

Risk Factors of Uterine Contraction After Ureteroscopy in Pregnant Women With Renal Colic

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Abstract

Background: Ureteroscopy is widely applied in pregnant women with renal colic, but such patients are easy to experience uterine contraction after ureteroscopy, There are many factors which may result to uterine contraction, this study aims to explore the risk factors of uterine contraction triggered by ureteroscopy in pregnant women with renal colic.

Methods: 101 pregnant women were retrospectively analyzed, all patients were hospitalized because of severe renal colic. All patients received ureteroscopy during which double J catheters were inserted into ureters for drainage. Patients received medical treatment individually according to their condition and the uterine contractions were detected by EHG within 12 hours after operation. Patients were classified as group A (uterine contraction, n=46) and group B (no uterine contraction, n=55) according to the presence or absence of continuously regular uterine contraction. Clinical characteristics were collected for further analysis, including history of deliveries, anesthesia methods, application of phloroglucinol or not, surgical time, Oxygen inhalation or not, pain relief or not after surgery, systemic inflammatory response syndrome(SIRS) occurred or not. A binary logistic regression analysis model was established to explore whether such clinical characteristics were relevant to uterine contraction after ureteroscopy.

Results: The incidence of continuously regular uterine contraction occurred within 12 hours after ureteroscopy in pregnant women was 45.54%, and the presence of uterine contraction was related to the following factors($P < 0.05$): history of deliveries(primipara versus multipara)(OR=6.593, 95%CI 2.231-19.490), surgical time(each quarter additional)(OR=2.385, 95%CI 1.342-4.238), application of phloroglucinol(yes versus not)(OR=6.959, 95%CI 1.416-34.194), pain relief after surgery(yes versus not)(OR=6.707, 95%CI 1.978-22.738), SIRS occurred after surgery(yes versus not)(OR=0.099, 95%CI 0.014-0.713).

Conclusion: Continuously regular uterine contraction is easy to occur within 12 hours after ureteroscopy in pregnant women. SIRS occurred after surgery is a risk factor for uterine contraction; on the contrary, no history of childbirth, shorter surgical time, application of phloroglucinol, pain relief after surgery are protective factors.

Background

Renal colic is one of the most common non obstetric pain and pregnancy emergencies in pregnant women^[1], and its incidence is between 1:200 and 1:2000, which could lead to severe pregnancy complications, such as abortion or premature delivery^[2]. The main cause of renal colic in pregnant women contains urinary calculi, in addition to ureteral stricture, fetal ureteral obstruction and so on. Symptomatic medications, double-J tube drainage or percutaneous nephrostomy are considered to be effective methods for renal colic^[3]. Due to the limitation of drug application for pregnant women and the trauma caused by nephrostomy^[4, 5], thereby ureteroscopy is widely considered to be a better choice. Uncontrollable pain and gradually aggravating ureteral obstruction are surgical indications of

ureteroscopy^[6, 7]. Complications such as urinary tract infection and so on are common after ureteroscopy, but regular uterine contraction is one of the most severe complications^[8, 9]. Persistent and intense uterine contraction, which could lead to cervical dilatation, is a sign of threatened preterm birth^[10], even if the length of cervical canal changes little^[11], the premature birth rate of pregnant women with renal colic could be as high as 12.8% according to previous literatures^[12]. Therefore, preventing the occurrence of continuously regular uterine contraction is the premise to avoid adverse pregnancy events. The occurrence of uterine contraction is interactive result of multiple factors^[13]. However, there is little literature about risk factors of uterine contraction caused by ureteroscopy. In this study, we retrospectively analyzed 101 pregnant women who were received ureteroscopy to explore the risk factors of uterine contraction after such surgery.

