

The Daily Resistive Index measurement useful tool in the estimation of the optimal time interval between two Shock Wave Lithotripsy sessions.

Esin Yencilek¹, Aysegul Sarsilmaz², Ozgur Kilickesmez³, Hakan Koyuncu⁴, Bilal Eryildirim⁵, Bengi Gurses², Yavuz Bastug⁶, Bilger Erihan⁴

¹Department of Radiology, Haydarpasa Numune Education and Research Hospital, ²Department of Radiology, Yeditepe University Medical Faculty, ³Department of Radiology Istanbul Education Hospital, ⁴Department of Urology, Yeditepe University Medical Faculty, ⁵Department of Urology Kartal Lutfi Kırdar Education and Research Hospital, ⁶Department of Urology Beykoz State Hospital, Istanbul, Turkey

Abstract

Objective: To monitor the impact of Shock Wave Lithotripsy (SWL) on the renal resistive index (RI) and to investigate the potential of the RI measurement for the estimation of the optimal duration between 2 SWL sessions. **Material and methods:** Thirty patients with single pelvis renalis stone were included. Participants were grouped according to their age as group 1 (<40 years, mean age 36.2±3.9 years) and group 2 (≥40 years, mean age 55.4±6.5 years). RI measurement was performed in all patients prior to SWL. After SWL, RI was monitored daily until RI returned to their pre-SWL values. **Results:** The mean stone size was 28.97±3.62 in group 1 and 10.08±4.67 mm in group 2 (p=0.077). Following SWL RI value of both groups increased and higher RI value was measured at 24th hour as compared with their pre-SWL values (p<0.001). In day 2 RI of the groups declined, but the differences were still statistically different from their pre-SWL RI values (p<0.001). However, on the third day, RI of group 1 was close to their pre-SWL level (p=0.143). But, in group 2, RI value returned to their pre-SWL limits on day 4 (p=0.229). **Conclusions:** RI measurement gives important data regarding SWL related acute renal trauma and should be used as an US marker for recovery after SWL.

Keywords: color Doppler ultrasound, kidney, resistive index, Extracorporeal Shockwave Lithotripsy.

Introduction

Urolithiasis is one of the most common urological diseases, and extracorporeal shock wave lithotripsy (SWL) is the method of choice in majority of cases [1,2]. Although it rarely leads to serious complications, SWL treatment is generally accepted as safe and has negligible side effects on long-term follow-up.

In urological practice, multiple SWL treatment sessions may be required depending on stone composition and patient-related factors. However, an “off period” (duration of stand-by time) between two SWL sessions is mandatory to protect the kidneys against repetitive shock wave-related trauma because acute adverse effects of SWL on renal morphology and function may occur despite its proven safety and efficacy during long-term follow-up. Therefore, urologists accept that there should be some duration between SWL sessions for kidney recovery. On the other hand, the ideal time interval between two SWL applications is not well known and remains under debate in the field of urology. It is not known exactly when the acute effect of shock waves on renal parenchyma returns to its normal limits. This question should be answered to estimate the ideal “off period” duration between sessions.

Received 02.12.2014 Accepted 10.01.2015

Med Ultrason

2015, Vol. 17, No 2, 175-179

Corresponding author: Esin Yencilek

Haydarpasa Numune Education and Research
Hospital, Department of Radiology
Tibbiye Cad. No: 40 Uskudar, Istanbul, Turkey
Phone: +902165423232, Fax: +902163360565
E-mail: e.yencilek@yahoo.com

Although some clinical experience with limited evidence indicates the feasibility of repeated SWL sessions within 1 day for ureteral stone, the interval required between repeated SWL sessions for renal stone treatment remains unknown [3]. Therefore, there is no consensus about the duration between SWL sessions. The “off period” varies between 5 days and 2 weeks in daily urological practice depending on urologist’s or SWL technician’s preference.

There are numerous radiologic reports concerning the impact of SWL on renal morphology and function [2,4-7]. Some of the studies focusing on the immediate vascular supply and total effective renal plasma flow of the kidneys indicated a transient decrease in these functions after SWL [4-7]. However, none of the radiological tools have been until now used to investigate the interval required between two SWL sessions. Measurement of the renal arterial resistive index (RI) of the kidney is a recently applied successful radiological tool that can reveal the resistance of intrarenal arteries, indirectly demonstrating effective renal blood flow [8-10]. Recently, it was shown that shock waves may decrease renal blood flow, which can lead to transient impairments in renal function after SWL [11].

