radical surgery for T1 rectal cancer had a significantly higher 5-year local recurrence rate (12% vs. 6%) and lower 5-year survival rate (70% vs. 80%) and 5-year disease-free rate (64% vs. 77%) (20).

**MITAS**

A new transanal local excision procedure, MITAS, has been developed with a specially designed anal retractor connected to the Octopus retractor holder, a staple device, and several newly developed techniques to excise a proximal tumor at a more distal site (21).

Operation was usually performed under spinal anesthesia with the patient in lithotomy or jackknife position, according to the site of tumor (4). The procedure consists of inserting into the rectum an originally designed E- or F-type anal retractor [a modified K-type anal retractor (22); Yufu Itonaga Co Ltd.] connected to the Octopus retractor holder, long type, 22 inches (Mednosbro AG).

To enable excision of a proximal tumor at a more accessible site, shortening or roll-in technique, intussusception, or invagination technique with retraction stitches is used. The retractor is inserted into the anus and rolled to pull the rectum in and permit easy access to the tumor in the proximal rectum (3). When the tumor is still beyond the surgical field, the retractor is opened and fixed and two Babcock forceps are used to pull the tumor gradually (21). Retraction stitches are passed under the tumor, from one side to the other with a minimum macroscopic margin of 5 mm from the boundary of the tumor to the adjacent normal mucosa by a 36-mm-long atrumatic needle with absorbable 1-0 thread, enough to retract the tumor fully and pull the rectum down. ENDO GIA (Tyco Co Ltd.) is used for excision and anastomosis while fully retracting the rectum with retraction stitches distally. Application of a stapler is usually done transversely, but oblique application is sometimes needed because of the difficult angle in the rectum or size of the tumor.

**TES**

**TEM**

Gerhard Buess of Germany pioneered TEM (Richard Wolf, Germany) in the early 1980s as a minimally invasive technique allowing the resection of adenomas and early rectal carcinomas unsuitable for local or colonoscopy excision, which would otherwise require major surgery (23). This method was basically designed from the idea of laparoscopic surgical techniques so-called because it is a minimally invasive technique and has proved to be useful for treating lesions in the mid or upper rectum. The main indications for TEM are rectal tumors that are out of reach for TAE and are unsuitable for endoscopic removal (24).
In addition, this technique can even be extended to lesions in the low sigmoid colon, with success reported up to 20 cm from the anal verge (25). Since its development, TEM has been also used for a variety of other rectal pathologies including neuroendocrine tumors, rectal prolapse, early stage carcinomas, and palliative resection of rectal cancers (26).

The orientation of the lesion is usually the deciding factor for the positioning. The lesion needs to be situated at the 180 degree angle of the scope view. For a posterior lesion, the patient is placed in a lithotomy position. For an anterior lesion, the patient is placed in the prone jackknife position. The operative technique for TEM involves three main components: a rigid operating rectoscope, a laparoscopic camera, and modified laparoscopic instruments. The operating rectoscope is typically 4 cm in diameter and varies from 12 to 20 cm in length. The rectoscope maintains an airtight seal at the anus once inserted in the rectum, and is held in place by the obligate articulating arm, which fixes the rectoscope to the operating table. The rectoscope has a port for the inflow of CO₂ for the pneumorectum, and an outflow for smoke evacuation during cauterization. The faceplate on the rectoscope has four ports through which a stereotactic telescope with connection to a three-dimensional video system and three modified laparoscopic instruments are connected to facilitate dissection and suturing (26). While the instruments are specialized, the operative steps are otherwise no different than for TAE. The lesion is then centered in the scope view. By using scissors, cautery, and graspers, a dotted line is burned around the target lesion with a 10-mm margin. Care must be taken to avoid trauma to and fragmentation of the specimen. After homeostasis is obtained, the defect is closed with absorbable full-thickness sutures in a transverse line. The defect can be left open to heal secondarily if the defect is posterior and surrounded by mesorectal fat. An anterior defect must be closed; closure is facilitated by placing a central stay suture and sewing from each corner to avoid tension on the suture line.

The limitation of TEM is that the equipment is designed to operate from the top-down; thus, the lesion must be oriented toward the floor to be compatible with the equipment. This means that the patient’s positioning is dependent on the tumor location, and sometimes specialized split-leg operating tables are necessary to resect anterior tumors requiring the prone jackknife position. Distal lesions near the sphincter are difficult to excise with TEM owing to the configuration of the equipment and inability to maintain insufflations of CO₂ to distend the rectum. This is the reason why TAE is easier to use for low-lying lesions. Other limitations to TEM focus on specialized equipment, which have a steep learning curve and high associated costs for the hospital (27,28).

