# Treatment and outcome of fibroepithelial ureteral polyps: A systematic literature review

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# Abstract

**Introduction:** Fibroepithelial polyps of the ureter are rare. Cases and small series are reported in the literature. The treatment of choice, outcome and appropriate follow-up regimen remain unclear. **Methods:** We conducted a systematic literature review of papers

reporting fibroepithelial polyps of the ureter in adult patients. Articles published before 1980 were excluded.

Results: The search yielded 144 papers, of which 68 met the inclusion criteria. A reference scan from the included 68 yielded an additional 7 new articles. In total, our study included 75 articles (68 + 7). A total of 134 patients were described. Most patients had a single lesion (range: 1-10). The median length of the polyp was 4.0 cm (range: 0.4-17.0). The percentage of polyps resected endoscopically increased from 0% before 1985 to 67% after 2005. Two perioperative complications were reported in 72 procedures (2.8%): a deep venous thrombosis and a case of mesenteric lymphadenopathy. Both of these occurred after open surgery. Follow-up data were available for 57 patients. The median follow-up was 12 months (range: 1-180). Four patients (7.0%) developed recurrent complaints: 2 had urinary stones, 1 had a ureteral stricture and 1 had recurrence of the polyp. Three of these events followed endoscopic resection, and occurred within a year after the procedure. Conclusion: Endoscopic resection of fibroepithelial polyps seems to be safe and effective. It is minimally invasive and should be considered the gold standard where endoscopic expertise is available. We advise follow-up imaging by computed tomographic intravenous urography after 3 months and ultrasound after 1 year to detect late complications.

# Introduction

Fibroepithelial polyps of the ureter are rare benign lesions of the urinary tract. They are non-epithelial benign tumours

of mesodermal orgin, a category that also includes leiomyomas, fibromas, neurofibromas and lymphangiomas. They are derived from mesenchymal tissue, in contrast with malignant transitional cell and squamous cell tumours which are derived from the epithelium.<sup>1</sup> In general, benign tumours account for 20% of all ureteral tumours.<sup>2</sup>

Most urologists will only infrequently encounter patients with a fibroepithelial polyp. They rely on the medical literature as guidance for treatment and follow-up. However, most published papers are case reports or describe small patient series. One larger review dates from 1980 and is outdated in the era of flexible ureteroscopy.<sup>3</sup> Several recent review articles focus on fibroepithelial polyps in children.<sup>4,5</sup> The aim of this study was to provide a systematic overview of the literature on fibroepithelial polyps in adult patients. In this way, evidence-based choices can be made for treatment and follow-up of patients with this condition.

#### **Methods**

A systematic literature review was performed of fibroepithelial polyps of the ureter. We conducted a search of the PubMed, EMBASE and Cochrane Library databases for English language literature published between January 1980 and January 2014. Papers published before 1980 were disregarded as they date from an era before the widespread use of (flexible) ureteroscopy. The search terms used were 'fibro-epithelial OR fibroepithelial AND ureter OR ureteral.'

Papers were included if they presented one or more patients aged 18 years or older with one or more fibroepithelial polyps of the ureter. Patients with only polyps in the renal pyelum or only in the bladder were excluded. The reference lists of the selected articles were then searched for more papers that met the inclusion criteria.

Statistical analysis was done using SPSS 17.0 for windows (SPSS Inc, Chicago, IL). Exploratory univariate analysis was performed to identify factors influencing the choice for endoscopic or open resection of polyps. The Pearson Chi Square test was used to compare proportions. The independent samples t-test was used to compare normally distributed continuous variables. The Mann-Whitney U-test was used to compare continuous variables that were not normally distributed.

# Results

### Search results and study characteristics

The primary search yielded 144 results. Of these 144 papers, 7 concerned different conditions other than fibroepithelial polyps or were review papers. Children only were described in 47 articles and no distinction was made in another 2 papers. Eight papers described fibroepithelial polyps in other locations than the ureter (urethra, bladder and renal pelvis). Three papers were veterinary studies and 9 papers were only published as abstracts (presentations at conferences). Therefore, of the 144 papers, we excluded 76.

The reference lists of the remaining 68 articles were searched for additional papers of interest. This search yielded another 7 studies, which were included. In total, 75 articles met the inclusion criteria (Fig. 1).<sup>1-3,6-77</sup> Most papers (56/75) presented a single patient. The largest series was reported by Williams and colleagues<sup>67</sup> and included 15 patients. Together, the 75 papers described 134 adult patients with one or more fibroepithelial polyps of the ureter.

