1 A case of duodenal perforation and pancreatic bleeding after flexible 2 ureteroscopy for right renal pelvis UTUC

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15 Abstract

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17 Upper tract urothelial carcinoma (UTUC) can be managed by flexible ureteroscopy (f-URS) and tumor laser ablation if kidney-sparing surgery is possible. This procedure can be affected by minor 18 19 to serious complications, including life-threatening sepsis, ureteral strictures, and ureteral and renal 20 pelvis injuries. Here we present the case of a 53-year-old man with history of high grade right renal pelvis and bladder tumor who undergone multiple endoscopic treatments and has already refused 21 radical surgery. We performed f-URS and laser ablation with Thulium: YAG laser for UTUC 22 23 recurrence of right renal pelvis, but the procedure was stopped due to significant bleeding which 24 impaired vision. Postoperatively the patient developed hematemesis and hemodynamic instability due to duodenal lesion and active bleeding documented at CT scan. An emergency exploratory 25 26 laparotomy was performed to drain hemoperitoneum, repair duodenal lesion and concurrent radical 27 right nephroureterectomy was carried out. A second surgery was necessary for repairing duodenal 28 fistula. After 1 week the patient presented again with recurrent hematemesis and hemorrhagic shock. 29 He underwent angiography and selective embolization of the duodenal branch of superior 30 mesenteric artery and as well as branches of gastroduodenal artery successfully and the patient 31 recovered with no other complications. This is the first case of duodenal perforation and pancreatic 32 bleeding due to flexible ureteroscopy and laser ablation of right renal pelvis urothelial carcinoma.

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Key words: UTUC; laser ablation; bleeding; duodenum.

37 Background

39 Upper tract urothelial carcinoma (UTUC) is an uncommon neoplasm and accounts for only 5-10% 40 of all urothelial carcinomas [1]. Kidney-sparing endoscopic management of UTUC is a feasible option in patients with solitary kidney and/or impaired renal function, as well as in cases of bilateral 41 42 or low-risk tumours [2]. Flexible ureteroscopy (f-URS) can be used to perform diagnostic biopsies 43 as well as ablate the tumour in the ureter or renal pelvis using laser technology. Potential risks of this procedure range from minor complications such as UTI and haematuria to serious 44 45 complications including life-threatening sepsis, ureteral strictures and ureteral and renal pelvis injuries. [3][4]. Here we present the first described case of duodenal perforation and pancreatic 46 47 bleeding due to flexible ureteroscopy and laser ablation of right renal pelvis urothelial carcinoma. 48

49 Case description

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51 A 53year-old man, with significant medical history of peptic ulcer disease on treatment with protonpump inhibitor (PPI), was referred to our Urology Department for a high-risk upper tract urothelial 52 cancer (UTUC) involving right renal pelvis and ureter. He was first treated in 2017 for right renal 53 pelvis tumour and histology showed pTa G3 urothelial carcinoma. He refused right 54 nephroureterectomy. During a follow-up period of 2 years, he underwent at least six endourological 55 procedures (ureteroscopy and biopsies and endoluminal instillation of Mitomicin C through MJ 56 stent) to treat recurrent UTUC. He has also undergone multiple endoscopic bladder resections 57 58 (TURB) for concurrent high grade bladder cancer followed by intravesical immunotherapy with BCG. Considering the extent of the disease, multiple recurrences and the limitation of the 59 endoscopic treatment, he was repeatedly offered right radical nephroureterectomy (RNU) but 60 61 patient declined radical treatment.

In February 2019, he underwent right f-URS. Wireless and sheathless "no-touch" technique was
performed using an 8.5 F flexible digital ureteroscope (Flex-XC, Karl Storz, Tuttlingen, Germany)
[5]. Multiple papillary lesions were found in all the major calyces and the renal pelvis. Biopsy was
performed with a tipless 1.9 F nitinol basket, followed by tumor ablation with a 200 µm fiber for

66 Thulium: YAG (Tm:Yag) laser (Cyber-TM, Quanta System, Samarate, Italy). The laser was set at 67 10 watts for the procedure. After an initial bloodless ablation, endoscopic vision deteriorated due to 68 development of significant bleeding. Laser power was increased up to 30 watts in the effort of 69 controlling the ongoing bleeding, but unsuccessfully. Decision was then made to terminate the 70 procedure and a single J ureteral stent was placed.

