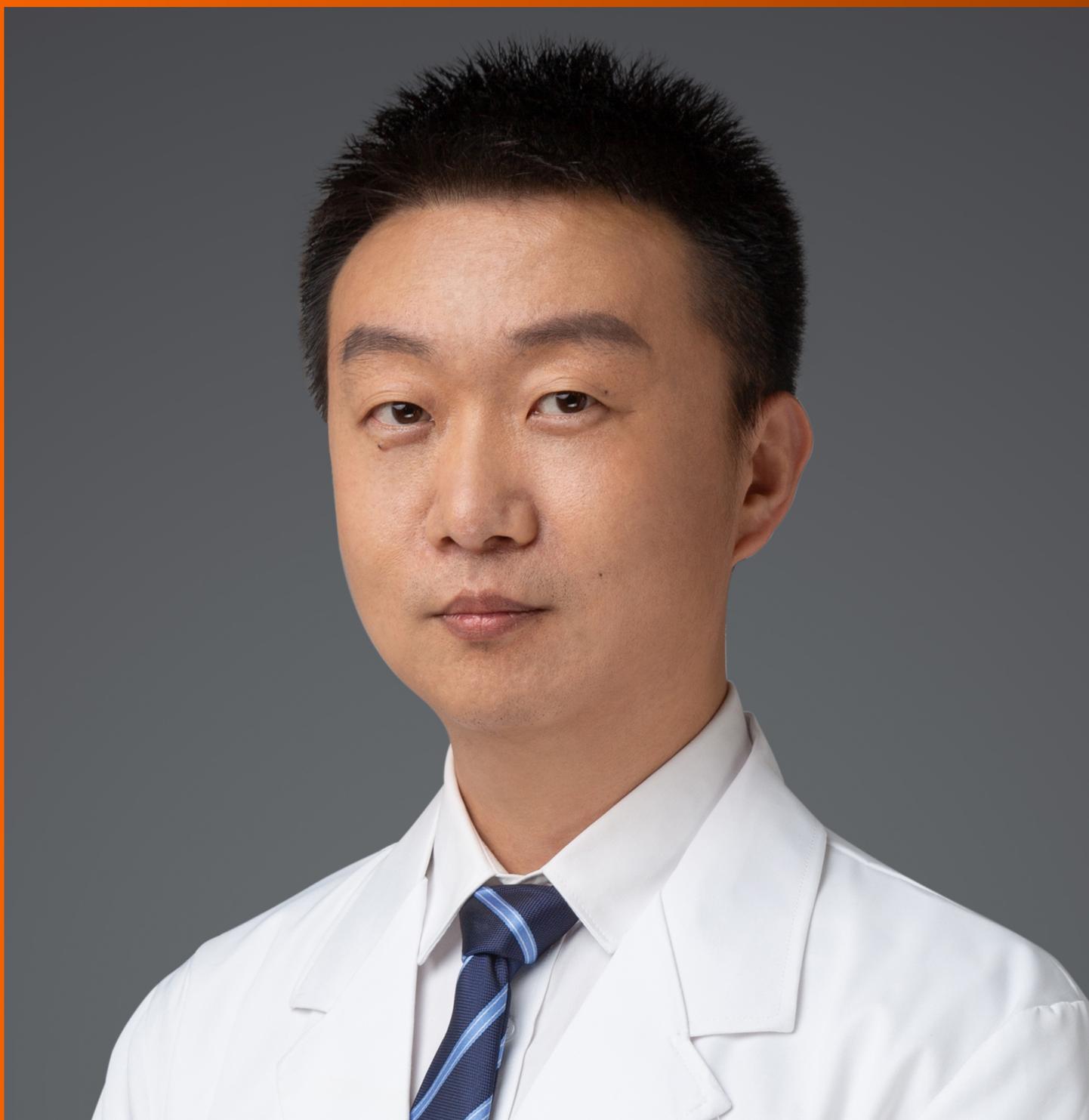


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## Point-of-care ultrasound for the early diagnosis of emphysematous pyelonephritis: A case report and literature review

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### Abstract

#### BACKGROUND

Emphysematous pyelonephritis (EPN) is a rare but fatal necrotic infection of the kidney, which usually leads to septic shock. Therefore, early diagnosis and optimized therapy are of paramount importance. In the past two decades, point-of-care ultrasound (POCUS) has been widely used in clinical practice, especially in emergency and critical care settings, and helps to rapidly identify the source of infection in sepsis. We report a rare case in which a “falls” sign on POCUS played a pivotal role in the early diagnosis of EPN.