In the present study, we aimed to evaluate and monitor renal RI changes after high-energy shock wave application in patients with pelvis renalis stones by using color Doppler ultrasonography (CDU), to show the ability of daily RI measurement during post-SWL follow-up, and to find the safe time interval between the first and second SWL sessions.

Material and methods

Adult patients with normal laboratory results (normal liver and kidney function tests, prothrombin time, activated partial thromboplastin time), no urinary tract infection, and a radiologically proven normal urinary system (no congenital anomaly, no obstruction) were enrolled in the study after approval by the local ethics committee and providing written informed consent. Patients older than 18 years of age and having unilateral single pelvis renalis stone smaller than 20mm were included for the study. The diagnosis of calculi was confirmed by non-contrast computed tomography (CT) and contrast enhanced images were taken to evaluate the urinary system anatomy in same CT session. Multiple stones, any kidney lesion other than stone, history of previous stone treatment (SWL or surgery), metabolic disease threatening kidney function such as hypertension or diabetes, obstructive stone, and body mass index > 30 were the exclusion criterias of the study. All participants underwent

a physical exam, urine culture to exclude urinary tract infection, and routine biochemical tests. Then, patients were grouped according to age into group 1 (<40 years) and group 2 (≥40 years).

Shock wave monotherapy was performed by using an electrohydraulic lithotripter under analgesia that was achieved by intramuscular diclofenac sodium prior to SWL therapy. Upper limit for SW received by patients was 3000 at a frequency of 1–1.5 Hz and 18 kV for the first SWL therapy session. The energy was increased in a stepwise manner with a pulse rate of 80/minute to the maximum intensity after 500 SW.

Before the SWL session, the patients were evaluated using renal CDU (General Electric, Lociq 9, Milwaukee, USA) to measure their interlobar or arcuate RI. RI values of the upper, middle, and lower parts of the kidney were measured, and the arithmetic means of the three values were defined as the RI of the kidney. The RI of the contralateral kidney was also assessed using the same protocol. After the SWL therapy, all patients were followed for a daily RI measurement until their RI values returned to their pre-SWL limits. The daily RI values of the groups were compared with each other and their pre-SWL RI values.

Statistical Package for the Social Sciences (SPSS) version 17 was used for the data analysis. Repeated measures analysis of variance, paired t tests, chi square test, and Mann Whitney-U tests were used for the statistical analysis. Values of $p < 0.05$ were considered statistically significant at the 95% confidence level.

Results

Thirty patients were included in the study. Most of the stones in our study were on the left side (12 in the right and 18 in the left kidney). Table I represents the patients’ demographic characteristics according to groups.

Before SWL monotherapy, RI values of the treated and nontreated kidneys were close to each other within the groups ($p = 0.192$ group 1, $p = 0.347$ group 2). In the treated kidneys, the RI value increased significantly after SWL session in both groups compared to their pre-SWL values ($p < 0.001$). Although a slight decrease was seen on day 2 the mean RI values of both groups remained high comparing with their corresponding pre-SWL RI values ($p < 0.001$). However, on day 3, the mean RI value of group 1 was not significantly different from their pre-SWL value ($p = 0.143$), and the difference in 3rd day RI values between the two groups (group 1 and 2) was statistically significant ($p < 0.001$). The mean RI value of group 2 returned to the normal limit (pre-SWL RI) on day 4 ($p = 0.229$), and the difference between group 1 and

Table 1. Demographic characteristics of the Younger (Group 1) and Older (Group 2) groups.

	Group 1	Group 2	p value
Age (years)	36.2±3.9	55.4±6.5	<0.001
Female	3 (27.3%)	6 (31.6%)	0.064
Male	8 (72.7%)	13 (68.4%)	0.071
BMI	26.2±4.2	28.3±3.0	0.085
Kreatinin (mg/dl)	0.97±0.3	1.08±0.7	0.261
BUN	13.2±3.8	15.7±4.7	0.063
Stone size (mm)	8.97±3.62	10.08±4.67	0.077
SWL duration (min)	45.1±5.8	46.9±8.8	0.125
SBP (mmHg)	110±10.5	128±15.2	0.074
DBP (mmHg)	66±4.8	72±9.9	0.088
Number of Shock Waves	2859±100	2921±57	0.095
Total Energy, kW	2.98±0.49	3.09±0.45	0.182

