



doi: 10.1093/gastro/gow023

Advance Access Publication Date: 31 July 2016

REVIEW

Pelvic autonomic nerve preservation in radical rectal cancer surgery: changes in the past 3 decades

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Abstract

The advent of total mesorectal excision (TME) together with minimally invasive techniques such as laparoscopic colorectal surgery and robotic surgery has improved surgical results. However, the incidence of bladder and sexual dysfunction remains high. This may be particularly distressing for the patient and troublesome to manage for the surgeon when it does occur. The increased use of neoadjuvant and adjuvant radiotherapy is also associated with poorer functional outcomes. In this review, we evaluate current understanding of the anatomy of pelvic nerves which are divided into the areas of the inferior mesenteric artery pedicle, the lateral pelvic wall and dissection around the urogenital organs. Surgical techniques in these areas are discussed. We also discuss the results in functional outcomes of the various techniques including open, laparoscopic and robotic over the last 30 years.

Key words: total mesorectal excision; pelvic autonomic nerve preservation; urinary dysfunction; sexual dysfunction

Introduction

Surgical management for rectal cancer is challenging due to the narrow pelvis and extreme proximity to contiguous organs; hence, recurrence rates are commonly reported. The advent of total mesorectal excision (TME) [1,2] together with minimally invasive techniques such as laparoscopic colorectal surgery [3–6] and robotic surgery [7–9] have not only improved surgical results but have also improved surgical technique, operative ability and surgical visibility. Nonetheless, the improved survival of patients due surgeons' increased ability to resect rectal cancers completely is not without problems. In particular, bladder and sexual dysfunction may be particularly distressing for the patient and troublesome to manage for the surgeon when they do occur.

The incidence of urinary dysfunction may be as high as 27% and includes difficulty emptying the bladder as well as urinary incontinence. Sexual dysfunction may also reach 11–55% after

TME. For females, the inability to achieve orgasm, dyspareunia and reduction in vaginal lubrication may be distressing, even for some of the more elderly females. Many male surgeons do not realise that this is an important quality of life factor, especially for younger female patients. For males, nerve dysfunction may include erectile dysfunction, absence of ejaculation or retrograde ejaculation [10–13]. The increased use of neoadjuvant and adjuvant radiotherapy is associated with poorer functional outcomes. When radiotherapy is indicated, however, it cannot be withheld just because of the fear of nerve dysfunction. One of the main risk factors, poor surgical technique with resultant iatrogenic sexual and urinary dysfunction, however, may be prevented by thorough and practical understanding of pelvic nerve anatomy [14].

This review attempts to evaluate current understanding of the anatomy of pelvic nerves and the differences in outcomes of the various techniques including open, laparoscopic and robotic.

Submitted: 28 June 2016; Revised: 18 July 2016; Accepted: 28 June 2016

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Surgical Pelvic Nerve Anatomy

Sexual and urinary dysfunction may be due to somatic and autonomic pelvic nerve damage. The lower pelvis can be divided into 2 anatomical levels: one above the levator ani (supplied by autonomic nerves) and the other below the levator (supplied by somatic innervation via the pudendal nerves) [15,16]. The autonomic plexus includes the superior hypogastric plexus (SHP) comprising sympathetic nerves, the inferior hypogastric plexus (IHP) comprising mixed sympathetic and parasympathetic nerves and the pelvic splanchnic nerves (which are parasympathetic nerves) [14,15].

The principles of sharp dissection and direct visualization of all structures during surgery of the rectum apply in all cases regardless of whether open, laparoscopic or robotic techniques are applied. The well-known TME technique requires high ligation of the inferior mesenteric artery pedicle and careful sharp dissection in the 'holy plane' down to the pelvic floor with transection of the rectum (with adequate margins) followed by endto-end anastomosis. Limitations of direct visualization may result from the anatomical constraints of the narrow bony pelvis, especially with a very curved, prominent sacrum together with a narrow true pelvis which is often especially narrow in the android pelvis. This may be made worse in patients with high body mass index with bulky pelvic sidewall and mesorectal fat. Pelvic adhesions from radiotherapy or tumour inflammation or gross tumour adherence to contiguous organs may also require surgeons to perform extrafascial dissection to achieve good circumferential radial margins or clearance—and thereby increase the risk of damage. The possible need for lateral node dissection may also lead to further injury.

Inferior mesenteric artery (IMA) pedicle

Nerve anatomy

The SHP is located anterior to the body of the L5 vertebra and is found on the left anterolateral side of the aorta and its bifurcation [15]. The SHP arises from pre-aortic sympathetic trunks alongside the T10-L3 vertebrae, descends along the sacral promontory and then bifurcates into the bilateral hypogastric nerves [17]. These run 2 cm medial to the ureter and common iliac artery on both sides [13] (Figure 1). The hypogastric nerves move obliquely and anteriorly towards the rectum and then along the side of the rectal fascia where they are covered by the endopelvic fascia. These hypogastric nerves finally end as afferent fibres of IHP at the level of the intersection between the vas deferens and the ureter [15,16,18].

