Complete laparoscopic nephroureterectomy with intravesical lockable clip

Milan Hora¹, Viktor Eret¹, Tomáš Ürge¹, Jiří Klečka¹, Ivan Trávníček¹, Ondřej Hes², Fredrik Petersson^{2,3}, Petr Stránský¹

Charles University Hospital, Department of ¹Urology and ²Pathology, Pilsen, Czech Republic ³Department of Pathology, National University Health System, Singapore

KEY WORDS

nephrectomy ▶ ureter ▶ urothelial carcinoma ▶ laparoscopy

ABSTRACT

Introduction. We present a cohort of patients with lowstage pelviureteric neoplastic disease who underwent complete laparoscopic nephroureterectomy (CLNUE) with intravesical lockable clip (IVLC). Due to the absence of a standard technique of NUE, the study was not randomized.

Materials. From 1/2010 to 1/2012, 21 patients were subjected to CLNUE-IVLC. The first step was transurethral excision of the ureterovesical junction with Collin's knife deep into the paravesical adipose tissue. The ureter was grasped with biopsy forceps and the distal end of the ureter was occluded with lockable clip. The applicator was introduced through a 5 mm port inserted as an epicystostomy. The patients were rotated to flank position and CLNUE followed. The endoscopically introduced clip on the distal ureter is proof of completion of the total ureterectomy.

Results. The mean operation time was 161 (115-200) min. In four (19.0%), the application of the clip failed and CLNUE was completed with non-occluded ureter. In three cases, subsequent laparoscopic nephrectomy was converted to open surgery. In two cases, the distal ureterectomy was completed with pluck technique through a lower abdominal incision that was also used for extraction of the specimen. There were four complications (Clavien II 2x, IIIb, V). Follow-up was available for all – mean 10.6 (range: 0-25) months. One died of disease generalization within 11 months.

Conclusion. CLNUE-IVLC is fast and safe. If needed, the endoscopic phase can be switched to open NUE. Disadvantages include: the need to change the position of the patient, the risk of inability to apply the clip on the distal ureter, and the risk of an unclosed defect of the urinary bladder.

INTRODUCTION

Despite expanded indications for conservative surgery of urothelial tumors (UT - formerly transitional cell carcinoma) of the upper urinary tract (UUT), radical nephroureterectomy (NUE) with complete removal of the distal ureter including the bladder cuff is the standard surgical technique used for most patients with UT of the UUT. Choosing the best procedure for this group of patients in everyday clinical practice is frequently a challenging task. While laparoscopic nephrectomy as a part of NUE was first described in 1991 [1] and it is to-date broadly accepted [2], the approach to the distal ureter and the timing of the ureterectomy are still disputed [3]. Several techniques have been developed to remove the distal intramural part of the ureter during laparoscopic NUE and it is very difficult to choose the best procedure for a given patient in everyday clinical practice.

Complete laparoscopic NUE (CLNUE) has previously been performed with a stapler [4]. There is a risk of residual tumor at the stapling site and titan clips may constitute a nidus for the formation of cystolithiasis [5, 6].

Two groups [7-9] have described a technique for the division of the ureterovesical junction with a thermo "sealing" system (Ligasure[®] Atlas[®]). We considered this modification of CLNUE as an excellent method and we have previously performed this on 14 patients. However, we found a significant risk of incomplete resection of the intramural part of the ureter [10]. For this reason, we have been searching for another method of CLNUE.

Exclusive laparoscopic sharp excision of the bladder cuff with intracorporeal suturing [11-15] appears too difficult for us. Thus, we have decided to start CLNUE with excision of the ureterovesical junction with Collin's knife followed by CLNUE. The main problem with this procedure is the risk of occlusion of the ureter to prevent spillage of urine containing tumor cells during laparoscopic pluck nephroureterectomy. Several methods have been described to date: (1) cauterization of the ureteric ostium only, (2) endoloop [16, 17], (3) Hem-o-lok[®] clip [18-20], or (4) fibrin sealant [21].

