Acute Renal Infection in Adult Part 1: An Overview of What the Radiologist Needs to Know

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Received June 13, 2019
Accepted June 24, 2019

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Abstract

Acute renal infection or acute pyelonephritis (AP) denotes the process of inflammation of the renal parenchyma and its collecting system and the urothelium following infection. Uncomplicated AP commonly affects otherwise healthy, young women without structural or functional urinary tract abnormalities and without relevant comorbidities. More severe and complicated AP occurs in patients with a structurally or functionally abnormal genitourinary tract, or in persons with a predisposing medical condition like immune compromised state and diabetes. Complicated AP is characterized by a broader spectrum of clinical presentations, a wider variety of infecting organisms, and a greater risk of progression to a complication, such as intrarenal or perinephric abscess or emphysematous pyelonephritis and has the capacity to damage the organ and at times maybe life threatening. Role of imaging in renal infection is secondary, and in most situations, imaging is done to confirm the clinical diagnosis, map progression of disease in immune-compromised group of patients, or to evaluate for potential complications and therapeutic interventions. This article attempts to discuss the pathophysiology of AP from the standpoint of medical imaging and also brings out illustrative examples of various manifestations of AP and its complications. It provides imaging insight into various stages of inflammation, development of complication, and a roadmap for understanding AP through cross-sectional imaging.

Keywords
► acute pyelonephritis
► papillary necrosis
► pyonephrosis
► renal abscess

Introduction

Acute renal infection or acute pyelonephritis (AP) denotes the process of inflammation of the renal parenchyma and its collecting system and the urothelium following infection. Uncomplicated AP commonly affects otherwise healthy, young women without structural or functional urinary tract abnormalities and without relevant comorbidities. More severe and complicated AP occurs in patients with a structurally or functionally abnormal genitourinary tract, or in persons with a predisposing medical condition like immune compromised state and diabetes. Complicated AP is characterized by a broader spectrum of clinical presentations, a wider variety of infecting organisms, and a greater risk of progression to a complication, such as intrarenal or perinephric abscess or emphysematous pyelonephritis and has the capacity to damage the organ and at times maybe life threatening. As histological specimens are difficult to obtain, clinical correlation with stages of inflammation and developing complication is nearly impossible. In day-to-day...
clinical practice, the primary goal of imaging is to provide information about the nature and extent of the disease and to identify significant complications, for example, gas-forming infection, abscess, and urinary obstruction.\(^2\)

Ultrasonography (USG) is the most widely used screening tool, although it is limited in its ability to map disease progression and suffers from operator bias. Currently, computed tomography (CT) with or without CT urography and magnetic resonance imaging (MRI) find wide acceptance for diagnosis of AP and its potential complications. This article attempts to discuss the pathophysiology of AP from the standpoint of medical imaging and also brings out illustrative examples of various manifestations of AP and its complications. It provides imaging insight into various stages of inflammation, development of complication, and a roadmap for understanding AP through cross-sectional imaging.

**Acute Pyelonephritis: Terminological Conundrum**

Numerous verbal expressions have added to the confusion about the terminology used to describe the extent and severity of AP. Popular terminology used by radiologists was based solely on imaging appearance without histological confirmation. The advanced generalized form of AP in diabetic patients, usually with a nonfunctioning kidney on excretory urography, was originally described as acute bacterial pyelonephritis.\(^3\) The localized form of AP resulting in a mass-like appearance in imaging studies was given numerous names including acute focal pyelonephritis, acute focal bacterial nephritis, and acute lobar nephronia\(^4,5\) (►Fig. 1).

As cross-sectional modalities have evolved, subtle abnormal nephrographic findings were picked up variably termed as cellulitis, carbuncle, and renal phlegmon. In 1984, Talner et al\(^6\) proposed that all renal parenchymal abnormalities without formation of abscess attributable to acute infection maybe called AP. It was proposed that degree of involvement should be described by one or more of the following modifiers: (1) unilateral or bilateral, (2) focal or diffuse, (3) focal swelling or no focal swelling, and (4) renal enlargement or none.

