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**UROLOGY** 

# Beware the bolus size: understanding intrarenal pressure during ureteroscopic fluid administration

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**Background:** Flexible ureteroscopy (FURS) and laser lithotripsy for ureteric and renal calculi requires adequate irrigation for visualisation. This study aimed to evaluate how bolus administration of irrigant fluid impacts intrarenal pressure (IRP) during FURS. We also investigated how ureteral access sheaths (UAS) of varying sizes mitigate elevated IRP.

**Methods:** Using a porcine cadaveric model, IRP was evaluated using an arterial invasive pressure measurement system. Given a fluid column height (driving force) of 80 cm  $H_2O$ , and varying bolus administration (1, 2, 3, 5, 10 ml), IRP was studied with and without a UAS. An IRP of < 40 mmHg was considered the cut off for "safe" FURS. The flow (drainage capacity) of UAS was also evaluated. At varying fluid column heights, three sizes of UAS were used, 10/12 French size (Fr), 11/13 Fr and 12/14 Fr, all 36 cm long.

**Results:** Bolus administration with a UAS of < 5 ml with a starting fluid column height of 80 cm H<sub>2</sub>O was "safe" (< 40 mmHg). In contrast, where no UAS was used, bolus sizes as small as 2 ml produced "unsafe" peak pressures. The flow through a 10/12 Fr UAS was poor but improved greatly with larger UAS diameters.

**Conclusion:** This study suggests that 10/12 Fr UAS may be inadequate to maintain drainage from the kidney at acceptable pressures. Bolus fluid administration produces "unsafe" (> 40 mmHg) elevated IRP in the absence of a UAS. When a UAS is used, a fluid bolus of < 5 ml is likely "safe".

Keywords: renal stones, ureteroscopy, complications, basic science, access sheaths, intrarenal pressure

### Introduction

Nephrolithiasis (kidney stones) is one of the most common diseases in modern times, affecting one in ten people in the United States in their lifetime. Recurrence rates are high, and prevalence worldwide is increasing due to diet and lifestyle changes.<sup>1</sup>

Flexible ureteroscopy (FURS) and laser lithotripsy is a commonly performed urological procedure for ureteric and renal calculi. The interest in collateral damage during such endourological interventions is increasing. One aspect that has recently received more attention is the risk of complications associated with elevated intrarenal pressure (IRP).

Visibility is key to successful FURS. Given the small size of flexible ureterorenoscopes (typically in the range of 8–10 Fr diameter), obtaining adequate irrigation fluid flow is not always easy, especially when vision is hampered by bleeding, pus and debris in the collecting system. In order to improve visualisation, irrigant flow must be increased. Various devices such as syringes or foot pedals have been proposed to improve irrigant flow. These devices, while increasing flow, may contribute to elevated IRP.

Normal IRP is 0–15 mmHg.<sup>2</sup> Hinman and Redewill showed that reflux of urine via pyelovenous reflux occurs above IRP of 40 mmHg.<sup>3</sup> Elevated IRP has been linked to postoperative infectious complications.<sup>2</sup>

Ureteral access sheaths (UAS) are commonly employed during FURS. UAS and low flow rate irrigation has been demonstrated by Wilson and Preminger to reduce IRP.<sup>4</sup>

The aim of the study was to evaluate IRP during varying irrigant flow from typical clinical driving pressures of fluid, to investigate the impact of varying bolus size administration on IRP and in a parallel set of experiments to investigate the ability of a UAS to mitigate IRP and provide effective drainage.

### Methods

IRP measurements were taken in response to varying fluid irrigation heights and bolus volumes. Fresh porcine cadaveric kidneys were used. The animals were not sacrificed for the purpose of the study. The porcine kidneys with preserved ureter were less than six hours post-mortem. Three porcine kidneys were used. For each experiment, the average IRP from the two kidneys was calculated.

IRP measurements were taken directly by means of 20-gauge cannula inserted to the renal pelvis (Figure 1) and transduced using a Deltran Invasive Blood Pressure Transducer (Utah Medical Incorporation, USA). To measure the IRP, an arterial invasive pressure measurement system, GE Datex – Ohmeda Cardiocap 5 was used and was zeroed to atmospheric pressure.



