Original Article Effect of flexible cystoscopy combined with flexible ureteroscopy on short-term outcomes in patients with complex renal stones

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Abstract: Objective: To investigate the short-term efficacy of combining flexible cystoscopy with flexible ureteroscopy in the treatment of complex renal stones. Methods: The medical records of 145 patients with complex renal stones admitted to Yan'an People's Hospital from February 2020 to February 2022 were retrospectively analyzed. Among these, 65 patients treated with flexible ureteroscopy alone constituted the control group. The research group consisted of 81 patients receiving both flexible cystoscopy and flexible ureteroscopy. Outcomes compared between the two groups included stone removal rate, operative time, time to ambulation, hospitalization duration, and intraoperative bleeding. Logistic regression analysis was used to assess the risk of stone retention. Results: In the research group, the stone removal rate was 85.19% and the residual stone rate was 14.81%, compared to a stone removal rate of 70.77% and a residual rate of 29.23% in the control group, with a statistically significant difference (P<0.05). The research group had a significantly longer operative time than the control group (P<0.05). However, intraoperative bleeding and hospitalization duration were significantly lower in the research group (P<0.05). There was no statistically significant difference in time to ambulation between the groups (P>0.05). Multivariate logistic regression analysis identified multiple stones (OR=3.581, P=0.013) as an independent risk factor for residual stones, while stone location outside the lower calyx (OR=0.305, P=0.021) and treatment with combined flexible cystoscopy and ureteroscopy (OR=0.398, P=0.160) were independent protective factors against residual stones. The area under the curve for predicting stone retention based on the number of stones, stone location, and treatment modality were 0.647, 0.642, and 0.606, respectively. Conclusion: The combination of flexible cystoscopy and flexible ureteroscopy in treating patients with complex renal stones significantly improves the stone clearance rate and postoperative patient recovery.

Keywords: Flexible cystoscopy, flexible ureteroscopy, complex kidney stones, stone retention

Introduction

Kidney stones are a prevalent condition within the urinary system, accounting for approximately 40% of urological interventions and affecting 8 out of every 1,000 adults annually [1]. Over the past few decades, their prevalence and incidence have consistently increased. A survey conducted in China revealed a prevalence rate of 6.4%, with a gender breakdown of 6.5% for men and 5.1% for women [2].

Complex renal stones, often characterized by their staghorn shape or multiple stones exceeding 2.5 cm in diameter [3], are increasingly common in clinical settings, likely occurring due to shifts in dietary habits [4]. Their intricate distribution, irregular shape, and considerable size make them prone to complications such as urinary tract obstruction and infection, potentially leading to renal failure and kidney damage, posing significant health risks [5]. Due to their size, high stone burden, or abnormal renal anatomy, complex renal stones present a significant challenge for surgical management, making them a recognized technical obstacle in clinical practice [6].

With advancements in endoscopic technology, percutaneous nephrolithotripsy has become a

widely used treatment for complex kidney stones. However, this procedure carries risks, including significant intraoperative and postoperative bleeding, which can lead to complications such as nephrectomy and liver, pleural, and intestinal injuries [7].

As medical equipment continues to improve and surgeons gain more experience, minimally invasive procedures such as flexible ureteroscopy, laparoscopic lithotripsy, and combined endoscopic surgeries have gained traction due to their benefits of causing less trauma and enabling quicker recovery [8, 9]. The advantages of these minimally invasive surgeries include reduced postoperative pain, shorter hospital stays, and a faster return to normal activities, making them increasingly preferred for treating complex renal stones.

The flexible ureteroscope and flexible cystoscope are valuable tools in urological procedures due to their unique characteristics [10]. The flexible ureteroscope, which can be maneuvered through the urethra, bladder, and ureter, has a bendable lens that allows it to navigate the natural curves of the urinary tract to access different parts of the kidneys, particularly the upper pole and calyces [11]. This makes it especially useful for examining and treating kidney stones in these areas.

On the other hand, the flexible cystoscope is designed to access the bladder through the urethra. Its soft, flexible body and bendable lens enable it to rotate within the urethra and bladder, allowing visualization of various parts of the bladder lining, including areas that are difficult to reach with a rigid scope [12, 13].