SBP: Systolic Blood Pressure; DBP: Diastolic Blood Pressure; min: Minute; mm: milimeters

Table II. The values of RI in treated and non-treated kidneys before and after SWL session

Groups		pre-SWL	24th hrs	2ndday	3rd day	4th day	p
Treated kidney	Group 1	0.588±0.11	0.679±0.07	0.643±0.11	0.597±0.14	0.592±0.08	<0.001
	Group 2	0.596±0.04	0.683±0.13	0.661±0.11	0.642±0.22	0.601±0.09	<0.001
Non-treated kidney	Group 1	0.581±0.09	0.583±0.41	0.579±0.22	0.592±0.14	0.575±0.32	0.09
	Group 2	0.585±0.05	0.593±0.13	0.591±0.08	0.586±0.16	0.590±0.11	0.12

2 was not significant by day 4 ($p=0.154$). Evaluation of RI in the contralateral nontreated kidneys revealed no significant change in RI values of pre-SWL versus early- and late-term post-SWL values in both groups (table II).

Discussions

Renal stones, one of the most common urological diseases, should be treated by SWL according to urological guidelines. SWL treatment success depends on technical, stone, and patient-related factors, and repeated sessions are frequently needed for stone disintegration. However, an “off period” is suggested between SWL sessions to protect the renal parenchyma against repetitive trauma caused by the shock waves [3]. The underlying reason for this suggestion depends on the fact that SWL causes acute trauma to the treated kidney parenchyma and its vasculature. On the other hand, the optimal duration of an “off period” between SWL sessions is debatable in urology. In present study, we aimed to show the capability of RI slope after SWL therapy to monitor shock wave-related trauma to kidneys with pelvis renalis stones in order to estimate the optimal “off period” time interval by age.

It is well known that shock wave-related high energy can potentially impair renal function. In the literature, the effects of shock waves on the morphology and function of treated kidneys have been subjected to several clinical and experimental studies [2,4-6]. Many were concen-

trated to clarify the short- or long-term effect of the shock waves on the renal parenchyma and the vast majority of the studies showed that the acute effect of SWL on the renal tissues was temporary due to its compensatory mechanism [12]. Today it is accepted that SWL treatment for renal stones is safe and reliable on long-term follow-up.

Although many studies have shown the efficacy and reliability of SWL, the optimal duration of the “off period” between SWL sessions has yet to be determined. This is an unanswered traditional question in urology because estimating the optimal safe time is important to protecting the renal parenchyma against repetitive shock wave-related trauma. Although some clinical experience with low degree of evidence indicates the possibility of repeating SWL session within 1 day for ureteral stones, we found no clinical study or experience concerning this aspect [2]. The degree of impact of shock waves on ureteral and renal tissue has not been largely studied. However, it is expected that renal tissue is more prone to SWL-related damage owing to its highly vascular thick parenchyma compared to that of ureteral tissue. Therefore, it may be logical to think that the “off period” for renal tissues in renal stone lithotripsy should be longer than that of ureteral tissue. The present study was the first to measure daily RI to demonstrate its decreases. Here we tried to monitor the slope of the RI decrease after SWL therapy and to develop an understanding of the time required for RI values of treated kidneys to return to normal limits.

Some radiological tools that were used to measure total effective renal plasma flow of the kidneys indicated a transient decrease in renal function after SWL [4,6,7]. Also, numerous radiological studies have shown the potential usefulness of imaging techniques in the assessment of early renal damage [7,13,14]. Almost all radiological studies in the literature have focused on the limitations of radiologic tools to understand SWL-related changes. Among these tools, CDU is easy to apply, cost effective, and has the potential to determine renal function and kidney microcirculation [8,9,15-17]. CDU was recently used as a noninvasive method to estimate renal function and has been used to obtain functional information after SWL sessions.

Although some authors showed no change in kidney RI, most studies reported increased RI values after SWL sessions [18-20]. The discrepancies between studies might be due to study group differences and study population heterogeneity. In some series, RI values were studied in stone-related obstructed cases but others were measured RI in patients without obstruction. Although there is some controversies about the impact of obstruction on RI value, obstruction may have potential to affect renal vascular resistance and causes renal perfusion changes [21-24]. Also, patients in the different studies were subjected to different numbers of shock waves. This may also help explain differences among studies showing the relationship between RI and SWL in different series. Therefore, here we studied non-obstructed cases to exclude the possible impact of obstruction on renal tissues. Additionally, the number of shock waves used in our series was similar between groups.