TEO
A few years later a cheaper alternative was introduced compared with TEM. The TEO (Karl Storz, Germany), which was a newer and simpler system, has become widely implemented. Indication of TEO is similar to TEM.

TEO platform was performed in a lithotomy position using a 30° forward-oblique telescope, adequate adjustment of the rectoscope, and curved laparoscopic operating instruments. TEO procedure was carried out under general anesthesia. After installation of the holding system, the anus was gently dilated, and the operating rectoscope (7.5 or 15 cm long, 4 cm in diameter) with obturator was inserted in the rectum with copious lubricant and fastened to the support arm attached to the operating table. The working attachment used with the rectoscope had two channels for instruments of 5 mm and one channel for instruments up to 12 mm. After achieving the pneumorectum with insufflations of CO₂ to 12 mmHg or more, a high definition (HD) 5-mm diameter endoscope with fiber-optic light transmission and 30° angled view was inserted through a 5-mm endoscope channel and the rectoscope was adjusted to achieve the best position for procedures (Figure 1A). Except the instruments are specialized, the operative steps are otherwise no different than those for TEM.

Similar to TEM, lesions near the anal verge are difficult to excise with the TEO. Standard laparoscopic instruments, equipment, and set up costs are lower, potentially opening the technique to any surgeon with previous laparoscopic experience. Hur et al. (29) performed initial experience of TEO and reported the mean operative time was 85 minutes, and the mean postoperative hospital stay was 4.5 days, a positive resection margin was documented for 9% patients. Furthermore, they demonstrated that according to the cumulative sum analysis, the operation time and hospital stay significantly decreased after 17 case experiences (29). Several studies have compared TEO with TEM for benign and malignant lesions and have shown satisfactory outcomes (30,31).

TAMIS
In recent years, TAMIS has become increasingly more popular. Reported by Atallah et al. (32) in 2010, the
The indications of TAMIS are the same as those of other TES. The main benefits of TAMIS are the relative ease of use and low cost, owing to the use of conventional laparoscopic instruments, including a laparoscopic camera, graspers, energy sources, and a standard laparoscopic CO$_2$ insufflator. It is ideal for lesions at 8–12 cm from the anal verge; however, it has been successfully performed in the lower and mid rectum (34). Distal lesions are covered by the transanal port, and excision of very proximal lesions can cause entry into the peritoneal cavity.

Mechanical bowel preparation or enema, antibiotics and antithrombotic prophylaxis are usually recommended. Anesthesia may be general or spinal. Lee et al. (35) have reported a series of 25 TAMIS procedures in which spinal anesthesia was used. In TAMIS, most lesions can be excised with a lithotomy position. However, we still recommend the prone position for the patients with large anterior lesions, especially if the distance from the anal verge is in a range in which there might be a risk of perforating the peritoneum. After the patient is positioned and the port is placed, pneumorectum is achieved with the standard CO$_2$ insufflator, with pressures ranging from 15 to 25 mmHg. Several different cameras can be employed, including those with a flexible tip. One common practice is to use the 5-mm, 30-degree bariatric camera with a right-angle light cord.

Figure 1 Procedure and position for TEO (A), single-incision assisted laparoscopic surgery port (B), GelPOINT path transanal access platform (C), glove port for robotic TAMIS (D). TEO, transanal endoscopic operation; TAMIS, transanal minimally invasive surgery.
adaptor. Conventional graspers, scissors, and electrocautery devices are employed, along with ultrasonic or bipolar energy devices as needed. Owing to the technical challenges of suturing in this confined working space, many techniques similar to TEM have been employed, including the use of clips and beads, barbed suture, and specialized suturing devices.

When compared with other platforms, TAMIS has several advantages. Devices that are used for TAMIS are more pliable than the 40-mm rigid scope used for TEM, and possibly lead to less impairment of sphincter function; the set-up time is significantly lower for TAMIS. Use of regular conventional laparoscopic instruments, as opposed to the fixed eyepiece of the TEM rectoscope, enables advancing the scope into the proximal rectum to look beyond the tumor. TAMIS is easily learned by surgeons because of its simplicity and similarity with conventional laparoscopic surgery, and it is a cost effective alternative to TEM (32). For some authors, the introduction of the TAMIS port into the anal canal makes it more complex than TEM or TEO (36). A disadvantage of TAMIS is that the rectoscope cannot be mobilized at the site of the lesion; rectal lesions behind a rectal haustral valve may be more difficult to access and remove. The longer channels associated with TEM and TEO equipment facilitates intraluminal rectal retraction. Moreover, an assistant is required to hold and manipulate the laparoscope during the TAMIS procedure.