On post-operative day (POD) 1, the patient presented with acute hematemesis and haematuria. CT
scan showed a large clot in the right renal pelvis clot. Duodenoscopy was also carried out in
emergency and the patient was found to have a 1cm perforation at the second part of the duodenum.
This was treated endoscopically with the application of 4 metallic clips (Fig 1).

- 75 Unfortunately, the next day, patient developed hemodynamic instability and a drop of haemoglobin
- from 13 to 9g/dL. An emergency exploratory laparotomy was performed with intraoperative
 findings of massive hemoperitoneum due to active bleeding from a 1cm perforation of the anterior
 duodenum wall and from the pancreatic head (Fig 2).
- After the abdomen was washed out, the pancreatic head bleeding was controlled with haemostatic sutures and the duodenal perforation repaired. Right radical nephroureterectomy was also performed at the same setting. The patient received 4 pints of packed cells transfusion during surgery. Post-operatively, the patient was clinically stable but admitted to the Intensive Care Unit for monitoring. On POD 8th, drain output was suspicious for enteric content, hence a CT scan and gastrografin swallow was performed. The scans demonstrated duodenal fistula. The patient was then brought back into the operating room for repair of the duodenal fistula (Fig 3).
- Patient recovered gradually following the second duodenal repair surgery. However, after 1 weeks
 the patient presented with recurrent hematemesis and haemorrhagic shock. CT angiogram
 documented recurrent active bleeding near the previous metallic clips used in the repair of the
 duodenal perforation. Patient underwent selective embolization of the duodenal branch of superior
 mesenteric artery and as well as branches of gastroduodenal artery successfully (Fig 4, Fig 5).
- Subsequently, patient recovered without further hematemesis and haemoglobin levels remained
 stable. He was discharged from hospital on the 30th post-operative day since first surgery.
- 93 Final histological examination of the right radical nephroureterectomy specimen was reported to be
 94 pTis urothelial carcinoma of the right renal pelvis. He is now still on follow up, with the last
 95 bladder recurrence in May 2022.
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98 Discussion and Conclusions

99 According to the current European Association of Urology (EAU) Guidelines, RNU remains the gold standard for high-risk UTUC[2]. Nevertheless, conservative treatment should be considered as 100 an option in patients with imperative indications for kidney-sparing surgery, such as in solitary 101 102 kidney, bilateral UTUC or chronic renal failure, as well as for clinically low risk UTUC [6]. In some cases where the patient chooses to avoid radical surgery, it is important to ensure that the 103 104 patient is aware of the possible risks of disease progression and the necessity of close endoscopic follow-up. To date, f-URS is useful for both endoscopic tumour ablation and a close postoperative 105 surveillance of UTUC after kidney-sparing treatment [7]. Endoscopic procedures in the upper 106 107 urinary tract are associated with the risk of trauma to the ureter and pyelocaliceal system[8]. These injuries are classified as a major complication of f-URS and usually reported to be due to the use of 108 a ureteral access sheath (UAS) [4]. In our case, we did not use a UAS, and therefore, the pelvic 109 wall perforation must have been due to tumor ablation with Thulium: YAG laser in the setting of 110 poor visibility due to significant bleeding during the tumour ablation. 111

In a review of URS complications by Linehan et al. [3], the authors found a rate of bleeding ranging from 1.6% to 27.3% but only a few were serious enough to require hospitalization and/or blood transfusion. Bleeding during URS treatment for UTUC was more frequently associated with patients who had previously received adjuvant instillations [3]. Renal pelvis wall perforation is an even rarer event, with rates varying from 1.3% to 7.4% of cases. There are also some disease117 related factors to consider, such as tumor location and invasiveness. In our case, the patient 118 presented with a large tumour involving the entire anterior wall of renal pelvis and all the calyces 119 and the diseased urothelium is more prone to injury and perforation.