While these instruments offer distinct advantages in their respective applications, the potential benefits of combining their use in treating complex renal stones or other urological conditions have yet to be fully explored. Therefore, this retrospective study collected data from patients with complex kidney stones treated with two different regimens to observe patient outcomes.

Materials and methods

In this retrospective study, we selected 168 patients from Yan'an People's Hospital who met the inclusion criteria from February 2020

to February 2022. Considering the exclusion criteria, 23 cases were excluded. Ultimately, data from 145 patients with complex renal stones were included. Based on the surgical approach, 65 patients treated with flexible ureteroscopy alone were designated as the control group, and 81 patients treated with a combination of flexible cystoscopy and flexible ureteroscopy were designated as the research group. The study was conducted with the approval of the Medical Ethics Committee of Yan'an People's Hospital.

Inclusion criteria

Patients diagnosed with complex kidney stones based on clinical history and imaging findings [14]. Patients with preoperative urinary tract infections treated with anti-infective therapy and well-controlled. Patients treated with cystoscopy combined with ureteroscopy or ureteroscopy alone. Age \geq 18 years old.

Exclusion criteria

Patients with a combination of acute and chronic infectious diseases. Patients with a previous surgical history of kidney stones. Patients with severe cardiovascular, cerebrovascular, pulmonary, hepatic, or other vital organ diseases. Patients with severe upper or lower urinary tract infections. Patients with systemic hemorrhagic disorders where coagulation had not been corrected.

All patients underwent ureteroscopy and double J stent placement (Shenzhen Kangyibo Company, China) prior to surgery. After general anesthesia, the control group was placed in the lithotomy position. A rigid ureteroscope (German Wolf, size F8/9.8) was inserted into the bladder to remove the double J stent. The ureteral opening was located, and a zebra guidewire was inserted into the ureter on the affected side. The rigid ureteroscope was advanced into the affected ureter and then removed. A flexible ureteroscope (German Wolf. size F12/14) sheath was then introduced to the renal pelvis. Upon locating the stone, the zebra guidewire was withdrawn, and a holmium laser fiber (Lumenis, USA) with a frequency and power setting of 20 Hz and 1 J was used to break the stone.

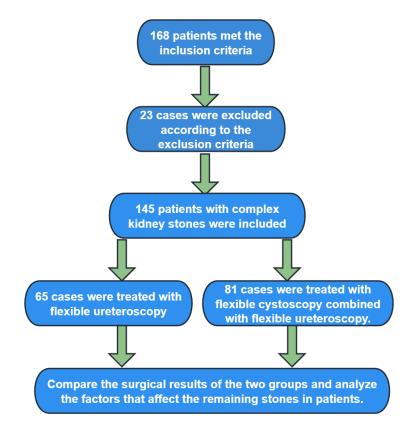


Figure 1. Flowchart.

In the research group, a flexible cystoscope (OLYMPUS, EVIS EXERA III CLV-190) was additionally used to explore each calyx. Residual stones were located and fragmented until no residual stones were detected in the calyx after repeated exploration. Both the double J stent and nephrostomy tube were left in place, and the operation was concluded.

The following factors were analyzed from the collected patient data: gender, age, stone diameter, duration of disease, BMI, site, number of stones, location of stones, degree of hydronephrosis, blood creatinine levels, stone clearance rate, operation time, intraoperative bleeding, time to ambulation, and length of hospital stay. The study flow is illustrated in **Figure 1**.

The one-time stone clearance rate was determined postoperatively. The criteria for onestage stone clearance were as follows: if the diameter of the residual stone was ≤ 4 mm and the anatomy of the upper urinary tract was normal on the 2nd and 3rd postoperative days, it was considered completely cleared. Otherwise, residual stones were noted.

Observation indicators

The primary observation indicators were the postoperative stone clearance rate and residual stone rate for both groups. Secondary indicators included surgery-related metrics such as operation time, time to ambulation, length of hospital stay, and blood loss. The risk of residual stones was evaluated using logistic regression analysis, and the Receiver Operating Characteristic (ROC) curve was drawn to assess the predictive value for residual stones.