Methods

101 pregnant women who received ureteroscopy in our hospital from Jan 2010 to Mar 2020 were retrospectively analyzed, including 46 primiparas and 55 multipara. The patients, at 20-36 weeks of gestation, were admitted because of severe renal colic. The diseases included ureteral calculi, renal calculus, ureteral stricture and ureteral obstruction caused by fetal oppression. The cervical was not opened and there was no continuously regular uterine contraction detected by EHG before ureteroscopy. If symptomatic treatment did not work well and the renal colic had little relief, then patients would receive ureteroscopy and double-J catheters drainage on the basis of informed consent. The anesthesia methods included general anesthesia(13 cases), lumbar anesthesia(39 cases) and spinal-epidural anesthesia(49 cases). After operation, EHG was used again to monitor continuously regular uterine contraction for 12 hours. Phloroglucinol was routinely applied in 42 patients; Oxygen inhalation was given to 61 patients; Renal colic was not completely relieved in 26 patients immediately after surgery but after several hours; Systemic inflammatory response syndrome (SIRS) occurred in 11 patients but nobody progressed to sepsis. According to the occurrence of continuously regular uterine contraction after operation, the patients were divided into group A and group B. Group A (n=46) presented regular contractions within 12 hours after surgery and received magnesium sulfate or ritodrine hydrochloride, while group B (n=55) did not present regular contractions within 12 hours after ureteroscopy. Patients' differences in age, weight and Gestational Week between the two groups were not statistically significant ($P > 0.05$) (Tab 1).

Diagnosis and treatment criteria were as follows: The uterine contraction were monitored continuously after admission until 12 hours after operation. Any uterine contraction lasted for more than 30 seconds with a interval less than 15 minutes would be considered as effective uterine contraction^[14]. If the length of cervical canal was found to become shorter, 25% magnesium sulfate or 0.01% ritodrine hydrochloride were given immediately to inhibit uterine contraction. In view of the side effects of uterine contraction inhibitors, the following indicators were needed to monitor closely during application of contraction inhibitors. (1) Breath frequency no less than 16 times / min; (2) heart rate no more than 120 times / min; (3) knee reflex existed; (4) urine volume not less than 17 ml / h^[15]. The diagnostic criteria of systemic inflammatory response syndrome were as follows: (1) body temperature $> 38.0^{\circ}\text{C}$ or body temperature $<$

36.0°C; (2) heart rate > 90 beats / min; (3) respiration rate > 20 breaths / min or excessive breathing with PaCO₂ < 32mmHg; (4) blood WBC > 12.0×10⁹ / L, or WBC < 4.0×10⁹ / L or immature WBC > 10.0%^[16].

Statistics: in addition to age, weight and gestational weeks, the other clinical characteristics of patients in the two groups were collected, including delivery history, anesthesia method, surgery time (each quarter additional), phloroglucinol application or not, oxygen inhalation or not, renal colic relief or not, and SIRS occurrence or not. The above clinical features were analyzed to explore whether they were related to the presence of uterine contraction (Table 2).

All statistical analyses were performed by SPSS version 23.0 (IBM, New York, NY). Measurement data were showed as mean±standard deviation ($\bar{x}\pm s$), rate or percentage were compared by chi-square test. A binary logistic regression model was established to explore the clinical characteristics which might be related to uterine contraction, adjusted odds ratio (OR) and 95% confidence interval (CI) were analyzed. The P value less than 0.05 was considered to be statistically significant.

Result

In all 101 patients, continuously regular uterine contractions which could be inhibited by magnesium sulfate or ritodrine occurred in 46 patients within 12 hours after operation. No abortion or premature delivery occurred in any patient. Binary logistic regression analysis showed that delivery history, surgery time, phloroglucinol application, relief of renal colic and SIRS occurrence were related to the presence of uterine contraction ($P < 0.05$). According to the statistical results, presence of SIRS after surgery (OR=0.099, 95% CI 0.014-0.713) was the risk factor for uterine contraction, meanwhile, no history of labor (OR=6.593, 95% CI 2.231-19.490), shorter surgery time(each quarter additional) (OR=2.385, 95% CI 1.342-4.238), phloroglucinol application (OR=6.959, 95% CI 1.416-34.194), complete relief of renal colic(OR=6.707, 95% CI 1.978-22.738) were protective factors for uterine contraction (Tab 3).