#### CASE SUMMARY

A 57-year-old man presented with fever and lumbago for 3 d prior to admission. He went to the emergency room, and the initial POCUS detected gas bubbles in the hepatorenal space showing a hyperechoic focus with dirty shadowing and comet-tail artifacts. This imaging feature was like a mini waterfall. His blood and urine culture demonstrated *Escherichia coli* bacteremia, and EPN associated with septic shock was diagnosed. The patient did not respond to broad-spectrum antibiotic treatment and a perirenal abscess developed. He subsequently underwent computed tomography-guided percutaneous catheter drainage, and fully recovered. We also review the literature on the sonographic features of POCUS in EPN.

#### CONCLUSION

This case indicates that a “falls” sign on POCUS facilitates the rapid diagnosis of severe EPN at the bedside.

**Key Words:** Emphysematous pyelonephritis; Point-of-care ultrasound; Ultrasound; Urinary

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**Core Tip:** Emphysematous pyelonephritis (EPN) is a rare but life-threatening infection, and its diagnosis and treatment remain challenging. Point-of-care ultrasound (POCUS) plays an important role in rapidly assessing critically ill patients at the bedside. Here, we report a “falls” sign on the initial POCUS examination in a patient diagnosed with EPN associated with septic shock. We suggest that the “falls” sign may act as an imaging feature for early diagnosis of EPN. The patient was successfully treated with computed tomography-guided percutaneous catheter drainage plus broad-spectrum antibiotic therapy.

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## INTRODUCTION

Emphysematous pyelonephritis (EPN) is a lethal necrotic infection of the kidney with the key features of a collection of gas in the renal parenchyma, collecting system, as well as perinephric tissues<sup>[1]</sup>. More than 60% of patients with EPN have poorly controlled diabetes mellitus<sup>[2]</sup>. EPN usually presents with a fulminant clinical course and leads to sepsis and septic shock. Misdiagnosis of EPN and delayed management are associated with a mortality rate up to 80%<sup>[3]</sup>. Over the past decade, computed tomography (CT)-guided percutaneous catheter drainage (PCD), advanced antibiotic therapy, and intensive care medicine have improved the clinical outcome with a decreased mortality rate of 21%<sup>[4]</sup>.

To date, point-of-care ultrasound (POCUS) is widely used in day-to-day clinical practice<sup>[5]</sup>. It seems more important in emergency and critical care where radiological examinations are time consuming or unavailable. POCUS has been defined as “the new stethoscope” challenging traditional diagnostic practice<sup>[6]</sup>. Sepsis is life-threatening organ dysfunction induced by infection, which remains a global health priority<sup>[7]</sup>. Bedside POCUS can be used to rapidly assess major organs, and helps to identify a septic source, especially acute pyelonephritis, and to speed up the diagnosis<sup>[8,9]</sup>.

In the present case, we report a “falls” sign on POCUS examination, which contributed to the early diagnosis of EPN. We discuss the use of POCUS in EPN and review the relevant literature. To the best of our knowledge, this is the first study to report the “falls” sign in EPN and to systematically discuss the POCUS features of EPN.

## CASE PRESENTATION

### Chief complaints

A 57-year-old Chinese man complained of fever and lumbago for the last 3 d.

### History of present illness

The patient presented to the emergency room with a history of sudden onset persistent right flank pain, fever and fatigue for 3 d. POCUS was performed immediately and EPN was initially diagnosed. The patient was transferred to the intensive care unit (ICU) due to septic shock and an abdominal CT scan was carried out.

### History of past illness

The patient had a 10-year history of poorly controlled diabetes.