**Acute Pyelonephritis: Must Know Clinical Issues**

Clinically AP is suggested by flank pain, nausea, and vomiting, fever (>38°C), or costovertebral angle tenderness, and it can occur in the absence of symptoms of cystitis.\(^7\)

AP in pregnant women may not only develop into renal and respiratory insufficiency, but also there is added risk of frequent preterm labor.\(^8\) It is of great importance to diagnose AP confidently and pick up complications early to provide effective treatment. Most men with febrile urinary tract infection (UTI) develop concomitant prostatitis.\(^9\) Prostate infections are notoriously difficult to cure and need long-term antibiotics. A diagnosis of AP in a man should remind us to image the prostate, preferably by transrectal USG or a diffusion-weighted MRI (DWI). Diabetic patients with AP are at risk of developing hypo- and hyperglycemia, hyperosmolar dehydration, or ketoacidosis\(^10\) and also infection by gas-forming organisms with a high mortality (emphysematous pyelonephritis). Emphysematous pyelonephritis is characterized histologically by acute pyogenic infiltration with microabscesses and the development of acute renal failure.\(^11\) Laboratory findings of AP include urinary pus cells in excess, granular or leukocytic casts, bacteruria, and a positive urine culture. Blood tests may show leukocytosis with a neutrophilic shift, elevated erythrocyte sedimentation rate, elevated C-reactive protein levels, and occasionally positive blood cultures that grow the same organism as cultured from the urine.

**Acute Pyelonephritis: Pathology and Pathogenesis**

The heralding event is inoculation of the urinary bladder with an infectious organism, which then migrates up the ureter to the central collecting system. Vesicoureteral reflux (VUR) is the retrograde flow of urine from the bladder into the ureter and toward the kidney secondary to a dysfunctional vesicoureteric junction (VUJ). The VUJ normally acts as a one-way valve, allowing urine flow from the ureter into the bladder and closing during micturition, preventing back flow. Shorter intramural submucosal segment of distal ureter increases the likelihood of VUR. The ratio of ureteric submucosal tunnel length and ureteric diameter critically determines the valve action. In healthy individual, the ratio is 5:1, while in most patients with primary VUR, it is ~1.4:1. The ascent of infection from bladder to kidney occurs even in...
the absence of VUR, owing to special virulence properties of the bacteria such as the adhesin P fimbriae and endotoxins. The bacterial endotoxins are believed to inhibit ureteral peristalsis by inhibiting the adrenergic nerves that supply the smooth muscle of the ureter. The ureter as a result lays flaccid creating a functional obstruction and urinary stasis. The ureteric obstruction compromises the forward flow of urine, which is a normal protective mechanism against upper UTI. Infection originates from the lower urinary tract and ascends via subepithelial lymphatic channels or directly via the ureter. Escherichia coli (E. coli) is the most common pathogen. In elderly patients with a history of instrumentation, however, Proteus mirabilis is a frequent offender. E. coli has the ability to attach to urothelial surfaces via pili or fimbriae. Once the kidneys are infected there is either global renal enlargement or sometimes there is patchy areas of inflammation in different stages. Typically, the pelvicalyceal system is involved first (pyelitis). The renal interstitial inflammation starts from the medullary region toward the cortex. Hematogenous infection occurs less commonly, where the renal cortex is involved first.

Complications of AP include abscess formation and spread of infection to perirenal space and beyond.

**Imaging of Acute Pyelonephritis and Its Complications**

**When Do You Image Acute Renal Infection?**

Mostly UTIs are treated without imaging as the diagnosis is straightforward. In confounding situations where significant clinical remissions are not seen following antibiotic therapy of 3 days, imaging is called for. In the situation when a definite diagnosis of AP is not established or when patients present with recurrent episodes of infection, renal imaging is indicated because there is more likelihood of stones, obstruction, abscess, or a congenital anomaly.

Imaging is also requested in a select group of patients where risk of complications runs high.