Surgical technique

Ligation of the IMA begins with tenting the sigmoid colon. The promontory and aortic bifurcation are identified, and the surgeon should be aware that the peritoneal layer covering this area holds the hypogastric nerves (Figure 2). In a medial-to-lateral approach, the incision is made medial to the right iliac vessels and may extend up to the duodenojejuenal flexure. Care is taken to push back the sympathetic pre-aortic nerves together with the retroperitoneal structures, such as the left ureter and left gonadal vessels located just below the peritoneum, using sharp dissection to expose the IMA origin. Technical points:

- i. Ligation of the IMA should be performed 1.5-2 cm from its aortic origin to avoid damage to SHP fibres lying in front of the aorta [15,18].
- ii. Avoid mass clamping of the IMA which may increase damage to the left trunk of the SHP due to its closer proximity

- compared with the right trunk of SHP located in the aortocaval plane [19].
- iii. Preserve Gerota's fascia during mobilisation of the ureter and gonadal vessels as these contain SHP fibres [20].
- iv. At the level of the sacral promontory, the transition from mesosigmoid to mesorectum is an area where damage is possible to the presacral plexus and hypogastric nerves. To avoid the wrong plane of dissection, it is important to dissect only immediately posterior to the superior rectal artery and to remain within the plane anterior to the parietal presacral fascia. It has been suggested that this should be approximately 2 cm anterior to the promontory [20,21].

Lateral pelvic wall

Nerve anatomy

The IHP receives pelvic parasympathetic fibres from roots S2-S5 (splanchnic nerves or erector nerves of Eckard). These nerves are covered by the parietal fascia, pierce the endopelvic fascia, cross the retrorectal space and form branches into the rectum via the lateral ligaments [15]. Fibres from the IHP also innervate the seminal vesicles, prostate and bladder, cervix and vagina. These nerves are responsible for penile erection, detrusor contractility, female arousal and vaginal lubrication. The pelvic parasympathetic nerves join the sympathetic hypogastric nerves in a Y-shaped connection to form the pelvic plexus (Figure 3).

The lateral ligaments are thought to be a condensation of endopelvic connective tissue located on the anterolateral side of the subperitoneal rectum. Some authors suggest that 'lateral ligaments' do not exist and are not distinct anatomical structures; instead, these ligaments are perhaps an artefact produced by surgical dissection. These ligaments may or may not contain an insignificant or small middle rectal artery [15,16,22]. Other authors suggest that the lateral ligaments join the parietal fascia to the fascia propia and that part of the IHP lies within the lateral ligaments, giving branches to the rectum. These arise bilaterally about 2cm below the peritoneal reflection [15,23]. Others, however, suggest that all lymphatics are located within the mesorectum and are arranged around the superior rectal artery with no lymphatic drainage into the lateral ligaments [24].

Surgical technique

The opinion that anatomic dissection of the rectum does not require over-zealous lateral dissection arose from this latter understanding. Dissection of the lateral ligaments is best done after posterior dissection has gone as far down to the pelvic floor as possible. [16] With adequate traction and countertraction, the hypogastric nerves can be visualised as they enter the deep layers of the parietal fascia to the IHP. It is suggested that the adherent nerves can be eased off the mesorectal fascia laterally. The surgeon should avoid hooking the tissue laterally with the finger; neither should there be a need to cross-clamp the middle rectal pedicle to avoid damaging the nerves [25,26].

Dissection around urogenital organs

Nerve anatomy

The pelvic plexus is a network of nerves, located at the level of the lower third of the rectum, that innervate the rectum, bladder, seminal vesicles, prostate, ureters, membranous urethra, corpora cavernosa, uterus and vagina [15,21]. In men, neurovascular bundles (of Walsh) from the pelvic plexus include the

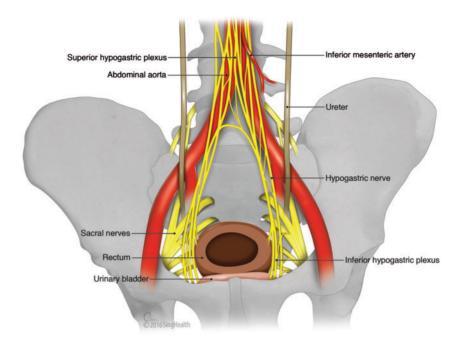


Figure 1: General overview of anatomy of the autonomic nerve distribution. The superior hypogastric plexus around the inferior mesenteric artery descends to the sacral promnotry and bifurcates into hypogastric nerves. These usually run 1-2 cm medial to the ureters and cross the common iliac arteries and S1 in the sacrum.



Figure 2: Inferior hypogastric nerve with branches to the rectum on a robotic view with medial-to-lateral dissection approach.

cavernous nerves which are responsible for erectile function and run laterally outside Denonvilliers fascia, lie at the lateral corners of the seminal vesicles at the 2 and 10 o'clock positions and eventually lie anterior to the postero-lateral border of the prostate and continue onto the periprostatic plexus [25]. In women, the nerves enter the vesicovaginal and rectovaginal septa and run underneath the crossing point of the ureter and uterine artery [21].