We chose to close the ureter with a lockable Hem-o-lok[®] clip, which was introduced through a 5 mm intravesical port in the suprapubic area rather than endoscopically [18,19,20]. We have labeled this technique as CLNUE-WILC (with intravesical lockable clip). In this study we characterize the applicability of this technique and report our experiences.

MATERIAL AND METHODS

The study is prospective, but due to the absence of a standard technique for NUE, the study was not randomized and comparative. From 1/2010 to 1/2012, 38 patients with suspected UT of UUT were indicated for surgical treatment. Four underwent conservative surgery (one Ho:YAG ablation, one ureteroscopic resection, one nephroscopic resection with resectoscope, and one open resection of the ureter) and 34 NUE. Thirteen underwent some type of open surgery (advanced cases with open nephrectomy or laparoscopic nephrectomy with open ureterectomy for tumor of the distal ureter) and 21 NUE by CLNUE- IVLC (main inclusion criteria: not suitable for conservative treatment, no tumor of distal ureter, no advanced tumor by CT, no contraindications to laparoscopy, or no concomitant bladder tumor).

Follow-up (months)	0.0	24.7	11.1	20.9	19.3	17.5	17.1	15.8	15.4	14.9	14.5	14.0	9.0	7.3	7.1	4.0	3.5
Histology	nc	0-PRCC	nc	nc	CRCC	inflamma- tion	nc	nc	nc	nc	NC	nc	nc	nc	nc	Meta- stasis of colorectal cancer	UC
Grade	-	2	. 	2	-	0	2	-	2	2	2	-	2	2	2	-	ę
рТ	7	1a	σ	2	3a	0	2	-	č	a+CIS	2	ŋ	-	-	-	4	e
Follow-up	Death from heart failure in day of operation		UC of the right ureter and ostium, death for generalisation to skeleton in 11.1 months								3Mo after NUE, hemodialysis started						
Open approach, complications	Open distal ureterectomy					2 nd post-op day open PE for bleeding				Open nephrectomy (history of suphrenic abscess)		Open distal ureterectomy		Open distal ureterectomy	Abscess of wound – Staphylococcus aureus	Open nephrectomy (fixation to surrounding structures)	Open nephrectomy (fixation to surrounding structures)
Notes			History of recurrent BT (pT1G2), TURBs, BCG					History of recurrent BT, TURBs, BCG		History of TU-RB fro BT pT1G2, ESKD, hemodialysis	UC of L ureter at the same time, Ho:YAG laser ablation, ESKD						
Failed Weck	-		-			-	-										
Hospital stay	NA	œ	~	12	80	14	6	თ	80	13	œ	7	9	11	18	=	7
Blood loss	100	0	200	100	100	150	150	50	0	500	150	100	100	150	10	500	600
Time of surgery	200	165	140	180	170	180	150	115	115	150	130	190	140	140	180	180	190
Open distal ureterectomy	-									-		-		-			
Open nephrectomy																-	-
CLNUE		-	-	-	-	-	-	-	-		-		-		-		
Localization of tumour	MU	٩	٩	۵.	۵.	ЧD	٩	۵.	٩	MU	٩	MU	۵.	٩U	۵.	Ð	۵.
Side	_	Ж	_	_	æ	œ	_	۳	Ж	۳	<u>م</u>	_	_	٣	~	£	٣
Age	84.1	55.8	68.0	72.1	77.9	65.9	81.6	59.5	76.1	67.2	82.9	52.6	70.0	66.9	71.2	58.0	65.4
Sex	Σ	Σ	Σ	Σ	Σ	Σ	Σ	Σ	Σ	Σ	Σ	Σ	Σ	Σ	Σ	Σ	Σ
Number	-	2	с	4	Ð	9	7	œ	6	10	7	12	13	14	15	16	17