Patients with poorly controlled diabetes mellitus, acquired immunodeficiency syndrome, renal transplantation, or other immunocompromised disease states are considered at risk of developing a complicated UTI, including development of renal or perinephric abscess. Therefore, imaging may be required when such patients present initially. Special mention must be made about the case of diabetes. Apart from risk of abscess formation and extrarenal infection spread, diabetics warrant early imaging in UTI. Up to 50% may not have the typical flank tenderness that helps to differentiate AP from a lower UTI. Absence of flank pain in diabetics might be a critical confounding factor and vigil against missing the narrow window of opportunity to initiate early treatment must be kept.

**Role of Ultrasonography in Acute Pyelonephritis**

USG is one of the first imaging investigations to be ordered because it is easily available and gives a quick look helping to rule out any predisposing factors like obstruction as well as intra or perirenal collections. However, signs of early AP may not be visible on USG and mild swelling as well as loss of corticomedullary differentiation present in such cases is subjective and may be missed.

A large study from South India on role of emergency ultrasound screening (EUS) in the evaluation of 1,218 patients with clinically suspected AP found that nearly 49% had a normal EUS, while 51% had at least one major or minor abnormality. The frequency of hydronephrosis, renal calculi, emphysematous changes, and renal abscess was 19.1, 8.9, 2.1, and 1.9%, respectively. Only 5.9% patients required emergency percutaneous nephrostomy or drainage of an abscess. They concluded that a large proportion of AP patients has only normal or minor abnormalities and do not need additional screening and intervention.

**CT versus MRI—How Do You Image Acute Pyelonephritis?**

The American College of Radiology appropriateness criteria for AP (revised 2018) state that CT abdomen and pelvis with IV contrast or CT abdomen and pelvis without and with intravenous (IV) contrast are usually appropriate for imaging complicated patients in the setting of AP. It also considers imaging investigation as “usually not appropriate” in uncomplicated cases. CT is a sensitive tool for AP and its sequelae, and it also provides a global assessment of the abdomen for coincidental pathology. Renal obstruction by urolithiasis is well picked up by CT apart from its ability to provide superior anatomic detail for underlying congenital, as well as acquired renal abnormalities.

Ability of contrast-enhanced CT to predict parenchymal changes, change in renal perfusion, and function are useful predictors of clinical outcome. Studies comparing CT with USG conclude that additional benefit of CT lies in its ability to detect parenchymal abnormalities in patients with AP that are generally missed by USG. CT scores over MRI in its ability to detect calculi and gas in pyelonephritis. Soulen et al emphasized validity of waiting for 72 hours prior to obtaining CT in patients with AP. In their series, 95% of patients with uncomplicated AP became afebrile within 48 hours of appropriate antibiotic therapy, and nearly 100% did so within 72 hours. In situations where use of iodinated contrast material must be avoided, MRI is fast emerging as an alternative to CT in imaging AP.

**Role of DWI**

In biological tissues that are highly cellular (such as tumors), the higher density of cell membranes restricts the diffusion of water protons. This restriction to diffusion manifests as high signal intensity on DWI and corresponding lower apparent diffusion coefficient (ADC). Use of ADC can serve as a noninvasive sensitive marker for renal fibrosis as an apparent decrease in ADC is associated with increased number of fibroblasts. Currently, it has been first author’s institutional practice to obtain DWI in situations where use of IV contrast is risky in diabetics with borderline glomerular filtration rate, pregnant women suspected of UTI, children with UTI showing prolonged fever and not responding to treatment. Comparison of DWI versus contrast-enhanced MRI was done for pediatric
population with equivalent outcome for both techniques. Rathod et al did a prospective study with DWI in a series of cases and found a higher sensitivity of DWI (95%) compared with noncontrast CT (67%) and contrast-enhanced CT (88%) in the diagnosis of AP. Areas of AP show significantly lower ADC values than normal renal cortical parenchyma. Renal abscesses show significantly lower ADC values than areas of AP. Similar observations of high sensitivity, specificity, and accuracy (each at 95%) with DWI in uncomplicated AP were reported by other studies too. Another exciting application of DWI is that of differentiation of pyonephrosis from hydronephrosis. Low ADC values of pyonephrosis debris were found particularly helpful for pregnant patients in the second and third trimesters who may have physiological hydronephrosis due to compression by the gravid uterus by Chan et al. Prediction of renal function by DWI is found to be a reproducible technique by Thoeny et al.