Surgical technique

Dissection of the infra-peritoneal rectum from the prostate, seminal vesicles and vagina which are areas at high risk of nerve injury [15,21]. Heald et al in their original technique considered that the Denonvilliers fascia should be resected in TME surgery as it forms the anterior surface of the mesorectum [2]. It is however suggested that this extra-mesorectal resection should only be performed if there are concerns about a

compromised circumferential resection margin, especially for anterior-based tumours. In such a case, care during dissection should be taken along all dissection planes within the propria fascia of the rectum. Once below the rectoprostatic fascia, the correct plane is along the muscular wall of the rectum [15,26,27]. For posterior and lateral rectal tumours, it is suggested that the entire rectoprostatic fascia be left intact [28] (Figure 4). There are, however, additional difficulties in identifying Denonvilliers fascia in elderly patients following neoadjuvant radiotherapy or in those patients with a narrow bony pelvis or very large bulky tumours.

During the perineal phase of an abdominoperineal resection, care should be taken especially at the level of the prostate which contains the dorsal nerve of the penis and plays an important role in erection and ejaculation [13,29]. This is commonly damaged during excessive traction or excessive or prolonged cautery in the anterolateral plane of dissection.

Comparison of Functional Outcomes with Regards to Open, Laparoscopic and **Robotic TME**

The introduction of TME was important not only for reducing local recurrence but also for preserving urogenital function [2]. The emphasis on sharp pelvic dissection in anatomical planes of the pelvis must be obeyed regardless if the technique is open, laparoscopic or robotic TME. However, postoperative sexual and urinary dysfunction occurs due to inadvertent avulsion or direct injury of nerve plexus.

Laparoscopic rectal surgery is technically difficult, and advanced laparoscopic surgical skills are required. As technology has improved over the years and high-definition cameras replaced traditional optic lenses and reduced 'smoky' images due to use of energy devices, the advances in dissection have produced convincing evidence of short-term benefits of reduced pain, shorter hospital stays and earlier returns to normal work

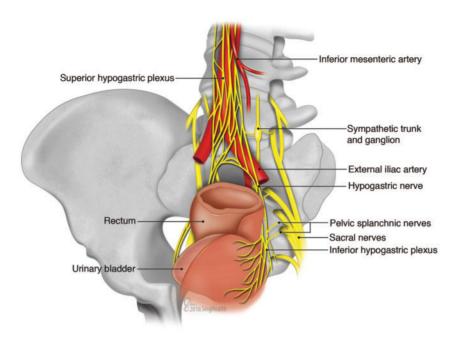


Figure 3: Anatomy of the pelvic autonomic nerves with relation to rectum. The inferior hypogastric plexus comprises nerves from the hypogastric and pelvic splanchnic nerves at lateral pelvic wall.

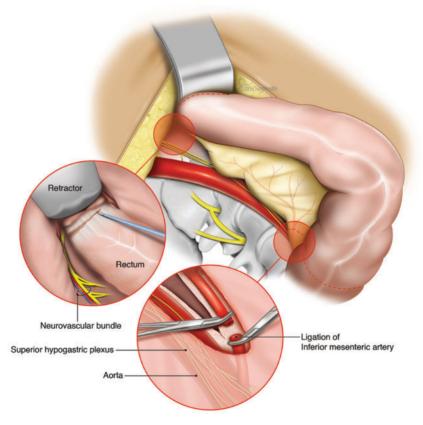


Figure 4: The relationship of the rectum and pelvic autonomic nerves during open surgery when standing on the patient's left. The ligation of the inferior mesenteric artery should be performed 1.5-2 cm from its origin from the aorta to avoid damaging the superior hypogastric plexus. At the pelvis, for posterior and lateral tumours, dissection should be directed below the Denonvillliers fascia to avoid damaging the neurovascular bundles that run along the tip of the seminal vesicle (2 and 10 o'clock directions).

without compromising oncology outcomes. Energy devices, however, produce more heat laterally in contrast to electrocautery, and initial concerns were increased risk of nerve damage.

Certain limitations continue to plague laparoscopic surgery, however, particularly the use of unarticulated rigid instruments. A good assistant is especially important during TME because two-dimensional view and poor ergonomics can negatively impact any surgeon attempting to visualize nerves during rectal dissection. The marked interest in robotic surgery was largely due to several reasons: (a) the EndoWrist function with seven degrees of freedom; (b) the magnified view of the operative field; (c) the decrease of inherent operator tremor and (d) the precision of instruments was said to be associated with improved functional outcomes. Next, we review the current published literature and compare the outcomes with respect to the different techniques of rectal surgery.

Urinary dysfunction

The studies of open TME are largely cohort studies. In older studies between the years1990 and 2002, the median follow-up ranged from 3 to 41 months, and the number of study subjects ranged from 68 to 199 patients (Table 1) [11,12,30-36]. While reporting incidences and outcomes varied with different classifications, the overall incidence can be as high as 33%. When nonconventional TME was performed, 19% vs 7% of subjects who underwent standard TME had increased difficulty in voiding [35,36].

In newer studies between the years 2003 and 2013, the median follow-up ranged from 7 to 218 months (Table 1) [37-48]. The number of study subjects ranged from 20 to 292 patients, although 3 studies were performed at the same study center [37,38,42]. When autonomic nerve preservation (ANP) was practiced, the incidence of urinary dysfunction ranged from 2.1 to 24.4% [37-41,45-48]. This was significantly higher (between 22.4 and 79.1%) if ANP was not observed during surgery [37,38,42,45-48]. Long-term indwelling catheter use was 2.1-10.8% with ANP, but 30-40% of these patients had successful trial of void within 6 months and improvement of function [37,38,42]. In patients who had pelvic lymph node dissection (PLND), the incidence in these studies noted a high rate of urinary dysfunction [32,41,43], and patients may take 2-3 weeks to return to presurgical residual urinary volume [42].