2.7	2.2	1.6	0.3	10,6	7,5	0.0		₹
nc	nc	nc	nc	Mean	SD	Min.	Max.	pillary renal ce ne – of whole
т	с	-	-					ocytic pa OMO: Tim
-	e	-	e					C – onco were N(
								eft, M – male, Mo – month(s), O-PRC nostic ureteroscopy. Notes: All cases
Hem-o-lok size L, open distal ureterectomy (2 cm)								– end-stage kidney disease, L – I urothelial carcinoma. URS – diag
Previous URS and stent		Urinary leakage from drain 6 days						– urinary bladder tumor, ESKD - ureter proximal-middle. UC – (
				4	19.0%			of BCG, BI or. UP-UM
œ	9	n	8	9.4	3.0	6.0	18.0	instillation adder tumo
350	200	0	0	167.1	175.8	0.0	600.0	ravesical ion of bla
195	142	140	185	160.8	26.5	115.0	200.0	ory of int ral resect
-				7	33%			1а, BCG – hist 3 – transureth
	-	-	-	14	67%			r adenon aht. TUR
M	۵.	٩	٩					tomy fo is. R – ric
~	٣	œ	_					Irenalec Jal pelv
65.2	61.4	77.0	58.2	68.4	9.1	52.6	84.1	AE – ad P – ren
Σ	Σ	Σ	ш	Mean	SD	Min	Max	viations: / oma [29].
18	19	20	21					Abbre

Technique of CLNUE- IVLC: The procedure starts in the lithotomy position with endoscopy. The ureterovesical junction is excised transurethrally with Collin's knife (the paravesical adipose tissue must be clearly visible). From the suprapubic region, a 5 mm port is introduced in the urinary bladder. The stump of the ureter is grasped with biopsy forceps and on the end of ureter, a Hem-o-lok® clip size ML is applied (the applicator is introducible through the 5 mm port). In broader ureters, a 10 mm port and size L clips may be used. The patient is rotated to the flank position and a standard laparoscopic nephrectomy via a transperitoneal approach is performed [10, 22]. The transperitoneal approach is more familiar to us than the retroperitoneoscopic one. The ureter is not divided. One additional 5 mm port in the suprapubic region is introduced; the skin incision from the previous endoscopic phase is used. The gonadal vessels are transected. The peritoneum is incised above the ureter up to the urinary bladder. The ureter is dissected along and under the iliac vessels with a harmonic scalpel or Ligasure® Advance® or Blunt tip 35 mm®. This phase is delicate due to the relatively narrow operation space and the close relation of iliac vessels and the bowel. The ureter is completely separated and the Hem-o-lok® clip must be clearly visible to constitute proof of completion of the ureterectomy. The urinary bladder stays unclosed. A pelvic drain is placed through a port in the suprapubic area. The specimen is extracted in a bag through a lower abdominal muscle splitting incision. A catheter is left for seven days. A standard postoperative follow-up is conducted according to established protocols of UT of UUT. All CLNUE-IVLCs were performed by three endoscopically and laparoscopically experienced surgeons.

RESULTS

The clinicopathologic data and results are summarized in Table 1. A few points to emphasize include: four complications (Clavien II 2x, IIIb and V) [23] - wound infection at the site of extraction (*Staphylococcus aureus*), urine leakage from the pelvic drain for 6 days (bladder catheter was removed on the 8th postoperative day following cystoradiography that did not reveal leakage), one open prostatectomy on the 2nd postoperative day because of an enlarged prostate (BPH with hematuria), and one patient died of heart failure on the day of operation. In four cases (19.0%) application of the clip failed and CLNUE was concluded with a non-occluded ureter and the risk of dissemination of tumor cells in urine paravesically. In the first case the patient had a ureteral stent and the ureter was incrassate, which is why we did not want to apply 10 mm port to facilitate the introduction of the Hem-o-lok® clip size L. In another case, the stented ureter was also incrassate and we introduced a 10 mm port to the urinary bladder and applied a size L clip without difficulty. There was one incidence with the inability to grasp the ureter by endoscopic forceps. In two cases the intramural ureters were cut by Collin's knife. In three cases the laparoscopic nephrectomy was converted to the open surgery (flank incision and lumbolaparotomy) - in one case this was due to extensive adhesions in the abdominal cavity (a history of open cholecystectomy with evacuation of subphrenic abscess), and in the other two cases it was due to advanced tumor growth with perirenal and periureteral adhesions. In three cases, laparoscopic nephrectomy was followed by open distal ureterectomy with the standard pluck method, because the laparoscopic approach was not feasible due to poor access of the laparoscopic instruments to the small pelvis. The incision was also used for extraction of the resected specimen. Four patients with non-UT histology were judged to be UT by preoperative imaging. Follow-up (mean 10, range: 0-22 months), including results of the control endoscopy, are known in all patients. One patient with UC of the renal pelvis pT2N0M0G2 had UC in the contralateral distal ureter and died after 11 months due to extensive metastatic disease, mainly to bone.

