Sectorial Technetium-99m-Dimercaptosuccinic Acid Scintigraphy for Monitoring the Effect of Extracorporeal Piezoelectric Lithotripsy for Calyceal Calculi on Regional Renal Function

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Key Words
Renal calyceal calculi · Extracorporeal piezoelectric lithotripsy · Technetium-99m-DMSA sectorial scintigraphy

Abstract
Objective: To apply a semiquantitative method for analysis of technetium-99m-dimercaptosuccinic acid (99mTc-DMSA) renal scintigraphy for monitoring the effect of extracorporeal piezoelectric lithotripsy (EPL) in patients with calyceal stones on regional kidney function and to check whether EPL had caused any deleterious effect on the target calyceal renal parenchymal function. Patients and Methods: Forty patients (mean age 35 years) suffering from calyceal stones documented by abdominal plain radiography, intravenous urogram or abdominal ultrasond were studied. All patients were treated by EPL. 99mTc-DMSA scan was performed before and 4 weeks after EPL. Sector analysis involved calculation of the relative function of the target calyx to the function of the ipsilateral kidney and the relative function of the treated kidney to global renal function. Results: The stone sizes were 6–11 mm in diameter and 11 were located in the upper, 13 in the middle and 16 in the lower calyx. After EPL, the overall stone clearance rate was 85% (100% for calculi in the upper and middle calyces, 62% for lower calyces). The sector analysis did not show statistically significant change of the relative regional (calyceal) or whole kidney function between the pre- and post-EPL 99mTc-DMSA scans. Using sector analysis, EPL appeared to be a safe modality and its usage was not associated with any untoward effect on calyceal or whole kidney function. Conclusion: Sector analysis of 99mTc-DMSA renal scan is a simple semiquantitative method for monitoring regional changes of kidney function after EPL for treatment of calyceal stone.

Introduction
Extracorporeal shock wave lithotripsy (ESWL) is currently a well-established procedure for the management of urinary stone disease which has greatly reduced the need for open stone surgery and percutaneous nephrolithotomy [1–5]. Variants of the shock wave generators commonly in use are the electrohydraulic, piezoelectric (EPL) and electromagnetic. Few serious side-effects have been reported due to shock wave trauma to the renal parenchyma,
which, fortunately, are transient in nature and usually recover without sequelae [6, 7]. However, in rare cases, permanent decrease in renal function after EPL can occur [8, 9]. Also, significant alterations of renal growth after ESWL have been reported in children [10–12]. Similarly, experimental studies in immature rats exposed to shock wave therapy demonstrated long-term effects on renal function and permanent histological damage [13].

In clinical practice, technetium-99m-dimercaptosuccinic acid (99mTc-DMSA) scintigraphy, a non-invasive procedure, has been used to evaluate the impact of various pathological processes on the renal cortex including cortical functional integrity and quantitation of differential renal function [14]. 99mTc-DMSA scintigraphy has been the most commonly used technique to study renal scarring after injury and stone clearance has been the most commonly used technique to study renal scarring after injury and stone clearance [10–12]. Also, significant alterations of renal growth after EPL can occur [8, 9]. The patients had a full clinical and biochemical evaluation before and after EPL. Lithotripsy was performed using a Richard Wolf Piezolith 2300 (Knittlingen, Germany). This device employed an ultrasound system for stone localization and piezoelectric energy for its disintegration. The power of the shock waves was gradually increased over the first 10 min to 100% of the machine capacity (output setting up to 4), which gave a peak pressure of approximately 700 bar (70 MPa) of focal zone 3 × 10 mm². The shock wave application was triggered as continuous pulses, up to 2 pulses per second. Four to five EPL sessions were required for every patient. The mean number of shock waves was 3,000 per session.