For laparoscopic TME, there were 2 randomized controlled trials comparing laparoscopic vs open TME techniques which reported functional outcomes [10,49]. While function was not the primary endpoint, there were no significant overall differences in urinary function between the laparoscopic and TME groups. This finding was also supported by other recent cohort studies (Table 1) [50-53,59]. Of note, Jayne et al. and Hur et al. reported that voiding function would reduce after surgery with some restoration by 6 months and full recovery to preoperative levels by 12 months [49,51]. The incidence of minor urinary dysfunction was low regardless of technique (7.3% open TME vs 5.4% laparoscopic TME) [50]. When laparoscopic TME was compared with robotic TME, Kim et al. noted that restoration of urinary function was faster at 3 months [54]. There are, however, no differences seen in bladder function between the 2 groups 12 months after surgery in other studies as well [55-58]. In all studies the incidence of permanent IDC was very low at 0.5%.

Sexual dysfunction

For open TME studies, the incidence of sexual dysfunction ranges from 11% to 85%. If ANP is performed, the incidence of male erectile dysfunction ranges from 1.5% to 49.15%, and the incidence of male ejaculatory dysfunction ranges from 12% to 44.2% (Table 2) [11,12,29–36,39–41,43–48,58,60–62]. There were, however, 2 older studies that noted a very high incidence of erectile dysfunction (69%) and ejaculatory dysfunction (85%) even with ANP [30,32]. If ANP is not performed, the incidence can be as high as 100% [40]. Only a few studies have reported female sexual dysfunction (due to low questionnaire response rate of female candidates compared with male counterparts). The incidence of orgasmic dysfunction reported in females is significantly lower at 0 to 10.5% [12,34,35,45,58], with only 2 studies reporting incidence of dyspareunia ranging from 4.15% to 66.7% [35,45]. If pelvic lymph node dissection is performed, the incidence of dysfunction can range from 21.5% to as high as 90% [33,41].

Post-laparoscopic TME results in 12.8% to 57% erectile dysfunction, 40% to 43.7% ejaculatory dysfunction and 7.1% to 41% overall male sexual dysfunction (Table 2) [10,49-57,59,63]. When comparing the incidence of sexual dysfunction following laparoscopic TME with open TME, the benefit of minimally invasive surgery is inconsistent. Jayne et al. and Quah et al. randomized trials suggested worse sexual functioning in laparoscopic rectal surgery (23.9-41%) compared with open surgery (10.8-18%) [10,49]. However, Liu's randomized trial demonstrated lower incidence of sexual dysfunction in laparoscopic TME with ANP vs open procedure (11.6% vs 16.9%); it is worth noting that the result shows no statistical significance [63]. In a more recent cohort study and a non-randomized trial, there is no statistically significant difference in the incidence of both erectile and ejaculatory dysfunction among laparoscopic vs open TME [51,56], but one study noted worse functioning of both sexes in the open group [50]. When robotic surgery was compared with laparoscopic TME, Kim et al. and Park et al. noted no difference in function between both arms [54,58], but D'Annibale et al. noted that while the erectile function worsened 1 month after surgery, it was almost completely restored at 12 months in the robotic group but only partially restored in the laparoscopic group with an incidence of 57% [55].

Conclusion

It is evident from this summary of outcomes that there is still more work to be done. It must be remembered that colorectal surgery underwent a huge wave of change in the time frame of the conducted studies. There was a radical shift from open to minimally invasive and robotic techniques, as well as a worldwide application of neoadjuvant chemoradiotherapy for locally advanced rectal cancers, which was all largely happening in the preceding 10-15 years. Many of the cohort studies also appear to have strong selection bias and perhaps include many institutions' learning curves for the minimally invasive techniques. In addition, many of these studies have a short follow up of < 2 years with a relatively small number of subjects. We must therefore interpret all study results cautiously. While open TME results initially appeared to have high urinary and sexual dysfunction rates, the later studies showed no differences in outcomes between laparoscopic and open techniques in terms of urinary dysfunction. There appeared to be worsened sexual functioning in the earlier studies for laparoscopic TME, but no difference was observed in the later studies. The argument of

Table 1. Incidence of urinary dysfunction when comparing open, laparoscopic and robotic total mesorectal excision (TME)