Figure 1: A 20-gauge cannula was placed into the renal pelvis to measure IRP using an arterial invasive pressure measurement system

Water for irrigation boluses of 1, 2, 3, 5 and 10 ml were administered into the cadaveric ureter through a FURS (Pusen Uscope digital ureteroscope, Zhuhai Pusen Medical Technology Co., Ltd.) with a diameter of 3.1 mm (as measured by Mitutoyo Dial Callipers # 505-681, Mitutoyo, Japan) and a tip to Luer Lock length of 72 cm.

An irrigation set (standard Cysto-irrigation set – total length 220 cm) was attached to the irrigation port at a fluid column height of 0 and 80 cm. This was done with and

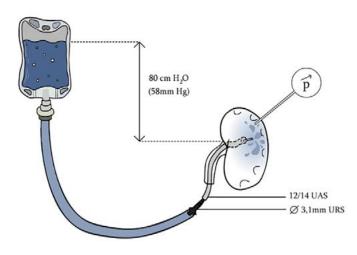


Figure 2: The IRP is assessed with varying driving pressure (80 cm  $H_2O$  is shown here) and with (as shown here) or without a UAS

without a UAS. The UAS was 36 cm long with a diameter of 12/14 Fr (inner diameter 12 Fr, outer diameter 14 Fr, Proxis, BARD/BD). The tip of the UAS was placed in the pelvis of the kidney.

The artificial choice of a column height 0 cm was motivated by wanting to assess the effect of fluid bolus administration in isolation, while a fluid column of 80 cm of  $H_20$  was used as 60–80 cm  $H_20$  is considered as the conventional height in clinical practice.

In the first experiment, starting pressure was measured before the bolus administration and the highest (peak pressure) following the bolus administration was recorded. The bolus was administered with a syringe by hand using a chronometer at a rate of 1 ml per second.

Gravitational hydrostatic pressure was recorded at a fluid column height of 0 cm and 80 cm of H<sub>2</sub>O (58 mmHg) with and without a UAS and with increasing volumes of bolus fluid administered (1, 2, 3, 5, 10 ml) (Figure 2).

Based on the work of Hinman and Redewill,<sup>3</sup> an IRP of > 40 mmHg was considered the cut off for pyelovenous reflux and potentially "unsafe".

In the second experiment, assessment was made of drainage capacity of UAS at varying fluid column height (driving pressure) while the UAS was in a straight position and bent/curved. Three sizes of UAS were used 10/12 Fr, 11/13 Fr and 12/14 Fr, all 36 cm long. Partial obstruction

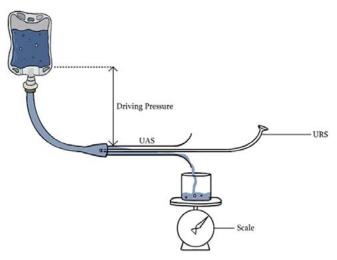


Figure 3: Flow (ml/minute and ml/hour) were assessed with varying diameters of UAS at a fixed 80 mm  $H_2O$  driving pressure



Figure 4: Straight and curved (illustrated here) UAS were used to understand if the curved UAS impacted UAS drainage

was caused by the 3.1 mm diameter Pusen FURS (Figure 3). The minute flow was measured, and the hourly flow was calculated.

Lastly, the second experiment was repeated, but this time measurements were taken through straight UAS and then through curved UAS (Figure 4).

# Results

At a driving force of 0 cm  $\rm H_2O$ , only a bolus greater than 3 ml produced pressures above the predetermined "safe range" of < 40 mmHg. By comparison with the more typical clinical scenario, with a driving pressure of 80 cm  $\rm H_2O$ , any bolus of > 1 ml produced "unsafe" pressures (Table I).

In comparison, when a UAS was used, at a driving pressure of 0 cm  $\rm H_2O$  only a bolus of 10 ml resulted in pressures above 40 mmHg. However, if the driving pressure was 80 cm  $\rm H_2O$ , a bolus of 5 ml and above caused "unsafe" pressures (Table II).