**CT Protocol for Acute Pyelonephritis**

Craig et al. and Stunell et al. have studied pyelonephritis by only precontrast and nephrographic phase CT in cases where obstruction is not suspected. Taniguchi et al. reported that scans using only the nephrographic phase had high accuracy with triphasic scans (which also included precontrast and excretory phases) for the diagnosis of AP and urolithiasis. This study reported an accuracy of nephrographic phase only CT of 90 to 92% in the diagnosis of AP and CT of 96 to 99% in the diagnosis of urolithiasis. Currently in our institutions, we justify use of unenhanced CT for detecting calculi, gas-forming infections, and hemorrhage. First and second authors institutional protocol includes a nephrographic phase at 60 to 90 seconds and excretory phase at 5 to 10 minutes post IV contrast injection studies to completely evaluate patients with renal inflammatory disease and also to define changes in the renal excretion of the contrast material that occur as a result of the inflammation. Papillary necrosis that is not uncommon is also better evaluated in the excretory phase. Our current experience also dictates that certain findings like streaky nephrogram, delayed parenchymal enhancement, and urothelial thickening are well picked up in excretory phase images.

Second author’s institutional protocol additionally includes a corticomedullary phase at 25 to 30 seconds post-IV contrast injection to cover any vascular complications like pseudoaneurysms, preoperative arterial mapping if any surgical intervention is required and assessing incidentally detected cysts and masses (Fig. 2).

**MRI Protocol for Acute Pyelonephritis**

At First author’s institution, DWI is used with multiple b values (−50, 600, 1,000). The protocol includes fat-saturated T2 in axial and T2 coronal, axial, and coronal LAVA (which gives in and opposed phase images, Water LAVA T1, Fat LAVA T1) and in most cases axial and coronal FS T1 following IV Gadolinium.

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**Imaging Findings of Acute Pyelonephritis: Illustrative Cases**

In first author’s institutional experience of over two decades and retrospective analysis of 100 cases of AP, the commonest imaging findings were global renal swelling (88%), perirenal fascial thickening (76%), perinephric fat stranding (56%), streaky nephrogram (44%), frank abscess (44%), microabscess (32%), gas in renal collecting system or parenchyma suggestive of emphysematous infection (16%), and abdominal wall involvement (8%).

**Streaky Nephrogram**

The common CT finding of striated or streaky nephrogram (Fig. 3) is seen as linear bands of alternating hyper- and hypotenuation orientated parallel to the axis of the tubules and collecting ducts. After administration of contrast material, this pattern of differential enhancement reflects the underlying pathophysiology of tubular obstruction caused by inflammatory debris within the lumen, interstitial edema, and vasospasm. All three of these pathophysiologic disturbances tend to decrease the flow of contrast agent through the renal tubule, which also helps explain the pattern of delayed and persistent enhancement seen 3 to 6 hours after administration of contrast material. The sites that originally
demonstrated reduced attenuation during the nephrographic phase transmute from hypoattenuation to hyperattenuation wedge-shaped defects because of the prolonged accumulation of contrast agent that slowly transits the compromised tubules. The learning point here is that though a streaky nephrogram is taught to be associated with AP, it is also common to be seen in case of renal vein thrombosis and acute renal or ureteric obstruction. Visualization of a thrombus in the renal vein and an anatomical obstruction or calculi in the ureter are not difficult to pick up on CT. In our experience, streaky nephrogram or hypoperfused areas may not be well seen in initial nephrographic phase in some situations, but may be better depicted in narrow window settings in the excretory phase images (►Figs. 4A–C, 5A and B).

Renal Swelling
The finding of renal swelling is best assessed by experience rather than measurement (►Fig. 6). Although renal swelling is a common feature, not all cases of AP present with swollen kidney. Perinephric fat stranding and perirenal fascial thickening are also found by the authors to be contributory but nonspecific findings.