The authors who introduced TAMIS went on to describe the use of a robotic platform for TAMIS in a cadaveric model in 2011 (37), and then extended that robotic TAMIS platform to live patients in 2012 (38). Since then, its use in humans has been described with both the GelPOINT Path platform (Applied Medical, USA) and a glove port (Figure 1D) (39).

The authors suggested that the transanal glove port facilitated the robotic setup, enabling flexibility and allowing docking of the cannulas away from the limited perianal workspace (40). Furthermore, the glove port provided a wider axis of movement for instruments inside the rectum, or allowed them to be easily rotated and/or crossed. Although robotic TAMIS has been shown to be feasible, this technique is still relatively new, and more studies are necessary before to widespread adoption.

**Postoperative complication of TES**

Regardless of procedures, the morbidity and mortality are lower than for radical surgery. Operative mortality is less than 0.5% and morbidity ranges from 4% to 30% in large series, depending on the inclusion of minor complications. The most frequent complications include acute urinary retention, bleeding requiring reoperation, abdominal perforation and recto-vaginal fistula (41). Kumar et al. (42) found that complications correlate with tumors located laterally and more than 8 cm from the anal verge. Pelvic sepsis, which occurs in about 3% of cases, is more common in lesions within 2 cm of the dentate line. The conversion rate to TAE is around 5%, and the main reason for conversion is technical difficulties (24,26). Peritoneal perforation, which was thought to represent a complication requiring conversion to laparotomy, can usually be salvaged with TES for experienced surgeons (43,44). A multicenter study by Baatrup et al. (45), performed by using database of 888 TEM procedures, found 22 perforations in the peritoneal cavity. They reported no association with major short term complications or adverse long-term oncological outcomes (45).

**Outcomes of TES**

Radical surgery with TME is still the treatment of choice for rectal cancer, offering patients the best results in terms of local recurrence and survival (46). However, TME is associated with significant mortality and morbidity (47). According to the experience in the last decades, TEM and TEO have been accepted as effective treatments in selected patients with early rectal cancer, with similar oncologic outcomes to and better functional effects than those of radical surgery (48,49). Recently, TAMIS has been proposed as an alternative technique; however, the experience with this approach for rectal cancer is still limited because of short follow-up (27,33,50-55).

TEM has a lower positive resection margin rate than TAE, with less fragmented specimens and better oncologic outcomes (56,57). Elmessiry et al. (50) showed that TAE was an independent predictor of local recurrence compared with TEM. The rate of reported positive resection margin in the surgical specimen in TAMIS was 4.4–6% (27,33,54), similar to those obtained with TEM, and seems to be related with the T stage (26,58,59). Some studies have compared TEM with radical surgery in early rectal cancer, showing similar results in terms of local recurrence and survival (60,61). In a meta-analysis, Winde et al. (62) demonstrated that the rate of local recurrence was higher with TEM (12% vs. 0.5%); however, no difference in survival was found.

TES seems to be a reasonable alternative to radical surgery in patients with low-risk rectal cancer (26,41,47,
principles may be somewhat cumbersome in cases of narrow dissection of the distal rectum according to TME, even in the most challenging cases (11). However, even in these studies there are considerable differences between low- and high-risk cancers (73). TES may be performed on patients with high-risk T1 or T2–3 tumors with poor life expectancy and severe morbidity, or those unfit for major surgery, or simply as palliative treatment in case of disseminated disease (47,61,63).

Salvage surgery for recurrence after TES demonstrated disappointing oncologic outcomes; the stage is usually more advanced than in primary lesions and may require multivisceral resection and an ostomy in up to 43% of cases. Survival is seriously compromised, with a 5-year survival ranging from 43% to 68%, dropping to 29% in patients with unfavorable histology (63,68,74). In contrast, Levc et al. did not find any difference in outcome between patients with rectal cancer undergoing immediate salvage TME after TEM and those undergoing primary TME (75). Despite contrasting conclusions, all authors warned that perforations into the original operating field during subsequent TME can occur owing to fibrotic changes to the bowel wall, which might allow microscopic tumor spillage.