In terms of drainage flow rate through various sizes of UAS at varying driving pressures, we found no appreciable difference between curved and straight UAS (Table III).

Flow through the 10/12 UAS was very poor (24 ml/hour) compared to 12/14 UAS (4 200 ml/hour) (Table IV).

### **Discussion**

The normal IRP is in the range of 0–15 mmHg (0–20 cm H<sub>2</sub>O).<sup>5</sup> Pyelovenous reflux has historically been quoted to occur above 40 mmHg (54 cm H<sub>2</sub>O),<sup>3</sup> while more recent studies suggest it can occur at lower level of 20 mmHg (27 cm H<sub>2</sub>O).<sup>6</sup> FURS is known to increase IRP to 60–100 mmHg in the absence of a UAS. If irrigation is forced, pressure rises above 300 mmHg are not unusual.<sup>2</sup> In a recent study, Jung and Osther demonstrated *in vivo* that during routine FURS in 12 patients for stones, the IRP averaged 54 mmHg and pelvic pressure peaks up to 328 mmHg occurred. In a 5-minute standardised period of simple FURS, 83 pressure peaks > 50 mmHg were measured.<sup>7</sup>

These elevated IRPs have been linked most importantly to infection, but also to haemorrhage, renal impairment and postoperative pain.<sup>2</sup> Pyelotubular reflux occurs

Table I: Peak IRP without UAS at varying bolus volumes with driving pressure of  $\theta c$  m  $H_2O$  (left) and 80 cm  $H_2O$  (right)

Bolus volume (ml)	Starting IRP (mmHg)	Peak IRP (mmHg)
1	5	8
2	4	19
3	3	44
5	5	115
10	5	148

Table II: Peak IRP with UAS (12/14 Fr) at varying bolus volumes with driving pressure of 0 cm  $\rm H_2O$  (left) and 80 cm  $\rm H_2O$  (right)

Bolus volume (ml)	Starting IRP (mmHg)	Peak IRP (mmHg)
1	1	1
2	1	1
3	1	4
5	1	14
10	1	47

Bolus volume (ml)	Starting IRP (mmHg)	Peak IRP (mmHg)
1	7	7
2	7	11
3	8	12
5	7	56
10	7	290

Bolus volume (ml)	Starting IRP (mmHg)	Peak IRP (mmHg)
1	35	36
2	36	78
3	34	116
5	37	116
10	35	320

Table III: Minute flow rate (ml/minute) through various sizes of UAS at varying driving pressures			
Size of UAS (in French ID/OD)	Minute flow at 20 mmHg (27 cm H <sub>2</sub> O) driving pressure	Minute flow at 35 mmHg (47 cm H <sub>2</sub> O) driving pressure	Minute flow at 60 mmHg (82 cm H <sub>2</sub> O) driving pressure
10/12 straight	5	6	6
10/12 curved	4	5	6
11/13 straight	48	62	94
11/13 curved	40	61	94
12/14 straight	71	124	191
12/14 curved	67	105	186

Table IV: Calculated hourly flow rate (ml/hour) though various sizes of UAS at varying driving pressures

Size of UAS (in French)	Calculated hourly flow at 20 mmHg (27 cm H <sub>2</sub> O) driving pressure (ml/hour)	Calculated hourly flow at 35 mmHg (47 cm H <sub>2</sub> O) driving pressure (ml/hour)	Calculated hourly flow at 60 mmHg (82 cm H <sub>2</sub> O) driving pressure (ml/hour)
10/12	24	30	36
11/13	2 400	3 660	5 640
13/14	4 200	6 300	11 160

at 20-30 mmHg and pyelovenous reflux occurs at 30-50 mmHg.<sup>2,3</sup>

Zhong et al. studied these infective complications demonstrating in a series of 260 patients undergoing FURS that a systemic inflammatory response (SIRS) was seen in 8.1%. Independent risk factors included: stone size, small-calibre UAS, irrigation flow rate and struvite calculi.<sup>8</sup> The large multicentre Endourology Society (CROES) FURS study reported infection postoperatively in 2.97% with severe sepsis seen in 0.3%.<sup>9</sup>