Statistical methods

Based on previous data and literature [15, 16], we estimated the stone-free rate of the control group to be approximately

68% and that of the research group to be about 88%. Using the formula n = $(Z_{\alpha/2} + Z_{\beta})^2 \times [p_1 (1 + p_2)^2)^2$ $(p_1) + p_2 (1 - p_2)]/(p_1 - p_2)^2$ estimate the sample size, where α =0.05, Z_{$\alpha/2$}=1.96, β =0.20, $Z_{p}=0.84$, p₁=0.88, p₂=0.68, we calculated that at least 64 subjects were needed per group. Accordingly, 65 individuals were included in the control group and 81 in the research group, based on the hospital's capacity. All statistical analyses were performed using IBM SPSS software version 20.0 (IBM Corp, Armonk, NY, USA). A p-value of less than 0.05 was considered statistically significant. For normally distributed measurement variables, data were described using the median ± standard deviation, and comparisons between two independent groups were made using the independent samples t-test. For categorical variables, chisquare analysis was employed. Logistic regression analysis was used to assess the risk of stone retention, with odds ratio (OR) values calculated for risk assessment. The ROC curve was utilized to analyze the best cutoff value for measures and the predictive value for the risk of stone retention.

	Control group (n=65)	Research group (n=81)	t/χ^2	Ρ
Gender				
Male	35	52		
Female	30	29	1.605	0.205
Age (years)	45.14±9.80	46.70±10.09	0.947	0.345
Stone diameter (mm)	15.42±6.16	14.22±5.41	1.227	0.222
Duration of the disease (months)	26.45±4.86	27.63±4.40	1.521	0.131
BMI (kg/m²)	23.24±2.01	23.13±2.52	0.276	0.783
Site				
Left kidney	34	39		
Right kidney	31	42	0.250	0.617
Number of stones				
Single-shot	26	36		
Frequent	39	45	0.292	0.589
Location of stones				
Renal calyces	40	45		
Non-renal calyces	25	36	0.531	0.466
Degree of hydronephrosis				
No or slight water accumulation	42	44		
Moderate or severe water accumulation	23	37	1.579	0.209
Blood creatinine (µmol/L)	64.90±8.25	63.51±8.16	1.016	0.312

Table 1. Comparison of baseline data

BMI: Body mass index.

	-			
	Stone	Clearance	Stones	Residual
	removal	Clearance	remain	rate
Control group (n=65)	46	70.77%	19	29.23%
Research group (n=81)	69	85.19%	12	14.81%
X ²				4.481
Р				0.034

Results

Comparison of baseline information

The baseline data of the two groups were compared, revealing no statistically significant differences in gender, age, stone diameter, disease duration, BMI, site, number of stones, stone location, degree of hydronephrosis, and blood creatinine levels (P>0.05), as shown in **Table 1**.

Comparison of surgical outcomes

Comparing the stone removal and residual rates between the two groups, the research group had a residual rate of 14.81%, signifi-

cantly lower than the control group's residual rate of 29.23% (P<0.05), as shown in Table 2.

Comparison of surgical and postoperative recovery

Comparing the operation and postoperative recovery metrics between the two

groups, the research group had a significantly longer operation time than the control group (P<0.05). Besides, the research group experienced significantly lower intraoperative bleeding and shorter hospitalization times compared to the control group (P<0.05). There was no statistical difference between the groups regarding the time to ambulation (P>0.05), as illustrated in **Figure 2**.

Analysis of optimal cutoff value for the measures

Since logistic regression requires dichotomous information analysis, we used the ROC curve to determine the optimal cutoff values for age, stone diameter, disease duration, BMI, and blood creatinine. These measures were classi-