Discussion

At present, the definition of preterm birth is still controversial. Flenady et al^[17]. pointed out that any childbirth within gestational age between 20 and 36 weeks could be defined as Preterm birth which was the main contributor of perinatal morbidity and the second leading cause of neonatal death in the world. The most common clinical symptom of urolithiasis during pregnancy was renal colic and mainly was observed after 20 weeks of gestation^[18]. In consideration of ritodrine hydrochloride's indications (only for pregnant women after 20 gestational weeks) and the difficulty for monitoring uterine contractions at early gestation^[19], thereby we only retrospective analyzed pregnant women at 20 to 36 gestational weeks. Johnson et al^[12]. had shown that the obstetric complications after ureteroscopy were about 4.0%, but in our study there was nothing severe complication except for that 45.5% of patients developed to continuously regular contractions after ureteroscopy. The incidence of uterine contraction in our study was higher than that reported by other scholars^[20]. The possible explanation were as follows: (1) All cases summarized in the study were patients with severe renal colic which could not be relieved by

painkiller in outpatient department. In view of the relevance between pain and uterine contraction, the incidence of such complication rose accordingly. (2) All patients were received uterine contraction monitoring by EHG for 12 hours after surgery, thus any slight contractions could be detected in time and analyzed in the study, but in fact, some uterine contractions in early stage could disappear without any medical treatment. There was not any severe complications such as abortion or premature delivery happened in this study, the results suggest that ureteroscopy was a safe and effective surgical method and routine monitoring of uterine contractions after ureteroscopy might help to avoid pregnancy complications.

Compared with primipara, the cervix of multipara is easier to expand, which means a sooner labor process^[21, 22]. Uterine contraction is stronger for multipara during labor and more analgesics are needed for painless delivery^[23]. Our study showed that multipara were more likely to develop to uterine contraction after ureteroscopy. The possible explanations for this results were as follows: (1) Compared with primipara, the oxytocin receptor which played a key role in uterine contraction had a higher expression on uterus in multipara^[24, 25]; (2) With the increasing of age, the incidence of pregnancy complications rose in step, multipara was more likely to develop to uterine contraction after operation^[26, 27].

Enlarged uterus due to pregnancy inevitably result in ureter compression because of the adjacent location between lower segment ureter and cervix^[28-30]. The operation of ureteroscope would inevitably stimulate the cervix, and repeatedly physical stimulation could trigger uterine contraction. Scholars had shown that pregnancy complications were prone to occur after long-time surgery^[31], and shortening operation time could reduce such complications^[32]. Our results also indicated that operation time was a risk factor for uterine contraction. This study showed that each additional a quarter increase, the adjusted odds ratio was as high as 2.385. This result also reminded that the surgeons should shorten operation time as much as possible when ureteroscopies are performed on pregnant women. The purpose of ureteroscopy was just to resolve urethral obstruction, not for removing stones.

Phloroglucinol, as a non-papaverine antispasmodic, only acts on spastic smooth muscles and has minimal impact on normal smooth muscles. Application of phloroglucinol during pregnancy does not increase the risk of fetal malformations^[33]. It was applied to relieve renal colic and inhibit uterine contractions for decades^[34, 35]. Our research showed that application of phloroglucinol after ureteroscopy could reduce the incidence of uterine contractions. In this study, only a few doctors prescribed phloroglucinol to patients after ureteroscopy. In view of the effectiveness and safety of this medication, we suggested that such drug could be applied to more patients.

If severe pain persists, human body would release prostaglandins which could promote cervical ripening and lead to smooth muscle contraction in uterine^[36, 37]. Pain could also cause stress response and then trigger a increasing release of catecholamines that plays an important role in uterine contractions^[38]. Furthermore, severe renal colic often accompanies with repeated vomit which may lead to uterus

compression. Mechanical stimulation could also induces uterus contractions^[39]. Our research also showed that unrelieved pain after surgery was a risk factor for uterine contractions. About a quarter of the patients in this study had little relieve in renal colic immediately after ureteroscopy. The result was coincide with other scholars^[3]. The possible explanations was that the double J catheter inserted into ureter during operation did not play a effective role in drainage soon after surgery. With painkiller prescribed, pain in these patients gradually subsided in 12 hours after operation.