Currently, an increase in RI values is expected after SWL therapy, which has been confirmed in many studies, though there are a few exceptions [11,18]. In the present study, we found higher RI values after SWL sessions in patients in both groups, comparable to that reported previously in literature. Increases in RI values were similar between groups as well. In the literature, increases in RI are time-dependent and the acute effect of the shock waves disappears after some time, most likely due to compensatory mechanisms of the kidney [22]. In agreement with the literature, the RI values in our series returned to almost pre-SWL values several days after SWL treatment. High RI values of the treated kidney lasted from a few days up to day 4 after SWL depending on age group.

The relationship between age and increase in RI values is also well studied. Almost all of the results showed that the increase in RI after SWL sessions is not related to age [25]. We also found high RI values immediately after SWL treatment in both groups. However, until

now, no clinical study has examined the RI decrease on a day-to-day basis; in fact, almost all previous studies measured RI over a predetermined time interval, which cannot determine the exact day of RI recovery to normal values (pre-SWL values). Therefore, we monitored the daily RI of each treated kidney and found that the mean RI values returned to their pre-SWL levels on day 3 in the younger group (<40 years) and on day 4 in the older group (>40 years). We do not have data to explain the underlying reason for the differences in RI recovery between age groups. However, in the literature, higher RI values in older versus younger patients were explained by decreased elasticity in the renal tissues of elderly patients [8]. This conclusion seems logical but should be confirmed by preclinical studies.

Limited studies have evaluated the untreated contralateral kidney in response to SW and data shows a debate in this issue. Some authors found no significant changes in RI values of untreated kidneys versus treated kidneys [26]. In contrast, Mitterberger et al found higher RI measurements after SWL in the contralateral kidney [8]. We found no significant increase in RI values of untreated kidneys after SWL. The difference in the literature can be explained by group heterogeneity, different numbers of shock waves, or the use of medication before SWL in the different study groups.

Although our data confirm the ability of RI to monitor renal blood flow and renal parenchymal RI and assess the optimal "off period" between the first and second SWL sessions, this study has some limitations. First, its main limitation is the few subjects included. Second, the quantitative data obtained from the CDU could not be compared with the histological findings; therefore, the exact relationship between RI value and renal tissue resistance or transient functional alteration after SWL is obscure. Lastly, two experienced radiologists measured the RI of the patients due to multicentricity of the study. However, despite the absence of such data, our results demonstrate the ability of RI to estimate the optimal "off period" in patients with renal stones, and that RI decreases with age among patients receiving SWL.

Conclusions

RI gives an idea regarding SWL related acute renal trauma and should be used as an US marker for recovery after SWL and for prediction in estimating the optimal period between two SWL procedures. However, further studies including more patients are required.