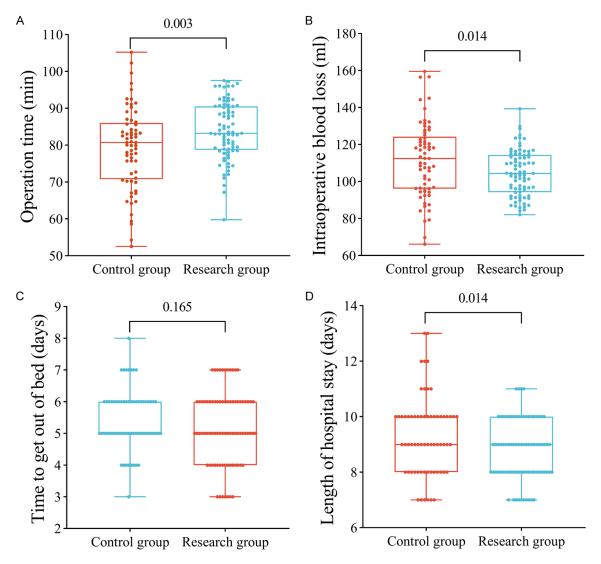


Figure 2. Comparison of surgical and postoperative recovery. A. The operation time was significantly higher in the research group than that in the control group. B. Intraoperative bleeding was significantly lower in the research group than that in the control group. C. The comparison of time to ambulation between the two groups was not statistically significant. D. The length of hospital stay was significantly lower in the research group. ns indicates P<0.05, * indicates P<0.05, and ** indicates P<0.01.

fied as dichotomous information based on the optimal cutoff values. The area under the curve (AUC) was 0.501 for age, 0.619 for stone diameter, 0.589 for disease duration, 0.571 for BMI, and 0.534 for blood creatinine. The optimal cutoff values were 41.5 years, 25.835 mm, 13.5 months, 21.705 kg/m², and 50.77 μ mol/L, respectively, as shown in **Figure 3** and **Table 3**.

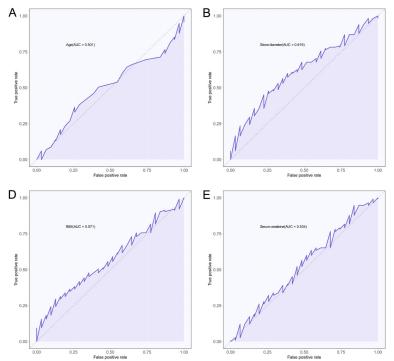
Analysis of single factors affecting stone retention

Univariate analysis of 31 patients with residual stones and 115 patients without residual

stones showed that stone diameter, number of stones, stone location, and treatment method were significantly associated with residual stones (P<0.05). In contrast, gender, age, disease duration, BMI, stone site, and degree of hydronephrosis were not significantly associated with residual stones (all P>0.05), as shown in **Figure 4**.

Multifactorial analysis

Multifactorial logistic regression analysis revealed that having multiple stones (OR=3.581, P=0.013) was an independent risk factor for residual stones. Conversely, stone location out-



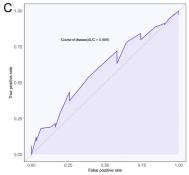


Figure 3. ROC curves of metrological data in predicting stone retention. A. ROC curve for age in predicting stone retention. B. Stone diameter in the ROC curve is used to predict stone retention. C. Disease duration in predicting ROC curves for stone retention. D. ROC curve of BMI in predicting stone retention. E. ROC curve of blood creatinine in predicting stone retention. ROC: Receiver Operating Characteristic, BMI: Body mass index.

Table 3. ROC curve results

	AUC	95% CI	Specificity	Sensitivity	Youden index	Cut off
(A person's) age	0.501	0.393-0.609	74.19%	36.52%	10.72%	41.5
Stone diameter	0.619	0.513-0.724	77.42%	45.22%	22.64%	25.835
Course of disease	0.589	0.479-0.699	74.19%	43.48%	17.67%	13.5
BMI	0.571	0.464-0.678	87.10%	29.57%	16.66%	21.705
Creatinine	0.534	0.415-0.652	16.13%	94.78%	10.91%	50.77

BMI: Body mass index.

side the inferior calyces (OR=0.305, P=0.021) and treatment with a combination of cystoscopy and ureteroscopy (OR=0.398, P=0.042) were independent protective factors against residual stones, as shown in **Figure 5**.

Predictive value of independent influences on stone retention

The predictive value of stone number, stone location, and treatment modality for stone retention was evaluated by plotting ROC curves. The AUC for stone number, stone location, and treatment modality were 0.647, 0.642, and 0.606, respectively, indicating modest predictive value, as shown in **Figure 6** and **Table 4**.