**Future perspectives**

Several new techniques and approaches are still under investigation, currently in preclinical or experimental stages, such as transanal natural orifice transluminal endoscopic surgery (NOTES), taTME, and robotic-TES (26,51,76). TES platforms seem to be safe for both transanal NOTES and taTME (77,78). Robotic technology can lower the difficulty inherent in TES platforms (79). However, clinical trials are necessary for full evaluation of these techniques.

**taTME**

The latest development in transanal approaches is taTME. Transanal TME was introduced in 2010 with the aim to cope with all these limits and improve the quality of mesorectal dissection even in the most challenging cases (11). Dissection of the distal rectum according to TME principles may be somewhat cumbersome in cases of narrow pelvis, bulging tumors, and obese patients. In adequate exposure and loss of the good plane of dissection require finding another alternative treatment approach. During the last decade, transanal approaches have been extensively used to overcome the inherent shortcomings of laTME (69,77). Among these emerging transanal techniques, taTME is a new minimally invasive procedure with the essential aim of improving oncological treatment quality and avoiding pelvic nerve injury in patients with mid or low rectal cancer. Given the encouraging outcomes of systematic investigation of taTME for patients with rectal cancer (80,81), taTME may be optimized as a surgical approach for rectal cancer. In comparison with conventional laTME, taTME defines the distal resection margin more precisely, with better visualization of the distal rectum, and allows the surgeon to perform deep pelvic dissection without the need for difficult retraction (82). However, the benefits of taTME compared with laTME must be confirmed before conducting multicenter randomized controlled trials (RCTs) and unifying taTME procedures. According to a meta-analysis study, the percentage of patients with complete mesorectum was 83.4% in the taTME group and 73.4% in the laTME group (83). In addition, achievement of complete plus near complete mesorectum was also greater in the taTME group (95.3% vs. 88.2%) (83). Hence, for patients with mid or low rectal cancer, taTME may achieve a complete or near complete resection of the mesorectum relative easily, compared with laTME. However, whether a higher quality of mesorectal resection will result in good survival remains unknown. Safety is always the most important issue for a new technique. The meta-analysis indicated a comparable rate of intraoperative complications and a significantly lower incidence of postoperative complications in the taTME group than in the laTME group (83).

As the new surgical technique of taTME is adopted increasingly by surgeons, the patient selection criteria will be crucial and will continue to inspire debate. Of note, the protocol published recently for a multicenter RCT comparing taTME with laTME (COLOR III) has formulated strict criteria for patient selection (84). According to the selection criteria of this protocol, T3 tumors with margins <1 mm to the endopelvic fascia, tumors with ingrowth in the internal sphincter or levator ani muscle, and all T4 tumors staged before preoperative therapy were excluded (84). However, the nature of the surgical candidates best suited to taTME treatment requires further studies.

In a matched case-control study from Taiwan, Chen...
et al. (85) demonstrated that compared with laTME, taTME not only achieves identical circumferential margin status without compromising other operative and quality parameters but also benefits patients by achieving a longer distal margin. Additionally, Denost et al. (86) performed a randomized trial in France, and reported that the rate of positive circumferential resection margin decreased significantly after taTME compared with abdominal low rectal dissection (4% vs. 18%; \(P=0.025\)). Currently, RCTs examining taTME are under way; the COLOR III study has been designed to compare taTME versus laTME for mid and low rectal cancer. TaTME is expected to be superior to laTME in terms of clear circumferential resection margins in case of mid and low rectal carcinomas (84). Although taTME is another new technique with great promise, the supporting data are preliminary, and further studies with larger cohorts of patients are needed to evaluate long term functional and oncological outcomes.

Conclusions

There have been significant advances in the transanal approach in the last 30 years. Appropriate patient selection is the key for good outcomes. These techniques have enabled mid and upper rectal lesion and sphincter salvage, leading to a better quality of life. From the point of view of technical advancement, it would be better to adopt this technique as a treatment option, and prospectively randomized comparison clinical trials should be conducted. When we are planning treatment for patients with early rectal cancer, the risk of local recurrence must be balanced with the quality of life. Nowadays, transanal approaches including taTME should be considered as good options for the treatment of rectal cancer because these techniques are definitely useful in selected patients.

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Footnote

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