#### Patient and polyp characteristics

Of the 131 patients with available data, 71 were female (55.9%). Most patients presented with flank pain and/or hematuria (Table 1). Intravenous ureterography was the most frequently used imaging modality (84.4%). Most patients had only 1 polyp (86.5%), but the maximum reported was 20 polyps in a single patient (both ureters were affected). The median size of the polyps was 4.0 cm (range: 0.4–17.0). Polyps were evenly distributed among the left and right ureter, with a few patients with bilateral polyps. Polyps were encountered over the entire ureter, but most frequently in the proximal ureter. The polyp caused hydronephrosis in 41.7% of patients. Urinary tract stones were present in 20.8% of patients and concomitant urological abnormalities, such as duplicated urinary tracts, were found in 7.0% (Table 1). A coincidental transitional cell carcinoma, located on the epithelial of the fibroepithelial polyp, was reported in one of the patients in this review.<sup>73</sup>

# Treatment, outcome and follow-up

Mode of treatment was reported for 113 patients (Table 2). Two patients were initially managed conservatively. One of these patients was being treated for advanced gastric cancer and had over 20 fibroepithelial polyps in both ureters.<sup>32</sup> The authors deemed it impossible to resect all of them without damaging the ureter. Therefore, the patient received bilateral nephrostomy catheters. No follow-up was reported. The other patient returned 11 months later with recurrent flank

#### **Table 1. Patient characteristics** Articles with available data (total = 131)† Age, years (median; min-max) 38 (18-76) 126 Female gender 71 (55.9%) 127 No. polyps 104 1 90 (86.5%) 2 4 (3.8%) 3 5 (4.8%) >4 5 (4.8%) 4.0 (0.4-17.0) Size, cm, median (range) 104 Side of polyp distribution 108 Left 55 (50.9%) Right 50 (46.3%) Both 3 (2.8%) 121 Level of polyp distribution Proximal ureter 55 (45.5%) Middle ureter 25 (20.7%) Distal ureter 41 (33.9%) Symptoms 116 Flank pain 77 (66.4%) Hematuria 50 (43.1%) (Recurrent) UTI 5 (4.3%) Other 3 (2.6%) 7 (6.0%) No symptoms 109 Imaging Ultrasound 23 (21.1%) IVU 92 (84.4%) СТ 35 (32.1%) **RPG/RUG** 42 (38.5%) Hydronephrosis 96 40 (41.7%) Intussusception 7 (7.3%) 96 Urolithiasis 21 (20.8%) 101 100 Other urological abnormalities 3 (3.0%) **UPJ** stenosis Duplicated upper tract 2 (2.0%) Cystic ureteritis 1 (1.0%)

tOf the total 134 adult patients, 3 did not have available data. UTI: urinary tract infection; IVU: intravenous urography; CT: computed tomography; RPG: retrograde pyelography; RUG: retrograde ureterography; UPJ: ureteropelvic junction.

1 (1.0%)

Ureterocele

pain and hematuria and was then treated by endoscopic resection.  $^{\rm 7}$ 

The method of endoscopic resection was further specified in 44 cases. Most polyps were resected using electrocautery or Holmium:YAG laser (Table 2). The remaining patients were treated by polypectomy via ureterotomy, partial ureterectomy or nephro-ureterectomy. The proportion of patients treated by endoscopic resection increased from none before 1985 to 66.7% after 2005 (Fig. 2). Two perioperatieve complications were reported<sup>33,40</sup> in 72 procedures (2.8%): one deep venous thrombosis<sup>40</sup> and one case of mesenteric lymphadenopathy.<sup>33</sup> Both of these occurred after open surgery.

Outcome data were available of 57 patients. Four patients developed recurrent symptoms (7.0%); 3 of these patients had initially been treated endoscopically and 1 by surgical resection. Two of these patients had recurrent urinary tract stones.<sup>9,13</sup> One had recurrent or residual growing polyp (after an incomplete initial resection).<sup>17</sup> One patient had a ureteric stricture.<sup>56</sup> Three of these late complications occurred within 1 year and a case of urolithiasis occurred after 3 years.