Discussion

Recent advances in medical technology have enabled the development of minimally invasive

strategies for managing kidney stones, generally yielding favorable outcomes [17, 18]. However, complex renal stones still pose significant challenges due to their size, the number of affected calyces, and often associated renal anatomical and functional abnormalities [19]. Key factors in managing these complex kidney stones include enhancing stone removal efficiency, reducing surgical trauma, and facilitating patient recovery [20].

This study compared the surgical outcomes of two groups and found that the stone clearance rate using ureteroscopy alone for complex renal stones was 70.77% [21]. However, when combined with cystoscopy, the clearance rate increased significantly to 85.19%. This improvement in stone clearance may be because complex renal stones are often large and irregularly shaped, making them difficult to remove entire-

Analyzing the effect of soft bladder microscopy combined with soft ureteroscopy

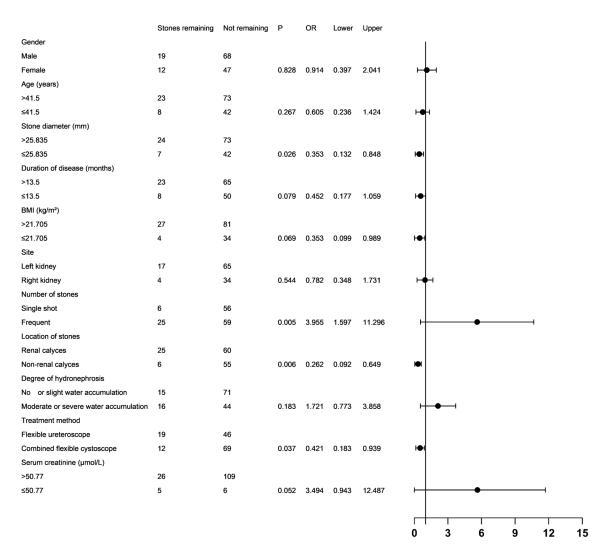


Figure 4. Forest plot of the results of the single-factor analysis. BMI: Body mass index.

ly with ureteroscopy alone [22]. Takazawa et al. [23] suggested that ureteroscopes are more suitable for stones smaller than 20 mm and that staged procedures should be considered for larger stones. Lv et al. [24] found that when the stone's long diameter is greater than 25 mm, the effectiveness of ureteroscopy diminishes. The combination of cystoscopy and ureteroscopy allows for a more comprehensive evaluation of stone location, size, and the presence of multiple stones. Cystoscopy provides direct visualization of the bladder, while ureteroscopy accesses the upper urinary tract. Bladder endoscopes can assist ureteroscopes in reaching difficult stone locations, such as the lower pole calyces, thereby improving the efficiency of lithotripsy and reducing the likelihood of residual stones postoperatively [25].

After comparing the surgery and postoperative recovery of the two patient groups, it was found that using a combination of ureteroscopy and cystoscopy led to significantly longer operation times than using ureteroscopy alone. However, the research group experienced significantly reduced intraoperative bleeding and hospital stays. The time to ambulation post-surgery was not significantly different between the two groups. Pan et al. [26] mentioned that combined treatment not only needs to consider the patient's stone clearance rate but also the impact on the operation time. Higher stone fragmentation efficiency can reduce the operation time and significantly lower surgical risks. The increased number of surgical steps due to using two endoscopes may have contributed to the longer operation times. Still, the operation

Analyzing the effect of soft bladder microscopy combined with soft ureteroscopy

	Stones remaining	Not remaining	Р	OR	Lower	Upper			
Stone diameter (mm)							I		
>25.835	24	73							
≤25.835	7	42	0.060	0.387	0.135	1.002	₩		
Number of stones									
Single shot	6	56							
Frequent	25	59	0.013	3.581	1.373	10.647	+	•	
Location of stones									
Renal calyces	25	60							
Non-renal calyces	6	55	0.021	0.305	0.102	0.799	H		
Treatment method									
Flexible ureteroscope	19	46							
Combined flexible cystoscope	12	69	0.042	0.398	0.160	0.954	H		
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Figure 5. Forest plot of the results of the multifactor analysis.