Literature	Year of publication	Study design	Procedure type	Subjects evaluated	Duration of follow- up (months)	Complications
Open TME Hojo et al. [30]	1990	Case series - Retrospective	TME and ANP	134	12	Multi-level sacrifice of HP and PP contributes to more severe urinary dysfunction and increasing bladder
Havenga et al. [29]	1996	Case series - Retrospective	TME	136		hypertonia No significant change in both male and female urinary
Sugihara et al. [31]	1996	Case series - Prospective	TME and ANP	199	12	Tunction Urinary dysfunction: 0% in intact HP and PP 3.9% in sacrifice HP and intact bilateral PP 6.5% in sacrifice HP and unilateral PP
Saito et al. [32]	1998	Case series	TME and ANP	167		30.8% in complete resection of pervic autonomic nerves No requirement for long-term indwelling catheter
Ishikura et al. [33] Maas et al. [35]	1999 2000	Case series - Prospective Case series - Prospective	TME and ANP with PLND TME and ANP with radical rectal resection	49	Median 41 Median 42	Urinary frequency in both sexes: 22%;
Nesbakken et al. [34]	2000	Case series - Prospective	TME	49	м	Minor incontinence in both sexes: 19% No significant change in urinary symptom score (both sexes) except for increase in female incontinence score
Maurer et al. [36]	2001	Case control	TME vs Non-TME conventional rectal excision	09	Minimum 3	No significant change in urinary symptom among both groups
Pocard et al. [12]	2002	Case series - Prospective	TME	20	Up to 60	Both sexes requiring indwelling catheter: 0%. No significant change in urinary symptom score (both sexes).
Kim et al. [11]	2002	Case series - Prospective	TME	89	Median 8.7	Males requiring long-term indwelling catheter: 0%. No significant change of male IPSS score.
Junginger et al. [38]	2003	Case series - Prospective	TME with pelvic plexus visualisation	150	Median 24	Overall urinary dysfunction (both sexes): 12%.
						Complete or partially visualised Any 4:5% vs non-Visual- ised: 38.5%. Required short-term indwelling catheter (both sexes) : 10.7%.
Kneist et al. [37]	2004	Cohort study	TME or partial mesorectal excision	229		Overall urinary dysfunction (both sexes): 4.1% with complete ANP vs 22.4% without ANP. Both sexes long-term indwelling catheter 8.8%.
Shirouzu et al. [40]	2004	Case control	TME with vs without ANP	292	Median 218	Overall urinary dysfunction (both sexes): < 20% with complete ANP us > 90% without ANP
Sterk <i>et a</i> l. [39]	2005	Case series - Prospective	TME	52	8	Urinary dysfunction (both sexes): 24.4% at 14 days and 8.1% at 3 months
Wang et al. [45]	2005	Case control	Open TME with vs without ANP	96		Female urinary dysfunction: 6.25% with ANP vs 31.25% without ANP (SD).

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Literature	Year of	Study design	Procedure type	Subjects	Duration of follow-	Complications
	publication			evaluated	up (months)	
						Residual urine > 50 ml: 10.41% with ANP us 31.25% without ANP. Significantly longer duration to recover from urinary summans in nations without ANP.
Laing et al. [46]	2006	Case control	Open TME with vs without ANP	236		Symptoms in patients without ANF. Overall male urinary dysfunction: 12.71% with ANP us 70.33% without ANP (SD only occurred in Duke A and
Kyo et al. [41]	2006	Case control	Open TME and ANP with us without PLND	37	Minimum 7	Duke B comparative groups) Male urinary dysfunction: 33% with PLND vs 9.5% with- out PLND
Dong et al. [47] Kneist et al. [42]	2007	Case series - Retrospective Cohort study	Open TME with ANP Open TME, operative bladder neurophysiology monitor with vs without ANP	12 4 62	Median 20	Failed urinary catheter removal on post-op day 3: 9.7% Urinary dysfunction (both sexes): 2.1% with complete ANP vs 60% without ANP.
						Both sexes required long-term indwelling catheter: 2.1% with ANP vs 33.3% without ANP. Significantly worse in urinary symptom score without ANP.
Akasu et al. [43]	2009	Case series - Prospective	LAR/APR with selected ANP and PLND	69	0.5	Residual urine > 50 ml at day 14 (SD)
						Bilateral pelvic-plexus preservation without PLND: 4% Bilateral pelvic-plexus preservation with PLND: 27% Unilateral pelvic-plexus preservation with PLND: 76% No pelvic-plexus preservation with PLND: 100%
Zhao et al. [48]	2011	Case control	Open TME and ANP vs TME alone	84	Up to 24	Male urinary dysfunction: 24.4% in TME and ANP vs 79.1% in TME alone (SD)
Cakabay et al. [44]	2012	Case series - Prospective	TME	20	12	Male urinary dysfunction: 25% Male requiring long-term indwelling catheter: 5%
Laparoscopic/robotic TME						
Quah et al. [10]	2002	Randomized controlled trial	Laparoscopic assisted vs open TME	80	Median 36	Urinary dysfunction (both sexes): 0%.
						Males required long-term indwelling catheter: 2.5% in laparoscopic νs 0% in open (no SD). Females required long-term indwelling catheter: 2.5% in laparoscopic νs 0% in open (no SD).
Jayne et al. [49]	2005	Randomized controlled trial	Laparoscopic colonic vs open rectal vs laparo- scopic rectal	246	Up to 76	Urinary dysfunction (both sexes): 21% in laparoscopic colonic, 35% in open/laparoscopic rectal.
Liang et al. [52]	2007	Case series - Prospective	Laparoscopic TME	74	Minimum 3	No significant change in urinary symptom score after 6 months. Urinary dysfunction (both sexes): 17.6%

(continued)