DISCUSSION

procedure (endoscopy, rotation of patient, laparoscopy); Non-UC histology: Tumors were described by radiologists on CT/MRI as a suspicious UC tumor.

It should be noted that the aim of this work is not to comprehensively discuss the whole complex problem of NUE. Our experience prompted us to review this topic in two recent publications [10, 22] and now we will focus the discussion on complete laparoscopic NUE with emphasis on the method of removing the distal part of the ureter including the bladder cuff. In our



Fig. 1. The steps of the endoscopic phase in lithotomy position. The ureterovesical junction is excised transurethrally with a Collin's knife to the paravesical adipose tissue. The stump of the ureter is grasped with biopsy forceps and the end of the ureter is clipped with a Hem-o-lok[®] clip size ML (an applicator is introduced through the 5 mm port inserted as an epicystostomy).

view, due to the disadvantages mentioned in the introduction, the method involving a stapler should be abandoned. The other options of distal ureterectomy as a part of CLNUE are as follows: (1) a thermosealing technique [7, 8,9], (2) the sharp excision of the bladder cuff with intracorporeal suturing (a purely laparoscopic technique) [11, 12, 15] including modification with a bulldog clamp [13], (3) robotic [24], or (4) purse string technique [14].

As pointed out previously, the thermosealing system technique has the risk of leaving the intramural part of the ureter intact [10]. The laparoscopic NE with ensuing sharp excision of the ureterovesical junction and closing of the defect with suture is an ideal but challenging method. This method has several modifications as mentioned above. We consider the technique with any variant intracorporeal suturing technically more challenging and time consuming [10, 22]. The variant with the da Vinci robotic system [24, 25] decreases the technical difficulty of intracorporeal suturing. The disadvantages of the da Vinci system include: high cost, lack of tactile sensation, long set-up time, and unavailability of the robotic system in many hospitals. An exotic technique is the pneumovesicum approach [26] in which three 5 mm ports are introduced to the bladder and insufflated with CO_2 – pneumovesicum (10-12 mm Hg). The distal ureter, bladder cuff, and intramural ureter are then completely dissected free using electrocautery. As soon as the distal ureter is dissected an endo-loop knot is used to ligate



Fig. 2. Ports for left side nephroureterectomy. Six ports, on the left side, but five usually suffice. The same skin incision as for the epicystostomy is used for the suprapuble port.

the ureter. We do not have experience with this technique and we feel this technique to be complicated. Sotelo et al. have used the pneumovesical technique as well. The ureter is excised through a laparoscopic single port introduced to the urinary bladder and the defect is closed with intravesical suture [27].

Due to the factors mentioned above we prefer a variant of excision of the ureterovesical junction but with another method for sealing of the ureter. We have long-term experience with excision of ureterovesical junctions using Collin's knife. Previously we used it as a pluck technique combined with open and thereafter laparoscopic or retroperitoneoscopic nephroureterectomy, and, later on, we also used it as a part of antegrade mini-invasive NUE [22].

We found Pathak et al.'s idea [18, 19] of endoscopically closing the excised ureter with lockable clip to be excellent, although the introduction of the Hem-o-lok[®] clip via endoscope appears to be difficult. Pathak et al. performed their technique in 25 cases with a mean total operative time of 164 (range: 105-235) minutes. No pelvic complications in were reported and there were no perivesical tumor recurrences with mean follow-up of 26 (range: 11-44) months. We have decided to apply the Hem-o-lok[®] clip size ML via an intravesical 5 mm port introduced through the suprapubic area. Regarding our first nine cases, we had found the exact same method described in the literature [20]. This work has strengthened our approach. Ahlawat et al. [28] published another three



Fig. 3. The dissected nephroureterectomy specimen during the operation. The tumor can be seen in the upper calyx with Hem-o-lok[®] size ML at the end of the ureter (see detail).

cases in 2011 with almost the same technique. The authors labeled this technique "suprapubic transvesical single-port technique for control of lower end of ureter during laparoscopic nephroure-terectomy".