**Personal and family history**

The patient had a 30-year history of smoking and drinking, which he had recently stopped.

**Physical examination**

On admission to the ICU, physical examination revealed a temperature of 38.8 °C, heart rate of 130 bpm, and blood pressure of 108/74 mmHg with a moderate dose of continuously pumped norepinephrine (0.56 µg/kg/min) and respiratory rate of 22 breaths/min. His heart beat fast without murmurs and lungs sounded clear without crackles. His abdomen was soft and was not tender. He had severe knocking tenderness in the right flank. These findings indicated septic shock provoked by acute pyelonephritis.

**Laboratory examinations**

**Table 1** shows the initial laboratory findings. Blood analysis revealed leukocytosis of  $10.37 \times 10^9/L$  with neutrophils of 81%, hemoglobin of 11.9 g/dL, and thrombocytopenia (platelet count  $69 \times 10^9/L$ ) induced by sepsis. Alanine aminotransferase (21 IU/L), aspartate aminotransferase (23 IU/L), and bilirubin (0.58 mg/dL) were normal. He had a slightly elevated serum creatinine level (1.66 mg/dL) indicating acute kidney injury induced by severe infection of the kidney and septic shock. Inflammation markers were significantly increased, including C-reactive protein (175.1 mg/L) and procalcitonin ( $> 100$  ng/mL). The glycosylated hemoglobin level (9%) was elevated, indicating poorly controlled diabetes. His urine analysis showed heavy pyuria with a white blood cell count of 325/µL. Arterial blood gas analysis on admission showed a pH of 7.43, partial pressure of carbon dioxide of 36.8 mmHg, partial pressure of oxygen of 64.4 mmHg, bicarbonate of 24.8 mmol/L, and an elevated lactate level of 2.9 mmol/L with room air, indicating septic shock. Blood and urine samples were sent for culture, with positive results of extended spectrum beta-lactamase-producing *Escherichia coli* bacteremia.

**Imaging examinations**

Emergency POCUS on day 3 after symptom onset showed hyperechoic spotted or patchy foci in the right hepatorenal space with dirty shadowing and comet-tail artifacts (**Figure 1A**). We called this imaging feature a “falls” sign to describe the shadowing and “comet tails” radiating from the gas gathering in the hepatorenal space. It also presented a mini waterfall in Chinese landscape painting style (**Figure 1B**). The typical imaging findings speeded up the initial diagnosis of EPN.

**Further diagnostic work-up**

An abdominal CT scan on day 3 after symptom onset revealed gas collection in the right perirenal space, an enlarged right kidney with perinephric fat stranding (PFS) (**Figure 2A**) and mild right hydronephrosis without urinary stones. The CT scan confirmed the initial diagnosis of EPN based on emergency POCUS.

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**FINAL DIAGNOSIS**

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EPN associated with septic shock was the final diagnosis based on symptoms, physical examination, and imaging findings. Gas in the right perirenal space may result from necrotic pancreatitis and extraperitoneal hollow organ perforation, such as perforation of the descending duodenum<sup>[10,11]</sup>. The patient had a soft abdomen without symptoms of enteroparalysis, and further CT scan showed upper urinary tract infection. Hence, duodenal perforation and necrotic pancreatitis were unlikely to be the cause of gas in the right perirenal space.