Exploratory univariate analysis was performed to investigate which factors may have influenced urologists in their choice for endoscopic or surgical resection of the polyp (Table 3). Polyps that were endoscopically resected were less likely to be in the proximal ureter, more likely to have concomitant urolithiasis, and more likely to have been operated after 2000. Larger polyps were more likely to have been removed surgically by open approach. However, successful ureteroscopic resection of polyps up to 16 cm was reported.<sup>56</sup>

Table 2. Treatment, out	Table 2. Treatment, outcome and follow-up			
	Value	Cases with data available (total = 134)		
Treatment		113		
Conservative	2 (1.8%)			
Endoscopic resection	49 (43.4%)			
Ureterotomy	26 (23.0%)			
Partial ureterectomy	26 (23.0%)			
Nephro-ureterectomy	10 (8.8%)			
Method endoscopic resection		44		
Holmium:YAG laser	16 (36.4%)			
Electrocautery	23 (52.3%)			
Mechanical	5 (11.4%)			
Perioperative complications	2 (2.8%)	72		
Follow-up		134		
Until discharge only	77 (57.5%)			
≤6 months	17 (12.7%)			
7 months–1 year	13 (9.7%)			
1–2 years	12 (9.0%)			
2–5 years	12 (9.0%)			
>5 years	3 (2.2%)			
Outcome		57		
Complaint free	53 (93.0%)			
Recurrent or remnant stones	2 (3.5%)			
Ureteric stricture	1 (1.8%)			
Recurrent polyp	1 (1.8%)			

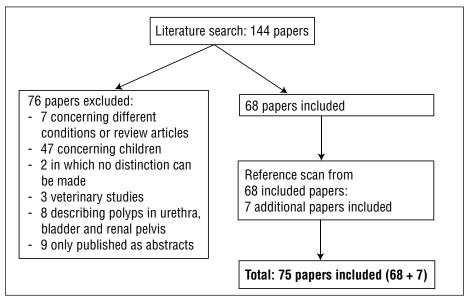


Fig. 1. Flow chart of included papers.

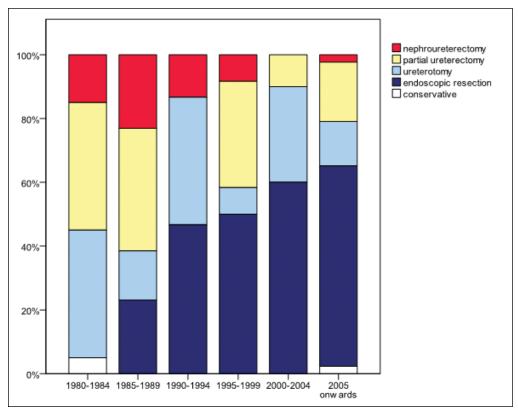


Fig. 2. Trends in treatment of fibroepithelial polyps between 1980 and 2014.

# Discussion

We have provided a comprehensive review of ureteral fibroepithelial polyp described in the English language literature since 1980. Ureteroscopic resection has become the most performed method of treatment. According to data, it is safe and effective. As the least invasive type of surgery, it should be considered the gold standard.

In 1980, Debruyne and colleagues analyzed 112 patients with fibroepithelial polyps treated since 1950.<sup>3</sup> Most complaints were flank pain and hematuria. The authors also describe predominance in male patients and a slight preference for the left ureter. These observations are not supported by the data presented in our review. This may be explained by the fact that Debruyne and colleagues included children in their review. Having been written in the pre-ureteroscopy era, the paper by Debruyne and colleagues advised that surgical resection of the polyp is always necessary. They noted that although local resection is preferred, 37% of patients underwent nephro-ureterectomy instead. Over the past 30 years, this percentage has decreased to almost zero (Fig. 2).

By exploratory univariate analysis we identified possible factors associated with urologists' choice for surgical rather than endoscopic resection. Besides operation after the year 2000, smaller polyp size, concomitant stones and distal location in the ureter were associated with endoscopic resection. The fact that proximal polyps were more likely surgically resected may be explained by the technical difficulty in achieving adequate endoscopic exposure of the polyp in that part of the ureter. Although larger polyps were more likely to be surgically resected, polyps up to 16 cm were successfully removed endoscopically. Therefore, size of the polyp should not be a contraindication for endoscopic resection.

Concomitant transitional cell carcinoma was reported in 1/108<sup>3,78</sup> cases described by Debruyne et al and in 1/134 patients in the current review. Considering the small chance of concomitant malignancy, we advise a form of endoscopic treatment that allows pathologic examination of the resected polyp. If malignancy is found, decision for nephro-ureterectomy or follow-up should be made by a multidisciplinary oncology team taking also the resection margins and grade of the tumour into account. Laser ablation of (parts of) the polyp may be considered after taking representative biopsies for pathological analysis.