During the operation of ureteroscopy, continuously intraureteral perfusion by saline is required. Increasing water pressure in ureter would force bacteria flow back into blood via renal pelvic-vein system, it is more likely to happen in patients with urinary tract infection. Bacteremia may result in spread of inflammation or even progress to SIRS which could develop to multiple organ dysfunction syndrome(MODS)^[40]. Inflammatory mediators would be release and spread if SIRS occurs, such mediators could lead to uterus contraction^[41, 42]. Our research also suggested that SIRS was a risk factor for uterus contraction. Pregnant women were more likely to develop to sepsis due to immunosuppression and down regulation of inflammatory response because of immunotolerance in pregnancy^[43]. pregnant women with SIRS or sepsis may easily be ignored because of un conspicuous fever and quick heart rate caused by ritodrine hydrochloride^[44], therefore, it is necessary to closely monitor the infection indicators, such as white blood cells, CRP, and PCT-Q and so on, in order to find potential infection in time.

Limitations

There are also some limitations in this study. Firstly, this study is a single center retrospective study, and the number of patients collected is not adequate enough; Secondly, there are many factors that may play a role in uterine contraction, such as scar uterus, but It's difficult to collect enough clinical feature by retrospective analysis and we will plan a cohort study in the future; Thirdly, in this study, all patients only received uterine contraction monitoring for 12 hours by EHG after surgery, but some contractions would not occur within 12 hours.

Conclusion

ureteroscopy is considered to be a safe and effective treatment for pregnant women, but the high incidence of uterine contraction after surgery still calls for more attention. In order to reduce the odds of uterine contraction after operation, surgeon are supposed to finish the operation as soon as possible, and prescribe phloroglucinol routinely after operation, and relieve pain and control infection effectively, and pay more attention to multipara.

Abbreviations

RS

Renal Stone; US:Ureteral Stricture; UO:Ureteral Obstruction

Declarations

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Author's contribution

Conceived and designed the Study: SLZ and GQL; Retrospective analyzed data: CJL; Data collection: JRL, YFZ and MJG, Paper Writing: CJL. All authors have read and approved the manuscript.

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Availability of data and materials

The data used and analyzed in this study are available from the corresponding author on reasonable request. The data that support the findings of this study has been upload as attachment in submission.

Ethics approval and consent to participate

The study received approval on May 26, 2020 by Ethics Committee of Foshan Maternal and Child Healthcare Hospital (protocol number FSFY-MEC-2020-022).. Written informed consent for participation was obtained.

Consent for publication

Not applicable.

Competing interest

The authors declare that they have no competing interests.

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Tables

Tab 1. Clinical features of patients in two groups

Group	Age (Y)	Weight (Kg)	Gestational Week(W)	Surgical time (min)	Disease species(Case)			
					Ureterolith	RS	US	UO
A (n=46)	28.0±5.7	58.0±8.3	26.4±4.4	28.8±16.3*	23	5	13	5
B (n=55)	29.8±5.2	61.6±9.6	25.8±4.7	23.8±11.1	39	5	5	6

*P<0.05

Tab 2. Clinical characteristics for logistics analysis

Clinical characteristics	Group A (case)	Group B (case)
History of delivery		
Primipara	30	16
Multipara	16	39
Anesthesia methods		
General Anesthesia	7	6
Lumbar Anesthesia	19	20
Spinal-Epidural Anesthesia	20	29
Surgical time (min)		
≤15	9	17
15≤30	22	25
30≤45	6	11
45≤60	6	2
60≤75	3	0
Application of Phloroglucinol		
Yes	36	49
No	10	6
Oxygen inhalation		
Yes	31	28
No	15	27
Complete relief of renal colic		
Yes	26	49
No	20	6
Occurrence of SIRS		
Yes	9	2
No	37	53

Tab 3. Risk factors for uterine contraction in pregnant women after Ureteroscopy

Risk factors	OR	95% CI	P
History of deliveries			
primipara versus multipara	6.593	2.231-19.490	0.001
Duration of surgery			
Short versus long	2.385	1.342-4.238	0.003
Application of Phloroglucinol			
yes versus no	6.959	1.416–34.194	0.017
Pain relief			
complete versus incomplete	6.707	1.978–22.738	0.002
SIRS			
occurred versus not occurred	0.099	0.014–0.713	0.022