Renal Microabscesses
Small microabscesses are seen in almost a third of cases of AP. These may coalesce and form larger abscesses and are well picked up by both CT and DWI (►Figs. 7 and 8).

Urothelial Thickening
Subtle urothelial thickening and increased urothelial enhancement in the walls of the renal pelvis as well as the ureter and along the mucosal layer of the urinary bladder can be well seen on CT and aid the diagnosis of AP especially in cases where renal changes are mild (►Fig. 9).

Renal Abscess
Renal abscess is a relatively rare consequence of treated cases of AP. But in poorly treated or inappropriately treated cases and in immunocompromised, especially in
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Postinflammatory Noninfected Fluid Collections
Postinflammatory cystic fluid collection may develop in an area of AP during or after antibiotic treatment (Fig. 12A and B). Renal abscess needs to be differentiated from postinfective sterile fluid collections that are sometimes seen following effective treatment. The principal differentiating point between renal abscess and postinflammatory sterile fluid is enumerated in Table 1.

Extrarenal Extension of Abscess
Spread of renal infection to perinephric spaces following abscess rupture is not uncommon. Abscess may extend to involve the perinephric spaces, the posterior abdominal wall, or psoas muscle (Fig. 13A and B).

Obstructing Pathology
Presence of obstructing calculus is a risk factor for developing infection. Obstruction of ureter causes increased tissue pressure in kidney that in turn impedes microcirculation and

within the abscesses, collecting system, and the parenchyma is a sign of fulminant infection and called emphysematous pyelonephritis. This will be dealt in the upcoming part 2 of our article (Fig. 11B).

Fig. 6 Coronal axial CT section in the nephrographic phase in a 23-year-old man reveals a significantly swollen left kidney with reduced enhancement (arrow).

Fig. 7 (A) An axial CT reveals swollen kidney with small microab- sesses (straight arrow), perirenal fascial thickening (elbow arrow), and dirty fat around kidney suggesting inflammation. (B) In a different case, microabsseses are (C) showing restricted diffusion in diffusion-weighted magnetic resonance imaging seen as hyperintense areas with corresponding lowered apparent diffusion coefficient (ADC) (arrow) in affected renal parenchyma. ADC value of inflamed parenchyma in right kidney measures 0.93 x 10^{-3} mm^2/sec to 0.95 x 10^{-3} mm^2/sec and in contralateral normal left kidney, it is 2.0 x 10^{-3} mm^2/sec.

Fig. 8 Panels A (axial) and B (sagittal) CT sections in a 68-year-old diabetic woman reveal clusters of tiny cysts representing microabsseses (straight arrows). Coalescing microabsseses evolving into a larger abscess are also seen (elbow arrow)

diabetics, renal abscess is a relatively frequent unwanted consequence. Hematogenous infections cause multiple abscesses (Fig. 10). CT is the modality of choice showing intrarenal loculations as well as extrarenal extent. There is excellent depiction of pus by DWI (Figs. 7B and C), which can be performed for patients who are unfit for contrast CT or where radiation exposure is a concern like in pregnant women and children. On CT renal parenchyma around the abscess cavity may appear poorly enhanced on the nephrogram phase due to edema. This hypoattenuating subjacent renal parenchyma may enhance slowly in delayed images excluding necrosis (Fig. 11A). Multiple abscesses may be seen where septic emboli seed the kidney. Presence of air
that leads to reduced ability to clear infection. It is important to image the urinary tract for any calculi, ureteric stricture, or mass that predisposes to urinary stasis and subsequent pyelonephritis (►Figs. 14–16).