In their review article, Debruyne and colleagues reported no recurrence or late complications.<sup>3</sup> The current study revealed 4 patients (7.0%) who developed late complaints: two with remnant or recurrent stones, 1 ureteral stricture and 1 with recurrent polyps. These complications warrant

 Table 3. Factors influencing choice for endoscopic treatment

treatment				
Factor	Endoscopic group	Surgery group	p value	
Female gender	31/47 (66.0%)	32/60 (53.3%)	0.218ª	
Age	$42 \pm 16$ years	36 ± 13 years	0.039 <sup>b</sup>	
Polyp size	3.0 (0.5–16.0)	6.0 (0.4–17)	0.009°	
Location proximal ureter	11/44 (25.0%)	32/59 (54.2%)	0.003ª	
Concomitant urolithiasis	13/39 (33.3%)	8/59 (13.6%)	0.020ª	
Published after 2000	34/50 (68.0%)	18/61 (29.5%)	<0.001ª	
<sup>a</sup> Pearson Chi Square; <sup>b</sup> Independent samples T-test; <sup>c</sup> Mann-Whitney U-test.				

follow-up imaging studies. As 3 of the 4 events occurred within a year of the initial resection, it is advisable to conduct follow-up imaging within 1 year (i.e., after 3 and 12 months). Some authors advise intensive follow-up-regimen with frequent urinalysis, ultrasound and intravenous urography (IVU) or computed tomography (CT) in the first years after surgery.<sup>36,56,68</sup> Considering the pattern of recurrent complaints (mostly within the first year) and the benign nature of the lesion, it is our opinion that a CT-IVU after 3 months and ultrasound of the urinary tract after 1 year are sufficient follow-up methods.

The pathogenesis of fibroepithelial polyps remains uncertain. Factors, such as obstruction, trauma, irritation, infection, exogenous carcinogens, hormonal imbalance, and allergy, have been proposed as causative agents. Stones may cause chronic irritation of the ureteral wall, with or without urinary infection, but their actual role is uncertain.<sup>5</sup>

The association with urolithiasis (20.8% in this study) could be explained in several ways. Firstly, the altered anatomy with ureteral polyps could promote the formation of stones. Secondly, urolithiasis is more likely to become symptomatic if there is already partial obstruction by a ureteral polyp. Thirdly, the presence of stones and the associated irritation of the urothelium could cause the formation of polyps. To our knowledge there is no apparent reason why chronic irritation of the ureter wall leads to transitional cell carcinoma in some patients and fibroepithelial polyps in others.

Although possibly even more rare than in adults, fibroepithelial polyps are occasionally found in the pediatric population. They account for 0.5% of ureteropelvic junction (UPJ) obstructions in children.<sup>79</sup> Pediatric polyps are often located in the UPJ and upper ureter (73.3%);<sup>80</sup> only a handful of cases are reported more distally in the ureter.<sup>81</sup> In contrast to the adult population, there seems to be a male preponderance (89%)<sup>79</sup> and a more frequent involvement of the left ureter (67%). Preoperative diagnosis of ureteral polyps in children is challenging because of the small lumen of the ureter. Malignant ureteral transitional cell tumours have not been reported in children.<sup>4,80</sup> Ultrasonography is still the first choice for the diagnosis, because it is harmless and convenient.<sup>79,80</sup> There are however insufficient data to estimate the specificity of ultrasound for this condition.

Our paper is the most complete overview of the treatment and outcome of ureteral fibroepithelial polyps in adults; however, it has its limitations. An important limitation is that we relied on data from case reports and small patient series. This made our results susceptible to publication bias. Publication is likely to be biased towards larger and multiple polyps. Therefore, the size and the number of polyps reported in this study were probably larger than the average in clinical practice. Regarding the complication rate, it could be hypothesized that there was a publication bias towards uncomplicated procedures. Another limitation of this study is that it does not include papers reported in languages other than English (although if we had included these non-English articles, the number of patients would have likely increased by a fraction). Moreover, it is unlikely that the general conclusions would have been affected.

# Conclusion

or stricture.

Endoscopic resection is a safe and effective treatment for fibroepithelial polyps of the ureter, and should be considered the gold standard where endoscopic expertise is available. Based on the pattern of late complications reported in literature, we advise control CT-IVU at 3 months and control ultrasound after 1 year to detect possible recurrence

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