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**TREATMENT**

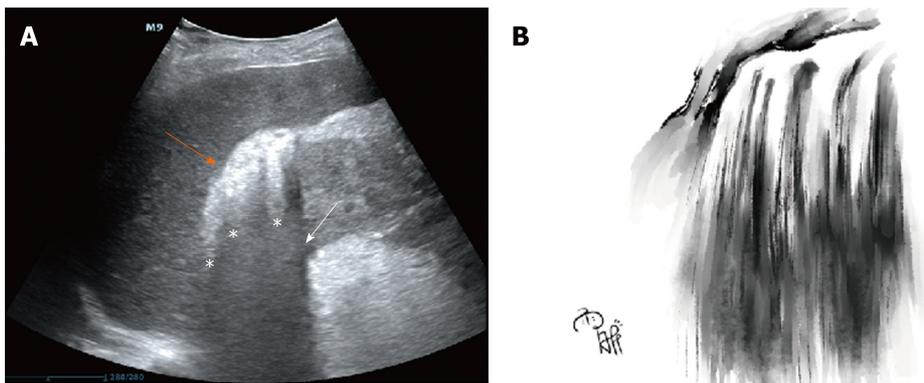
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The clinical course and vasopressor doses are shown in **Figure 3**. On admission to the ICU, the patient received fluid resuscitation, insulin infusion, vasopressor support, and 14 d of broad-spectrum antibiotic therapy including meropenem (3 g/d) and tigecycline (0.1 g/d). Septic shock did not respond to the initial therapy. A repeat CT scan was performed on day 7 after symptom onset (**Figure 2B**), and showed a more enlarged kidney with more PFS and gas plus an abscess in the right perirenal space. A

Table 1 Initial laboratory data consistent with sepsis

Variables	Results	Normal range
White blood cells	$10.37 \times 10^9/L$	$4-10 \times 10^9/L$
Percentage of neutrophils	81%	50%-70%
Hemoglobin	11.9 g/dL	11.5-15 g/dL
Platelets	$69 \times 10^9/L$	$100-300 \times 10^9/L$
Alanine aminotransferase	21 IU/L	9-50 IU/L
Aspartate aminotransferase	23 IU/L	15-40 IU/L
Total bilirubin	0.58 mg/dL	0.29-1.2 mg/dL
Creatinine	1.66 mg/dL	0.3-1.0 mg/dL
CRP	175.1 mg/L	0.068-8.2 mg/L
PCT	$> 100 \text{ ng/mL}$	$< 0.05 \text{ ng/mL}$
Glycosylated hemoglobin	9%	4%-6%
Urine white blood cells	$325/\mu\text{L}$	$0-5/\mu\text{L}$

CRP: C-reactive protein; PCT: Procalcitonin.