**Pyonephrosis**

Pyonephrosis meaning pus within the collecting system is a situation where obstructed dilated collecting system is laden with pus. USG depiction of debris within a dilated collecting system or documentation of diffusion restriction in MRI helps to establish diagnosis. CT may detect thickened walls of the renal pelvis and ureter, hyperdense urine within the collecting system suggestive of debris, fluid-fluid levels (pus-urine, gas-urine) (Figs. 9 and 10).
Table 1 Differentiating postinflammatory fluid and renal abscess

<table>
<thead>
<tr>
<th>Postinflammatory sterile fluid</th>
<th>Renal abscess</th>
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<tbody>
<tr>
<td>• &lt;3 cm in diameter</td>
<td>• No size criteria</td>
</tr>
<tr>
<td>• No wall thickening or</td>
<td>• Wall thickening and enhancement</td>
</tr>
<tr>
<td>enhancement</td>
<td>• Perinephric fat strand-</td>
</tr>
<tr>
<td>• Perinephric fat remains</td>
<td>• Aspiration reveals pus</td>
</tr>
<tr>
<td>clean</td>
<td>• USG reveals debris</td>
</tr>
<tr>
<td>• Aspiration shows brownish</td>
<td>• Restricted diffusion</td>
</tr>
<tr>
<td>fluid</td>
<td>in MRI</td>
</tr>
<tr>
<td>• USG reveals anechoic fluid</td>
<td>• No diffusion restriction in MRI</td>
</tr>
<tr>
<td>without debris</td>
<td></td>
</tr>
<tr>
<td>• No diffusion restriction in</td>
<td></td>
</tr>
<tr>
<td>MRI</td>
<td></td>
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Abbreviations: MRI, magnetic resonance imaging; USG, ultrasonography.

Infected Renal Cysts
Infected renal cysts cannot be reliably differentiated radiologically from an abscess. An infected cyst may also mimic a renal cyst complicated by intracystic hemorrhage on imaging studies. Percutaneous cyst aspiration and drainage urine-debris, or contrast-debris levels), and gas within the collecting system to contribute to a diagnosis (Fig. 17).
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Fig. 16 Sagittal curved CT reconstruction in the nephrographic phase (A) and diagrammatic representation (B) through the left kidney, ureter, and bladder in a 33-year-old lady with acute pyelonephritis and hydronephrosis show a ureterocele (long arrow) as the cause of the obstruction. The upper pole of the kidney reveals a poorly enhancing area suggestive of infection (short stout arrow). Three loose lying nonobstructive calculi are seen in the distal dilated ureter (elbow arrows).

Fig. 17 Oblique coronal CT reconstruction in the nephrographic phase through the left kidney and ureter of a 78-year-old man with chronic left ureteric obstruction by calculi (short arrow). The urine reveals high density (25 HU) suggestive of debris signifying a diagnosis of pyonephrosis. An abscess is seen leading from one of the posterior calyces into the abdominal wall muscles at the back (long arrow).

Fig. 18 Sagittal CT maximum intensity (A), volume rendered (B), and colored volume rendered (C) images through the left kidney in the excretory phase of a 50-year-old diabetic woman with acute pyelonephritis show deformed ballooned calyces with filling defects (straight arrows) along with medullary clefts and deformities of the calyces (curved arrows). A minimum intensity projection through the right kidney in the plain scan (D) of another 68-year-old man with deranged renal functions shows pelvicalyceal mild dilatation along with clubbing and deformity of several calyces (curved arrows).

under CT or US guidance may be indicated for diagnosis and treatment.

Papillary Necrosis
AP is one of several diseases such as diabetes, analgesic overuse, sickle cell disease, renal vein thrombosis, tuberculosis, and obstructive uropathy that cause renal papillary necrosis. CT findings include contrast filled clefts in the renal medulla in the excretory phase, nonenhanced lesions surrounded by rings of excreted contrast material representing sloughed off medullae, and medullary calcifications. Sloughed off papillae may also be seen in the renal pelvis or ureter and may cause ureteric obstruction (►Fig. 18).

An imaging algorithm for patients with AP has been provided in ►Fig. 19
Conclusion

Early detection of AP is important to avoid complications especially in vulnerable groups of patients. A USG helps to rule out obstruction in most patients. CT helps in patients who have complicated AP, have not responded to treatment, or are very sick. The sensitivity and specificity of CT for early detection of AP and mapping extra renal extent of disease process are high. MRI is reserved for patients where CT is contraindicated.

Conflict of Interest
None declared.

References

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