Reportedly, occlusion of the ureter may be performed instead of clip with electro-coagulation only (it is probably less reliable) or with fibrin sealant injection [21].

CONCLUSION

CLNUE- IVLC is a relatively simple, reproducible, and minimally invasive method with minimal risks of tumor spillage and seeding. The main disadvantage seems to be the risk of an unclosed defect of the urinary bladder, but based on our own experience, as reported in this paper and the available literature, we have not found any significant complications emerging from this. Another disadvantage is failure in applying the Hem-o-lok[®] clip, in which the technique is concluded without closing the ureter, and is generally thought to carry a higher risk of extravesical tumor recurrence, but as described recently, this technique has comparable oncological results to the open distal ureterectomy [3, 6]. Failures in clip application were experienced only in early cases and, with increasing experience, this problem was avoided. Importantly, if needed, the endoscopic phase can be transformed to open NUE or it can be combined primarily with open surgery. Some may consider conversion to open distal ureterectomy/nephrectomy as a failure of the method. The method whereby closing of the ureter is performed allows the procedure (NUE) to be completed safely in complicated cases (obesity, advanced cases, previous intraabdominal surgery etc.).

Acknowledgments:

The work was supported by the Czech government research project MSM 0021620819 and grant IGA NT 12010-5/2011.

REFERENCES

- Clayman RV, Kavoussi LR, Figenshau RS, et al: Laparoscopic nephroureterectomy: initial clinical case report. J Laparoendosc Surg 1991; 1 (6): 343-349.
- Capitanio U, Shariat SF, Isbarn H, et al: Comparison of oncologic outcomes for open and laparoscopic nephroureterectomy: a multi-institutional analysis of 1249 cases. Eur Urol 2009; 56: 1-9.
- Li WM, Shen JT, Li CC, et al: Oncologic outcomes following three different approaches to the distal ureter and bladder cuff in nephroureterectomy for primary upper urinary tract urothelial carcinoma. Eur Urol 2010; 57 (6): 963-969.
- Shalhav AL, Dunn MD, Portis AJ, et al: Laparoscopic nephroureterectomy for upper tract transitional cell cancer: the Washington University experience. J Urol 2000; 163 (4): 1100-1104.
- Matin SR, Gill IS: Recurrence and survival following laparoscopic radical nephroureterectomy with various forms of bladder cuff control. J Urol 2005; 173: 395-400.
- Macejko AM, Pazona JF, Loeb S, et al: Management of distal ureter in laparoscopic nephroureterectomy-a comprehensive review of techniques. Urology 2008; 72 (5): 974.
- Tsivian A, Benjamin S, Sidi AA: A sealed laparoscopic nephroureterectomy: a new technique. Eur Urol 2007; 52: 1015-1019.
- Simone G, Papalia R, Leonardo C, et al: Laparoscopic versus open nephroureterectomy: comparison of operative data and preliminary results of oncological outcome. Eur Urol Suppl 2007; 6: 221 (Abst. 793).
- 9. Simone G, Papalia R, Guaglianone S, et al: *Laparoscopic versus open nephroureterectomy: perioperative and oncologic outcomes from a randomized prospective study.* Eur Urol 2009; 56: 520-526.