Table 1. Continued						
Literature	Year of publication	Study design	Procedure type	Subjects evaluated	Duration of follow- up (months)	Complications
Kim et al. [54]	2012	Cohort study	Laparoscopic <i>vs</i> robotic TME	38	12	Urinary dysfunction (both sexes): 3.3% in robotic vs 5.1% in Lap (no SD)
McGloen et al. [50]	2012	Cohort study	laparoscopic vs open TME	143	Minimum 6	No significant change in unnary symptom score (both sexes)
D'Annibale et al. [55]	2013	Cohort study	Laparoscopic vs robotic TME	09	12	Male urinary dysfunction: 0%
Runkel et al. [53]	2013	Case series - Prospective	Laparoscopic nerve orien- tated mesorectal excision	274	12	Male requiring long-term indwelling catheter: 0.5%
Luca et al. [57]	2013	Case series - Prospective	Totally robotic TME	74	17.03	Male urinary dysfunction: 0% Female urinary dysfunction: 0%
Hur et al. [51]	2013	Non-randomized control trial	Laparoscopic vs open TME	97	12	Urinary dysfunction (both sexes): 5.4% in laparoscopic vs 7.3% in open (no SD). Required long-term indwelling catheter (both sexes): 0%. No significant changes in IPSS score between both
Zeng et al. [59]	2013	Cohort study	Laparoscopic vs open TME with ANP	81	9	groups. Overall male urinary dysfunction: 16.28% in laparoscopic vs 15.79% in open (no SD)
Park <i>et al.</i> [56]	2014	Case control	vs Robotic TME	64	12	No difference in IPSS scores

ANP. autonomic nerve preservation; PLND: pelvic lymph node dissection; IPSS: International Prostate Symptom Score; SD: statistical significant difference; HP: Hypogastric plexus; PP: Pelvic Plexus

Table 2. Incidence of sexual dysfunction when companing open, laparoscopic and robotic total mesorectal excision (TME)

Literature	Year of publication	Study design	Procedure type	Subjects evaluated	Duration of follow- up (months)	Complications
Open TME Hojo <i>et al.</i> [30]	1990	Case series - Retrospective	TME and ANP	39	12	Male erectile dysfunction: 69%
Havenga et al. [29]	1996	Case series - Retrospective	TME	136		Male ejaculatory dystunction: 85% Male ejaculatory dysfunction: 13%
						< 60 year-old sexual dysfunction (both sexes): 14% > 60 year-old sexual dysfunction (both sexes): 33%
Sugihara et al. [31]	1996	Case series - Prospective	TME and ANP	57	Median 53	Male sexual dysfunction with and without ANP: 66.7% Male sexual dysfunction with complete ANP: 29.6% Male erectile dysfunction with complete ANP: 3.8%
						Male erectile dysfunction with incomplete ANP: 20% Male ejaculatory dysfunction with complete ANP: 18.5% Male ejaculatory dysfunction with incomplete ANP: 100%
Enker et al. [58]	1997	Case control	TME and ANP in APR vs LAR	136	up to 60	Male sexual dysfunction in LAR: 14% Male sexual dysfunction in APR: 43%
						Male ejaculatory dysfunction in LAR: 12% Male ejaculatory dysfunction in APR: 15%
						Female sexual dysfunction: 14%
Saito et al. [32]	1998	Case series	TME and ANP	167		remale orgasmic dysrunction: 9% Unsuccessful in preserving sexual function
Ishikura et al. [33]	1999	Case series - Prospective	TME and ANP with PLND	15	Median 41	Male sexual dysfunction: 21.5%
Maas et al. [35]	2000	Case series - Prospective	TME and ANP with radical	47	Median 42	Male erectile dysfunction 8.3% with preserved SHP and
			rectal resection			100% without complete preservation of ANP. Male ejaculatory dysfunction 12.5% with preserved SHP
						and 100% with sacinics of 3rm. Female dyspareunia: 66.7%
Nesbakken et al. [34]	2000	Case series - Prospective	TME	49	9	Male erectile dysfunction: 16.7%
						Male ejaculatory dysfunction: 7.6%
Maurer et al. [36]	2001	Case control	TME vs Non-TME	09	Minimum 3	remare organized your cutour. 5 % Male sexual dysfunction increased from 33% preop to
			Conventional Rectal			93% postop in conventional group vs from 47% to 78%
			Excision			in TME group. Male erectile dysfunction increased from 25% preon to
						94% postop in conventional group vs from 42% to 74%
						in TME group.
						Male ejaculatory dysfunction increased from 12% preop
						47% in TME group
			Ę	Ç	1	Insufficient data for female sexual dysfunction analysis
Kim et al. [11]	2002	Case series - Prospective	IME	89	Median 8./	Male efectite dysfunction: 19.1% Male ejaculatory dysfunction: 13.2%
						Significant worsening of ILEF Score in all domains