The use of UAS has been shown by Traxer et al. to decrease postoperative sepsis rates by 50%. 10 Other than UAS, the factors confirmed to impact IRP include: the access sheath size, the ureteroscope diameter and the irrigation rate. 2

Strategies which reduce IRP and promote drainage should impact the incidence of infective complications and promote patient safety, particularly in those patients with infected struvite stones. Rehman et al. suggested that larger sized UAS lead to improved irrigation flow and lower IRPs in a human cadaveric model. Similarly, Wright et al. concluded statistically that a 12/14 Fr access sheath offered significantly improved irrigation whilst maintaining significantly lower IRP, when compared to a 10/12 Fr access sheath in a cadaveric porcine model.

Another strategy proposed by Ng et al. noted substantially poorer flow with the irrigation channel occupied by a guidewire or basket. The proposed placing of the guidewire "inside or alongside the UAS provides by far the highest flow rates without a rise in IRP".<sup>13</sup> Likewise, Wright et al. counterintuitively have shown that different laser fibres of apparently the same diameter reduce the flow to quite varying degrees.<sup>14</sup>

More recently, Fang et al. highlighted the importance of the ratio of endoscope-sheath diameter (RESD). They showed that to maintain a low intrapelvic pressure (IPP) and acceptable flow rate during FURS laser lithotripsy, RESD should be kept below 0.75. UAS length had less effect on the IPP and flow rate. This study goes on to note that certain manufacturers' ureteroscopes may be better suited to certain commercially available UAS in terms of IRP and flow characteristics. This work builds on an earlier paper by Monga et al. 16

Our study presents three major findings. Firstly, we have demonstrated that UAS not only reduces IRP in the setting of continuous irrigation but also with bolus fluid administration. Bolus administration of < 5 ml with a starting fluid height (driving force) of 80 cm H<sub>2</sub>O is "safe". This contrasts with the situations where no UAS is used here bolus volumes as little as 2 ml produced "unsafe" (> 40 mmHg) peak pressures. Secondly, flow through different sized UAS varies dramatically. The flow through a 10/12 Fr UAS is far poorer than through a larger UAS. This dramatic difference in flow in a larger UAS is a demonstration of Hagen-Poiseuille's law.<sup>17</sup> The law shows that as tube radius halves, flow reduces by 16-fold (i.e. flow is proportional to radius of the tube to the power of 4). This finding would appear to contradict Sener et al. whose findings showed in vitro that FURS via a 10/12 Fr UAS provided pressures below 30 mmHg.<sup>18</sup> Thirdly, despite good minute drainage capacity of the two larger UAS (Fr 11/13 and Fr 12/14), this ability of good flow is in sharp contrast to its limited ability to reduce IRP during bolus administration. The relationship between flow and IRP reduction requires further elucidation

Despite the increased flow of larger calibre UASs, it should be mentioned that their use is not without its own set of complications. Traxer and Thomas showed that marked ureteral wall injury is common after the insertion of a 12/14 Fr UAS and that this risk benefit should be born in mind when using large UASs.<sup>19</sup>

Our study has limitations. The work was done *in vitro* in porcine cadaveric kidneys. This limits its applicability to live human subjects. The lack of peristalsis from a living animal and the pressure from the surrounding tissues may make the results less useful in a clinical setting. It would be useful to attempt to reproduce these findings *in vivo*. We limited the study to a single set of devices which limits it applicability to other manufactured products.

### Conclusion

Visibility is key to successful FURS. Infective and other complications are under-recognised problems with FURS. Keeping IRP low while promoting flow appears to be critical in reducing these complications. This study suggests that 10/12 UAS may be inadequate to maintain drainage from the kidney. Bolus fluid administration produces "unsafe" (> 40 mmHg) elevated IRP in the absence of a UAS. When a UAS is used, fluid bolus < 5 ml is likely "safe". Further work is warranted to investigate how elevated IRP can better be mitigated.

# Conflict of interest

The authors declare no conflict of interest.

### Funding source

None.

# Ethical approval

The research did not involve human participants and/or animals and therefore did not require ethical approval.

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