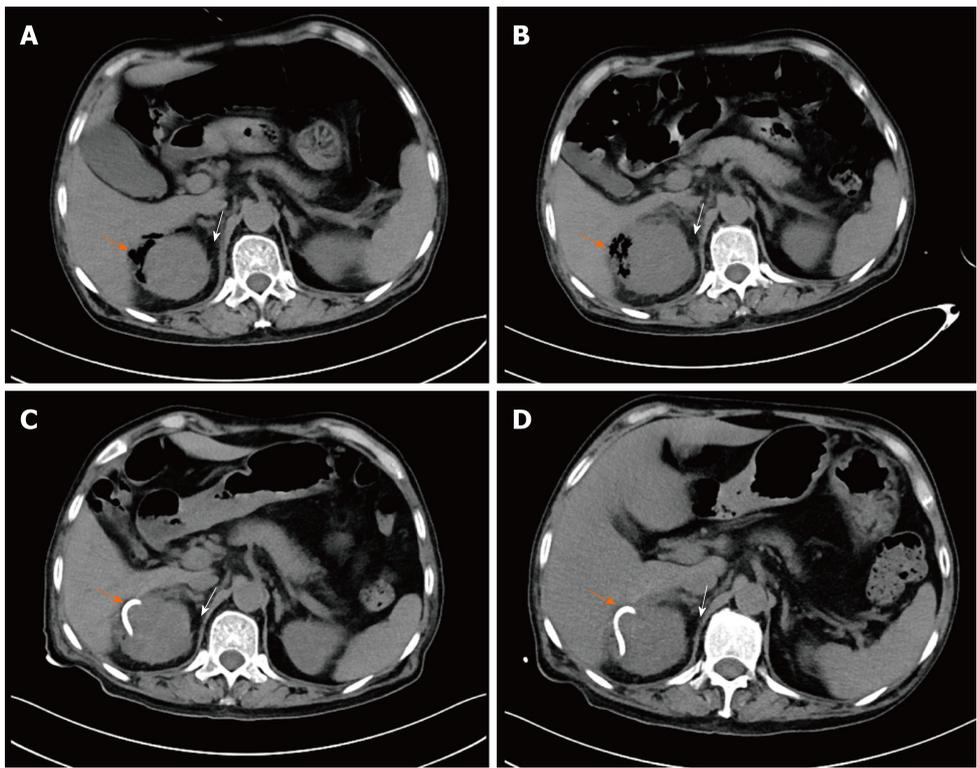


**Figure 1** Point-of-care ultrasound of a “falls” sign and a sketch of this sign. A: Image on day 3 after symptom onset showing hyperechoic spots or patches (orange oblique arrow) collecting in the right hepatorenal space with dirty shadowing (white oblique arrow) and comet-tail artifacts (white asterisks); B: Chinese landscape painting illustrating a mini waterfall by Yu-Xin Wang.

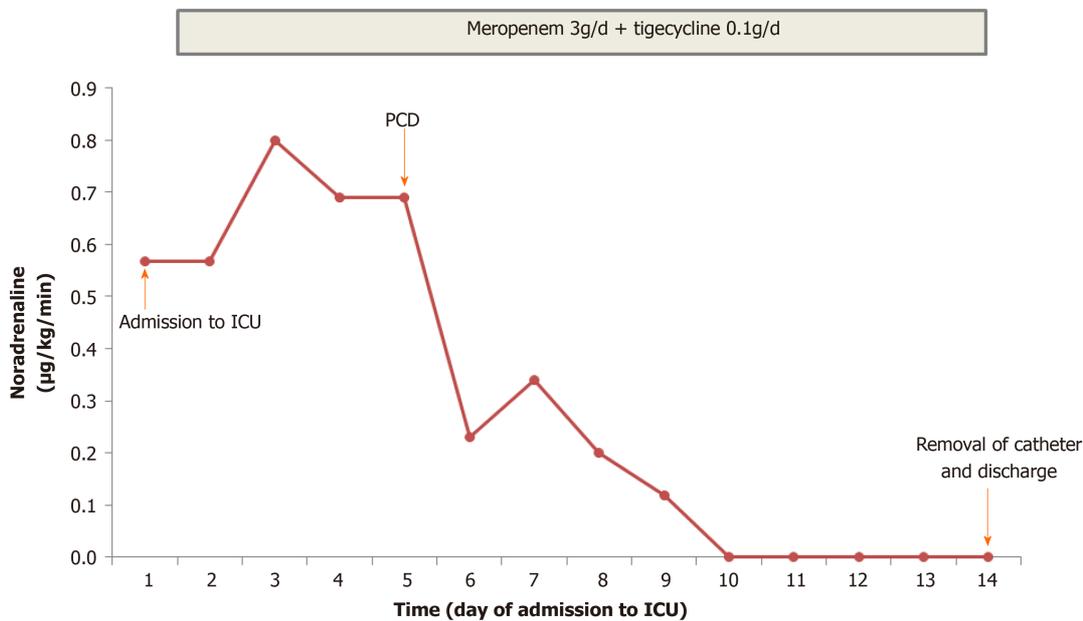
urological and interventional radiological consultation was obtained, and urgent CT-guided PCD was recommended for the patient on day 5 after admission. The culture from pus also yielded *E. coli* bacteremia. Double-J catheter (DJB) stenting was not advocated due to mild hydronephrosis of the right kidney and the absence of urinary stones.

## OUTCOME AND FOLLOW-UP

As shown in Figure 3, PCD associated with antibiotic therapy successfully reversed the clinical course. His clinical condition improved noticeably, and norepinephrine was discontinued within 5 d after initiating the combination therapy. CT reexaminations on days 9 and 11 after symptom onset (Figure 2C and D) revealed the pig-tail catheter in the right perirenal space and gas and abscess absorption. The patient was asymptomatic with a normal serum creatinine level and platelet count. The perirenal catheter was removed, and the patient was discharged with a 7 d course of oral levofloxacin (400 mg/d) on day 14 after admission. At 2 wk after discharge, a repeat urinary CT scan showed almost normal kidney imaging. The patient has been followed in an endocrinology clinic for his diabetes for 1.5 years. During follow-up, he remained healthy with stable blood glucose control and normal renal function. The