(continued)

ations	: 11.0% 1: 0%	.7% with ANP vs 63.5% : 44.2% with ANP vs 71.2%	% with ANP vs 100% with- : 35% with ANP vs 100%	.1%		rith ANP us 37.5% without 1: 10.5% with ANP us 45.9% 1ction: 12.5% with ANP us	tion: 49.15% with ANP us	% with PLND vs 10% with- : 90% with PLND vs 30%	7%))	vation without PLND: 5% vation with PLND: 44% ervation with PLND: 55% 1 with PLND: 100%	.3% in TME and ANP vs : 26.8% in TME and ANP vs	79.1% in TME alone (SD) Male erectile dysfunction: 5%
Complications	Male erectile dysfunction: 0% Male ejaculatory dysfunction: 11.0% Female orgasmic dysfunction: 0%	Male erectile dysfunction: 32.7% with ANP vs 63.5% without ANP (SD). Male ejaculatory dysfunction: 44.2% with ANP vs 71.2%	without ANP (SD) Male erectile dysfunction: 21% with ANP vs 100% without ANP. Male ejaculatory dysfunction: 35% with ANP vs 100%	Male erectile dysfunction: 55.1% Male eigenlatory dysfunction: 3.6%	Male sexual dysfunction: 35.2% Male erectile dysfunction: 33.3% with ANP vs 63.2% without ANP (SD) Male ejaculatory dysfunction: 43.8% with ANP vs 70.0%	without ANP (SD) Female dyspareunia: 4.15% with ANP vs 37.5% without ANP (SD) Female orgasmic dysfunction: 10.5% with ANP vs 45.9% without ANP (SD) Female sexual arousal dysfunction: 12.5% with ANP vs	54.16% without ANP Overall Male erectile dysfunction: 49.15% with ANP us	93.70% without ANF (3D) Male erectile dysfunction: 50% with PLND vs 10% without PLND (no SD) Male ejaculatory dysfunction: 90% with PLND vs 30%	without PLND (SD) Male erectile dysfunction: 37.7% Wele significant dysfunction: 42.9%	Male erectile dysfunction (SD)	Bilateral pelvic-plexus preservation without PLND: 5% Bilateral pelvic-plexus preservation with PLND: 44% Unilateral pelvic-plexus preservation with PLND: 55% No pelvic-plexus preservation with PLND: 100%	Male erectile dysfunction: 29.3% in TME and ANP vs 76.7% in TME alone (SD) Male ejaculatory dysfunction: 26.8% in TME and ANP vs	79.1% in TME alone (SD) Male erectile dysfunction: 5%
Duration of follow- up (months)	Up to 36		Median 218	e	Up to 84			Minimum 7		12		Up to 24	
Subjects evaluated	20	104	129	29	105 105	96	236	30	124	99		84	20
Procedure type	TME	Open TME with vs without ANP	TME with ANP vs without ANP	TME	Open TME Open TME with vs without ANP	Open TME with vs without ANP	Open TME with vs without	TME and ANP with vs with- out PLND	Open TME with ANP	Open LAR/APR with selected ANP and PLND		Open TME and ANP vs TME alone	TME
Study design	Case series - Prospective	Case control	Case control	Case series - prospective	Case series - Retrospective Case control	Case control	Case control	Case control	Case series - Retrospective	Case series - Prospective		Case control	Case series - Prospective
Year of publication	2002	2003	2004	2005	2005	2005	2006	2006	2007	2009		2011	2012
Literature	Pocard et al. [12]	Wang et al. [60]	Shirouzu et al. [40]	Sterk <i>et al.</i> [39]	Wang et al. [61] Wang et al. [62]	Wang et al. [45]	Laing et al. [46]	Kyo et al. [41]	Dong et al. [47]	Akasu et al. [43]		Zhao et al. [48]	Cakabay et al. [44]

(continued)

Table 2. Continued						
Literature	Year of	Study design	Procedure type	Subjects	Duration of follow-	Complications
	publication			evaluated	up (months)	