- Hora M, Stránský P, Eret V, et al: Complete laparoscopic nephroureterectomy with thermo-sealing system. CEJU 2010; 63 (2): 77-81.
- 11. Hemal AK, Kumar A, Gupta NP, Seth A: *Retroperitoneal nephroureterectomy* with excision of cuff of the bladder for upper urinary tract transitional cell carcinoma: comparison of laparoscopic and open surgery with long-term follow-up. World J Urol 2008 26 (4): 381-386.
- Hattori R, Yoshino Y, Komatsu T, et al: Pure laparoscopic complete excision of distal ureter with a bladder cuff for upper tract urothelial carcinoma. World J Urol 2009; 27 (2): 253-258.
- Cho HJ, Kim SJ, Yoon BI, et al: A Novel Bulldog Clamp Technique for Management of a Distal Ureter and Bladder Cuff During Laparoscopic Nephroureterectomy. J Endourol 2010; 24 (11): 1719-1720.
- Shoma AM: Purse-string technique for laparoscopic excision of a bladder mucosal cuff in patients with transitional cell carcinoma of the upper urinary tract: initial report with intermediate follow-up. BJU Int 2009; 104: 1505-1509.
- Ghazi A, Shefler A, Gruell M, et al: A Novel Approach for a Complete Laparoscopic Nephroureterectomy with Bladder Cuff Excision. J Endourol 2010; 24 (3): 415-419.
- Gill IS, Soble JJ, Miller SD, Sung GT: A novel technique for management of the en bloc bladder cuff and distal ureter during laparoscopic nephroureterectomy. J Urol 1999; 161: 430-433.
- 17. Agarwal DK, Khaira HS, Clarke D, Tong R: *Modified transurethral technique for the management of distal ureter during laparoscopic assisted neph-roureterectomy*. Urology 2008; 71: 740-743.
- Pathak S, Watcyn-Jones T, Lavin V, et al: A novel closed system laparoscopic nephroureterectomy for upper tract urothelial cancer. J Urol 2008; 179 (Suppl 4): 159A.
- Pathak S, Watcyn-Jones T, Doyle D, Oakley N: Laparoscopic nephro-ureterectomy for upper tract urothelial cancer: cystoscopic closed system pluck. Ann R Coll Surg Engl 2008; 90 (6): 524-525.
- 20. Sesmero JH, Redondo CC, Delgado MC, et al: *Intravesical sealing of the distal ureter in nephroureterectomy*. Arch Esp Urol 2010; 63 (3): 223-229.
- Mueller T, DaJusta D, Cha D, et al: Ureteral Fibrin Sealant Injection of the Distal Ureter During Laparoscopic Nephroureterectomy - A Novel and Simple Modification of the Pluck Technique. Urology 2010; 75 (1): 187-192.
- Hora M, Eret V, Ürge T, et al: Antegrade miniinvasive nephroureterectomy (AMNUE - laparoscopic nephrectomy, transurethral excision of ureterovesical junction and lower abdominal incision). Urol Int 2009; 83 (3): 264-270.
- Dindo D, Demartines N, Clavien PA: Classification of Surgical Complications. A New Proposal With Evaluation in a Cohort of 6336 Patients and Results of a Survey. Ann Surg 2004; 240 (2): 205-213.
- 24. Park SY, Neony W, Ham WS, et al: *Initial experience of robotic nephroureter*ectomy: a hybrid-port technique. BJU Int 2009; 104 (11): 1718-1721.
- Hu JC, Silletti JP, Williams SB: Initial experience with robot-assisted minimally-invasive nephroureterectomy, J Endourol 2008; 22 (4): 699-704.
- Guzzo TJ, Schaeffer EM, Allaf ME: Laparoscopic Radical Nephroureterectomy With En-Bloc Distal Ureteral and Bladder Cuff Excision Using a Single Position Pneumovesicum Method. Urology 2008; 72: 850-852.
- 27. Sotelo R, Ramírez D, Carmona O, et al: *A novel technique for distal ureterectomy and bladder cuff excision.* Actas Urol Esp 2011; 35 (3): 168-174.
- Ahlawat RK, Gautam G: Suprapubic transvesical single-port technique for control of lower end of ureter during laparoscopic nephroureterectomy for upper transitional cell carcinoma. Indian J Urol 2011; 27 (2): 190-195.
- 29. Ürge T, Hes O, Ferda J, et al: *Clinical characteristics of oncocytic papillary renal cell carcinoma.* World J Urol 2010; 28 (4): 513-517.

Correspondence

Milan Hora Department of Urology, University Hospital 13, E. Beneše Street 305 99 Plzeň, Czech Republic phone: +42 0 377 402 171 horam@fnplzen.cz