Renal scintigraphy was performed 2 h after intravenous administration of 185 MBq (5 mCi) of 99mTc-DMSA. Imaging was done in a supine position using a dual-head scintillation camera (Millennium, GE, USA) fitted with a low energy high resolution parallel hole collimator and interfaced to a computer. A 256 × 256 pixel matrix and a zoom of 2 was used. Static images of the abdomen in anterior, posterior and right and left posterior oblique were obtained for a preset time of 4 min. Quantitative analysis was done on the anterior and the posterior images to calculate the sector (calyceal) function by the geometric mean method. This involved drawing regions of interest around the target calyx identified from the structural imaging prior to the scan or on the static 99mTc-DMSA images. Another region is drawn on the whole kidney in addition to a background region. The product of the net count for the sector taken from the anterior and posterior views was obtained and the square root taken. The sector function was calculated by taking the ratio of sector geometric mean count to the whole kidney geometric mean count. The split renal function was also calculated using the standard geometric mean method in a similar way to that described for the sector analysis described above. 99mTc-DMSA scans were done before and 4 weeks after the EPL sessions. In both studies, the same method of analysis using the same regions of interest was applied.

### Statistical Analysis

The Student’s paired t test was used to calculate any statistical significance difference between pre- and post-EPL whole and sectorial renal function as measured by 99mTc-DMSA scan.

### Results

The stone sizes as determined by abdominal plain radiography, intravenous urography or abdominal ultrasound were 6–11 mm in diameter. The locations of the stones are given in table 1. Of the 43 patients, 40 completed the EPL sessions with successful fragmentation and clearance of the calyceal stone fragments from the upper and middle calyces. In the patients with lower calyceal calculi, residual stones were seen in 6 out of 16 cases (37.5%). The overall stone clearance was 34/40 (85%).

Typical 99mTc-DMSA renal scans obtained from patients are shown in figure 1a, region of interest for sectorial analysis in figure 1b and that of split renal function in figure 1c. The results of the sector and whole kidney analysis of the 99mTc-DMSA scan are shown in table 2.
Table 1. Distribution of stones and stone clearance rates among the group studied

<table>
<thead>
<tr>
<th>Location of calculus</th>
<th>Right kidney</th>
<th>Left kidney</th>
<th>Total</th>
<th>Stone clearance rate</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>male</td>
<td>female</td>
<td>male</td>
<td>female</td>
</tr>
<tr>
<td>Upper calyx</td>
<td>6</td>
<td>1</td>
<td>2</td>
<td>2</td>
</tr>
<tr>
<td>Middle calyx</td>
<td>4</td>
<td>3</td>
<td>4</td>
<td>2</td>
</tr>
<tr>
<td>Lower calyx</td>
<td>6</td>
<td>3</td>
<td>6</td>
<td>1</td>
</tr>
</tbody>
</table>

1 The overall stone clearance rate was 34/40 (85%).

Table 2. The effect of treating calyceal stones using EPL on calyceal and whole kidney function as assessed by $^{99m}$Tc-DMSA scan

<table>
<thead>
<tr>
<th></th>
<th>Upper calyceal stones (n = 11)</th>
<th>Middle calyceal stones (n = 13)</th>
<th>Lower calyceal stones (n = 16)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>pre EPL</td>
<td>post EPL</td>
<td>pre EPL</td>
</tr>
<tr>
<td>Affected kidney to both total kidney uptake, %</td>
<td>47.2 ± 3.2</td>
<td>48.1 ± 4.1</td>
<td>47.7 ± 9.2</td>
</tr>
<tr>
<td>Affected calyx to ipsilateral kidney uptake, %</td>
<td>21.6 ± 1.3</td>
<td>22.7 ± 4.2</td>
<td>24.1 ± 6</td>
</tr>
</tbody>
</table>

$p$ values were between 0.13 and 0.18 (not statistically significant).

Ipsilateral kidney uptake is taken to be 100%. The percent uptake shown in the second row is the percentage function of the affected calyx to the kidney function on the affected side.