Quals et al. [10] 2002 Randomized controlled trial open TME Laparoscopic rectal to spen TME 80 Median 36 Jayne et al. [49] 2005 Randomized controlled trial rectal to spen TME 246 Up to 76 Liang et al. [52] 2007 Case series - Prospective Laparoscopic rectal to spen TME 60 6 Liue tal. [63] 2002 Randomized control trial Laparoscopic us open TME 143 Minimal 6 Kim et al. [54] 2012 Cohort study Laparoscopic us robotic 60 12 Modisen et al. [55] 2013 Cohort study Laparoscopic us robotic 41 17.038 Luca et al. [57] 2013 Cohort study Laparoscopic us open TME 74 17.038 Runkel et al. [53] 2013 Cohort study Laparoscopic us open TME 50 12 Zeng et al. [53] 2013 Cohort study Laparoscopic us open TME 81 6 Zeng et al. [53] 2013 Cohort study Laparoscopic us open TME 81 6 Zeng et al. [54] 2013 Cohort study	Laparoscopic/ Robotic TME						
Randomized controlled trial Laparoscopic rectal us aparoscopic colonic			Randomized controlled trial		08	Median 36	Sexual dysfunction (both sexes) 23.9% in laparoscopic vs 10.8% in open. Male erectile dysfunction: 40% in laparoscopic vs 13.7% in open. Male ejaculatory dysfunction: 40% in laparoscopic vs 4.6% in open.
2007 Case series - Prospective Laparoscopic TME 60 2009 Randomized control trial Laparoscopic us open TME 119 with ANP 143 2012 Cohort study Laparoscopic us robotic 38 TME TME 144 2013 Case series - Prospective Totally Robotic TME 74 2013 Case series - Prospective Laparoscopic us open TME 74 tated mesorectal excision 1 Laparoscopic us open TME 50 trial Laparoscopic us open TME 50 trial Laparoscopic us open TME 81 with ANP 50 Laparoscopic us open TME 81 with ANP 64 Laparoscopic us open TME 81 Laparoscopic us robotic 60 Laparoscopic us open TME 81 with ANP 64 TME			Randomized controlled trial		246	Up to 76	Male sexual dysfunction: 41% in laparoscopic rectal, 23% in open rectal and 4% in laparoscopic colonic. Male erectile and ejaculatory dysfunction more common in laparoscopic and open rectal surgery but none in laparoscopic colonic. Female sexual dysfunction: 28% in laparoscopic rectal, 18%, open rectal 18%, and 8% in laparoscopic colonic.
2012 Cohort study Laparoscopic vs open TME 119 with ANP 2012 Cohort study Laparoscopic vs open TME 143 2012 Cohort study Laparoscopic vs robotic 2013 Case series - Prospective Totally Robotic TME 74 2013 Case series - Prospective Laparoscopic nerve orien 42 tated mesorectal excision 1 2013 Cohort study Laparoscopic vs open TME 50 trial Laparoscopic vs open TME 60 2014 Case control Laparoscopic vs robotic 60 TME			Case series - Prospective		09	9	Male erectile dysfunction: 37.5% Male ejaculatory dysfunction: 43.7% Female orgasmic dysfunction: 32.1% Female dyspareunia: 39.2%
2012 Cohort study Laparoscopic vs open TME 143 2012 Cohort study Laparoscopic vs robotic 38 TME TME 2013 Case series - Prospective Totally Robotic TME 774 2013 Case series - Prospective Laparoscopic nerve orien 42 tated mesorectal excision trial 2013 Non-randomized control Laparoscopic vs open TME 50 trial 2014 Case control Laparoscopic vs robotic 64 TME Case control Laparoscopic vs robotic 64 TME		60	Randomized control trial		119	12	Male sexual dysfunction: 11.6% in laparoscopic vs 16.9% in open (no SD)
2012 Cohort study Laparoscopic vs robotic 38 TME 2013 Cohort study Laparoscopic vs robotic 60 TME 2013 Case series - Prospective Totally Robotic TME 74 Case series - Prospective Laparoscopic nerve orien- 42 tated mesorectal excision Non-randomized control Laparoscopic vs open TME 50 trial 2013 Cohort study Laparoscopic vs open TME 81 with ANP TME TME			Cohort study		143	Minimal 6	Worse IIEF score in male open group, especially erectile function (SD). Worse FSFI score in female open group (SD).
al. [55] 2013 Cohort study Laparoscopic vs robotic 60 TME 2013 Case series - Prospective Totally Robotic TME 74 Case series - Prospective Laparoscopic nerve orien 42 tated mesorectal excision Non-randomized control Laparoscopic vs open TME 50 trial 2013 Cohort study Laparoscopic vs open TME 81 with ANP TME TME			Cohort study		38	12	Male erectile dysfunction: 13.3% in robotic vs 12.8 % in laparoscopic (no SD)
2013 Case series - Prospective Totally Robotic TME 74 Case series - Prospective Laparoscopic nerve orien- 42 tated mesorectal excision Mon-randomized control Laparoscopic vs open TME 50 trial 2013 Cohort study Laparoscopic vs open TME 81 with ANP TME TME		13	Cohort study		09	12	Male erectile dysfunction: 5.6% in robotic vs 57% in laparoscopic at 12 months(SD)
2013 Non-randomized control Laparoscopic vs open TME 50 trial 2013 Cohort study Laparoscopic vs open TME 81 with ANP TME TME	53]		Case series - Prospective Case series - Prospective	u	74	17.03 12	No change in IIEF or FSFI score Male erectile dysfunction: 18.2%
2013 Cohort study Laparoscopic vs open TME 81 with ANP 2014 Case control Laparoscopic vs robotic TME			Non-randomized control trial		50	12	Male sexual dysfunction: 22.7% in open vs 7.1% in laparoscopic. No significant change of IIEF score by 12 months in both groups.
2014 Case control Laparoscopic vs robotic 64 TME		13	Cohort study		81	9	Male erectile dysfunction: 27.91% in laparoscopic vs 28.95% in open (no SD). Male ejaculatory dysfunction: 25.58% in laparoscopic vs 23.68% in open (no SD).
			Case control		64	12	IIEF Scores for robotic group higher at 6 months compared to laparoscopic group

ANP: autonomic nerve preservation; PLND: pelvic lymph node dissection; SHP: superior hypogastric plexus; APR: abdominoperineal resection; LAR: low anterior resection; IIEF: International index of Erectile Function, FSH: Female Sexual Function Index; SD: statistical significant difference

robotic technique also remains largely undecided with perhaps improved shorter-term urinary function within 3 months, but no difference in 12 months. There is no similar benefit for sexual function as well. It is important that successful rectal surgery be based not only on histopathological oncologic outcomes but also on the quality-of-life indices of every patient being restored to near normal as soon as possible. The decision about which technique to use will largely depend on a multitude of factors including tumour factors, pelvic anatomy and perhaps cost effectiveness. Nonetheless, regardless of the technique to be applied, the rectal surgeon must be aware of anatomical landmarks and hazards while dissecting to avoid nerve injury and make a dedicated attempt to preserve for autonomic nerve functioning.

Conflict of interest statement: none declared.

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