Conflict of interest: none

References

- Bozgeyik Z, Kocakoc E, Sonmezgoz F. Diffusion-weighted MR imaging findings of kidneys in patients with early phase of obstruction. *Eur J Radiol* 2009; 70: 138-141.
- Muller MF, Prasad PV, Bimmler D, Kaiser A, Edelman RR. Functional imaging of the kidney by means of measurement of the apparent diffusion coefficient. *Radiology* 1994; 193: 711-715.
- Türk C, Knoll T, Petrik A, Sarica K, Skolarikos A, Straub M, Seitz C. EAU Guidelines on Urolithiasis 2014. Available at http://uroweb.org/wp-content/uploads/22-Urolithiasis_LR.pdf.
- Laissy JP, Menegazzo D, Dumont E, et al. Hemodynamic effect of iodinated high-viscosity contrast medium in the rat kidney. *Invest Radiol* 2000; 35: 647-652.
- Chan JH, Tsui EY, Luk SH, et al. MR diffusion-weighted imaging of kidney: differentiation between hydronephrosis and pyonephrosis. *Clin Imaging* 2001; 25: 110-113.
- Le Bihan D, Delannoy J, Levin RL. Temperature mapping with MR imaging of molecular diffusion: application to hyperthermia. *Radiology* 1989; 171: 853-857.
- Zhang H, Prince MR. Renal MR angiography. *Magn Reson Imaging Clin N Am* 2004; 12: 487-503.
- Aoki Y, Ishitoya S, Okubo K, et al. Changes in resistive index following extracorporeal shock wave lithotripsy. *Int J Urol* 1999; 6: 483-492.
- Nazaroglu H, Akay AF, Bükte Y, Sahin H, Akkus Z, Bilici A. Effects of extracorporeal shock-wave lithotripsy on intrarenal resistive index. *Scand J Urol Nephrol* 2003; 37: 408-412.
- Tatar IG, Teber MA, Ogur T, Kurt A, Hekimoglu B. Real time sonoelastographic evaluation of renal allografts in correlation with clinical prognostic parameters: comparison of linear and convex transducers according to segmental anatomy. *Med Ultrason* 2014; 16: 229-235.
- Knapp R, Frauscher F, Helweg G, et al. Age-related changes in resistive index following extracorporeal shock wave lithotripsy. *J Urol* 1995; 154: 955-958.
- Strohmaier WL, Carl AM, Wilbert DM, Bichler KH. Effects of extracorporeal shock wave lithotripsy on plasma concentrations of endothelin and renin in humans. *J Urol* 1996; 155: 48-51.
- Riccabona M, Ruppert-Kohlmayr A, Ring E, Maier C, Luisuardi L, Riccabona M. Potential impact of pediatric MR urography on the imaging algorithm in patients with a functional single kidney. *AJR Am J Roentgenol* 2004; 183: 795-800.
- Grenier N, Basseau F, Ries M, Tyndal B, Jones R, Moonen C. Functional MRI of the kidney. *Abdom Imaging* 2003; 28: 164-175.
- Kurt S, Tokgöz Ö, Tokgöz H, Voyvoda N. Evaluation of effects of Extracorporeal Shock Wave Lithotripsy on renal vasculature with Doppler ultrasonography. *Med Ultrason* 2013; 15: 273-277.
- Abd Ellah M, Kremser C, Pallwein L, et al. Changes of renal blood flow after ESWL: assessment by ASL MR imaging, contrast enhanced MR imaging, and renal resistive index. *Eur J Radiol* 2010; 76: 124-128.
- Ghadirpour A, Tarzamni MK, Naghavi-Behzad M, Abedi-Azar S, Koushavar H, Nezami N. Renal vascular Doppler ultrasonographic indices and carotid artery intima-media thickness in diabetic nephropathy. *Med Ultrason* 2014; 16: 95-99.
- Beduk Y, Erden I, Gogus O, Sarica K, Aytac S, Karalezli G. Evaluation of renal morphology and vascular function by color flow Doppler sonography immediately after extracorporeal shock wave lithotripsy. *J Endourol* 1993; 7: 457-460.
- Zolfaghari A, Ghadirpour A, Tarzamni MK, Goldust M, Mirabad MR, Nezami N. Renal vascular Doppler resistance after extracorporeal shock wave lithotripsy. *Ren Fail* 2013; 35: 686-690.
- Mitterberger M, Pinggera GM, Neururer R, et al. Multimodal evaluation of renal perfusional changes due to extracorporeal shock wave lithotripsy. *BJU Int* 2008; 101: 731-735.
- Gurel S, Akata D, Gurel K, Ozmen MN, Akhan O. Correlation between the renal resistive index (RI) and nonenhanced computed tomography in acute renal colic: how reliable is the RI in distinguishing obstruction? *J Ultrasound Med* 2006; 25: 1113-1120.
- Opdenakker L, Oyen R, Vervloessem I, et al. Acute obstruction of the renal collecting system: the intrarenal resistive index is a useful yet time dependent parameter for diagnosis. *Eur Radiol* 1998; 8: 1429-1432.
- Bertolotto M, Moro U, Gioulis E, Lodolo C, Lissiani A. Changes of renal resistive index in response to hydration and diuretic administration in normal subjects and in patients with small ureteral stone. *J Ultrasound Med* 1999; 18: 819-825.
- Lee HJ, Kim SH, Jeong YK, Yeun KM. Doppler sonographic resistive index in obstructed kidneys. *J Ultrasound Med* 1996; 15: 613-618.
- Mohseni MG, H Khazaeli M, Aghamir SM, Biniiaz F. Changes in intrarenal resistive index following electromagnetic extracorporeal shock wave lithotripsy. *Urol J* 2007; 4: 217-220.
- Juan YS, Huang CH, Wang CJ, et al. Predictive role of renal resistance indices in the extracorporeal shock-wave lithotripsy outcome of ureteral stones. *Scand J Urol Nephrol* 2008; 42: 364-368.