Discussion

The use of EPL represents a major advance in the treatment of urinary calculic disease due to its effectiveness in disintegrating renal stones and the relative non-invasive nature compared to surgery. Many researchers are still investigating its short- and long-term effects on the kidney [1, 4, 21] and as such many studies have been performed to evaluate the effect of EPL on renal function [4, 14, 22]. Gilbert et al. [9] evaluated several basic physiological parameters, including creatinine clearance, fractional sodium excretion, protein excretion, glomerular filtration rate, effective renal plasma flow and urinary osmolality before and after EPL to quantify the changes in renal function. They found that the treated kidney appeared to maintain its ability to dilute urine and conserve sodium. They also reported that the renal physiological parameters correlated well with renal uptake of $^{99m}$Tc-DMSA. Similar findings have been reported by Kaude et al. [22]. Groshar et al. [21] also found that EPL did not cause any effect on the renal cortical function of the treated kidney as determined by $^{99m}$Tc-DMSA uptake.

$^{99m}$Tc-DMSA renal scans have been shown to be ideal for evaluating long-term functional and morphological parenchymal sequelae of ESWL [16, 17, 23, 24]. Several studies have evaluated the renal parenchymal effects of ESWL using $^{99m}$Tc-DMSA renal scan. Scarring has been observed in kidneys with calculi treated with ESWL about 3 months after treatment, the degree of scarring depending on the number of shock waves administered [18]. However, some of these lesions have been found to be reversible [19, 20, 25–27]. These studies have been essentially performed in children to assess any lasting damage to the immature kidneys [25–27]. In adults similar studies have also reported transient effects due to ESWL on the kidneys as determined by $^{99m}$Tc-DMSA scans [9, 16, 18,
Fig. 1. a $^{99m}$Tc-DMSA static images of the abdomen in multiple views in a patient with a lower calyceal stone of the left kidney. Decreased uptake is seen in the lower pole (arrow) corresponding to the location of the stone (pre-EPL scan). LPO = Left posterior oblique; RPO = right posterior oblique. b Sector analysis showing the region of interest around the lower calyx (Sector) and the whole kidney (L). BG = Background. c Split renal function analysis showing regions of interest around each kidney (R, L) and a background region (BG).

20]. However, there have not been studies that have looked specifically at the function of the target area of the kidney as done in our study. A major disadvantage of studies assessing whole renal function using $^{99m}$Tc-DMSA scintigraphy to see if any deleterious effect is produced by ESWL is the dilution effect of the focal abnormality by evaluating whole kidney uptake and the well-known phenomenon of compensation of function by the unaffected contralateral organ or the unaffected part of the same organ. Hence, subtle changes in function of a target part of the kidney may not be detected by whole renal $^{99m}$Tc-DMSA scan. In this study we have evaluated specifically the effect of EPL on the function of the target calyx as well as that of the whole of the affected kidney using $^{99m}$Tc-DMSA. The ipsilateral kidney was chosen for monitoring the effect on the sector function as a change in the ratio of the sector uptake to that in the whole kidney between baseline and post-EPL, thus providing a more reproducible way for assessing regional function. A slight underestimation of the damage from lithotripsy may have been introduced due to a slight decrease in the whole renal function on that side, leading to a smaller denominator in the calculated ratio. The percentage uptake of the sectorial and affected kidney showed no significant changes in the uptake values before and after treatment.

The overall stone clearance rate obtained in this study was 85% and is comparable to the result from most centres [5, 7]. The stone clearance rates from the upper and middle calyces were 100%. As with big renal calculi, the position of the calculi may influence the clearance rate. Thus even though calculi in the lower pole or calyx of the kidney may fragment, they may not necessarily be cleared due to the effect of the anatomical position of the calculi as well as the negative effect of gravity.

Conclusion

Using $^{99m}$Tc-DMSA renal sector scintigraphy for monitoring the effect of EPL on the kidney, it was shown that EPL caused no significant damage to the function of either the target calyx or the whole kidney. Sector analysis provided a simple method for evaluation of regional renal function that would be more accurate for follow-up of localized injury to the kidney.
References