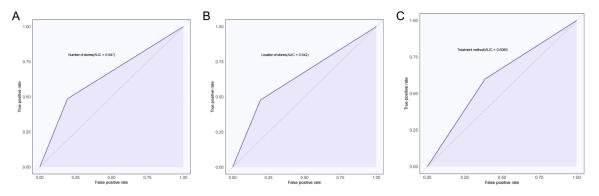


Figure 6. ROC curves for independent influences to predict stone retention. A. Number of stones predicts the ROC curve for stone retention. B. Location of stones predicts ROC curves for stone retention. C. Treatment method predicts the ROC curve for stone retention. ROC: Receiver Operating Characteristic.

Table 4. ROC curve results

	AUC	95% CI	Specificity	Sensitivity	Youden index	Cut off
Number of stones	0.647	0.562-0.731	80.65%	48.70%	29.34%	0.5
Location of stones	0.642	0.558-0.727	80.65%	47.83%	28.47%	0.5
Treatment method	0.606	0.508-0.705	61.29%	60.00%	21.29%	0.5

AUC: area under the curve.

duration remained within two hours, which did not increase patient surgical risk. Despite the longer surgical time, the combined use of the two flexible endoscopes can minimize intraoperative bleeding due to their minimally invasive nature, which reduces tissue damage [27]. Additionally, more precise endoscopic maneuvering and improved visualization mitigate the risk of complications, thereby shortening hospital stays. Zhao et al. [28] noted that combined cystoscopy minimizes postoperative renal impairment. Yang et al. [29] combined cystoscope lithotripsy with the treatment of patients with ureteropelvic junction obstruction and calyceal stones. They mentioned that when the ureteropelvic angle is very small or the patient has severe hydronephrosis, the cystoscope can more easily reach the lower renal calyx, improve the efficiency of stone removal, avoid the risk of secondary surgery, and reduce the patient's bleeding risk and medical economic burden.

In the involved 146 patients, 31 had residual stones postoperatively. To better understand the causes of residual stones, it is essential to identify suitable patients preoperatively to enhance treatment success. Univariate and multivariate logistic regression analyses showed that multiple stones were an independent risk factor for stone retention. In contrast, stone locations in non-lower calyces and the treatment modality of cystoscopy combined with ureteroscopy were independent protective factors against stone retention.

Multiple stones indicate the presence of more than one stone in different parts of the kidney, including regions that are challenging to visualize or access, increasing the risk of residual stones [30]. Stones in non-lower calyceal regions are more accessible to locate and manage endoscopically. In contrast, anatomical limitations make stones in the lower calyces more challenging to remove [31]. Stones in non-lower calyceal regions are more accessible through ureteroscopes and cystoscopes, reducing the likelihood of residual stones.

The combination of cystoscopy and ureteroscopy provides a comprehensive view, enabling better stone localization and improving stone removal rates. Cystoscopy directly visualizes the bladder and upper urinary tract, while ureteroscopy accesses individual kidney calyces. The combined use of both scopes provides a more thorough assessment. The study evaluated the predictive value of these three independent factors for postoperative stone retention, with the AUC of the ROC curves being 0.647, 0.642, and 0.606, respectively. This suggests that identifying the number of stones. their location, and the treatment modality can more effectively predict and reduce the risk of stone retention.

This study has several limitations. The data collection period was relatively short, and the overall sample size was small, potentially leading to biased results due to subjectivity and selection bias in case selection. Additionally, intraoperative bleeding was influenced by factors such as fluid infusion, leading to potential bias in the results. The stone clearance rate was evaluated 2-3 days postoperatively, but this assessment could be influenced by factors such as intestinal gas interference and stone accumulation. Current studies suggest assessing surgical outcomes and stone retention 1-3 months postoperatively, but this is challenging due to patient loss to follow-up or incomplete case data [32, 33]. Additionally, the cost of flexible cystoscopes is relatively high [34]. Many primary hospitals in some developing countries do not have access to flexible cystoscopes, so lower-cost treatment plans should be developed in the future.

In conclusion, the application of cystoscopy combined with ureteroscopy in treating patients with complex renal stones improves the stone clearance rate and postoperative recovery, with a more significant effect compared to ureteroscopy alone.

Disclosure of conflict of interest

None.

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