The choice of laser in the ablation of the tumour also contributes to the risk of injury to the 120 collecting system and hence the characteristics of the laser must be considered. Proietti et al 121 122 evaluated the effects of both Tm:YAG and Holmium:YAG (Ho:YAG) lasers on upper urinary tract urothelium, with a focus on incision depth and coagulation area[9]. This study showed a lower 123 penetrative power for Tm:YAG (due to lower peak-power, better water absorption and continuous 124 mode) compared to Ho:YAG, with a higher coagulation effect without excessive carbonization of 125 126 tissue. Despite these advantages in choosing Tm: YAG laser, we encountered major bleeding in our case, resulting in poor vision that led to the inadvertent perforation of the renal pelvis with the 127 128 involvement of the duodenum and the pancreas.

In addition, our patient had a history of peptic ulcer disease, and this underlying pathology may
have contributed to the fragility of the duodenal wall. The second segment of duodenal "C" lies
over the right renal hilum, hence its proximity resulted in the injury with the renal pelvis perforation
(Fig 6).

Duodenal perforation is a rare condition but is associated with high morbidity and mortality, 133 ranging from 8% to 25% [10]. Isolated duodenal injuries after trauma are rare and pancreas is 134 frequently injured concomitantly due to their close anatomical relationship. The second segment of 135 the duodenum is the most commonly injured part (36%). When isolated minor CT findings are 136 discovered the clinical case can be managed conservatively with close monitoring, otherwise 137 patients usually require surgical intervention. Endoscopic management with supportive medical 138 therapies is first line therapy and is highly effective. However, approximately 10% of all patients 139 will either continue to bleed or experience re-bleeding within 48 h of the endoscopic treatment. 140 141 While surgical therapy has historically been considered the next line of treatment for upper GI refractory bleeding, angioembolization has now become the next line of therapy[11]. Given the dual 142 supply to the duodenum from the celiac trunk (GDA), as well as the superior mesenteric artery 143 144 (through the inferior pancreaticoduodenal arcades), embolization that is distal to the site, and 145 proximal to the bleeding is needed for effective embolization[11] The main complication after 146 trans-arterial embolization is bowel ischemia. Although the upper gastrointestinal tract has a rich 147 collateral blood supply, ischemic complications can still occur in 7 to 16% of cases [12] in which unfavourable evolution is very likely. 148

In the literature we find only very limited number of cases of iatrogenic injury of the duodenum
during endourological surgery: one in a patient with indwelling right ureteral DJ stent [13] and 4
cases of duodenal perforation during percutaneous nephrolithotomy [14].

Here, we describe the first case of duodenal injury occurring during operative ureteroscopy for 152 UTUC. The injury was due to laser ablation of the tumour in poor visibility condition on 153 background of tissue fragility. Urologists should pay special attention when using laser in 154 endourology, especially during soft tissue treatment: the last generation high power lasers are very 155 effective but also able to deliver an energy that may clearly exceed the amount needed becoming 156 dangerous and counterproductive especially in case of severe bleeding. In this scenario, when 157 controlling of the bleeding in the upper urinary tract is not achievable, always consider stopping the 158 procedure in order to avoid life-threatening complication like this and involving interventional 159 160 radiologists.

Early recognition, diagnosis and timely intervention are crucial in the management of these rare butserious complications.

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Fig.1: EGDS and application of metallic clips for duodenal bleeding.



Fig. 2: CT scan and hemodynamic instability: active bleeding and hemoperitoneum.



Fig. 3: Abdomen X-Ray with gastrografin: duodenal fistula

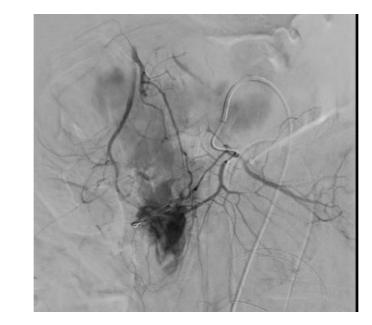


Fig. 4: Active bleeding from mesenteric and gastroduodenal artery at angiography.

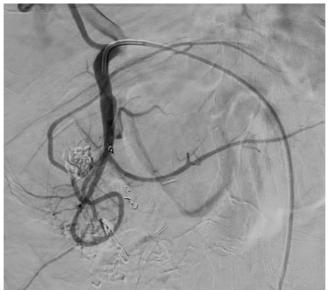


Fig. 5: Haemostasis after selective angioembolization.