**Figure 2 Comparison of computed tomography scans of the right kidney.** A: Image on day 3 after symptom onset showing gas bubbles (orange oblique arrow) in the right perirenal space and an enlarged kidney with perinephric fat stranding (PFS) (white oblique arrow); B: Image on day 7 after symptom onset showing gas bubbles plus an abscess in the right perirenal space (orange oblique arrow) and a more enlarged kidney with more PFS (white oblique arrow); C: Image on day 9 after symptom onset showing a pig-tail catheter (orange oblique arrow) in the right perirenal space and an enlarged kidney with PFS (white oblique arrow); D: Image on day 11 after symptom onset showing a pig-tail catheter (orange oblique arrow) in the right perirenal space and a normal-size kidney with clear perinephric fat (white oblique arrow).



**Figure 3 Clinical course and vasopressor doses.** Meropenem and tigecycline were prescribed on days 1-14. Percutaneous catheter drainage (PCD) was performed on day 5. The perinephric catheter was removed and the patient was discharged on day 14. ICU: Intensive care unit.

patient was satisfied with his care.

## DISCUSSION

EPN is a type of life-threatening upper urinary tract infection with a high mortality rate and the hallmark of the presence of gas<sup>[1]</sup>. It has become a challenge worldwide, especially in developing countries with poor health care access<sup>[12]</sup>. There is a growing amount of literature focusing on EPN; however, most is limited to case reports. The major predisposing factor of EPN is uncontrolled diabetes, which decreases renal tissues perfusion and impairs host immune response<sup>[13]</sup>. In addition, the hyperglycemic environment facilitates the growth of facultative anaerobes. The most common causative organism in EPN is facultative anaerobic Enterobacteriaceae, especially *E. coli* and *Klebsiella pneumoniae*, which is in common with urinary tract infections<sup>[14]</sup>. Gas is produced by the pathogenic organism *via* fermentation of glucose and lactate in necrotic tissues<sup>[14]</sup>. In addition to diabetes, other risk factors for EPN include obstructive nephropathy, urolithiasis, chronic renal failure, hypertension, and immunosuppression<sup>[15,16]</sup>. A CT scan is recommended for most patients with EPN during the clinical course<sup>[17]</sup>.

In 2000, Huang published a pioneering clinicoradiological classification based on CT findings<sup>[18]</sup>. This classification has been assessed and used with widespread acceptance<sup>[12]</sup>. It classifies EPN into localized EPN (Classes 1 and 2) and extensive EPN (Classes 3 and 4), and shows the correlation between the class of EPN and its management<sup>[14,18]</sup>. Classes 1 and 2 indicate gas in the collecting system only and gas in the renal parenchyma only. Class 3A and B indicate the expansion of gas into the perinephric space and pararenal space, respectively. Class 4 indicates EPN in a solitary kidney or in bilateral kidneys<sup>[18]</sup>. In our case, the initial CT scan showed solitary gas in the right perirenal space only but no gas in renal parenchyma. The CT findings did not correspond with any of the classes in Huang's radiologic classification. We suggest that EPN with solitary gas in the perirenal space is a special type of localized EPN, which can be successfully managed with PCD associated with antibiotics.

As this case shows, common clinical manifestations of EPN include fever, flank pain, pyuria and sepsis-associated presentations such as shock and thrombocytopenia<sup>[19]</sup>. However, some rare cases have an insidious onset and unusual presentation. EPN may be completely asymptomatic, or may present with only nonspecific symptoms such as generalized weakness, polydipsia, and hiccups<sup>[20-22]</sup>. Also, EPN can pose a challenge for timely diagnosis by mimicking intestinal obstruction and hollow organ perforation<sup>[23,24]</sup>. Class 3B EPN often involves adjacent retroperitoneal organs, especially the psoas muscle<sup>[25]</sup>. However, there is a great diversity of the extension of gas in rare Class 3B cases including the pancreas, spine, thigh and biliary system<sup>[26-29]</sup>. Additionally, EPN is complicated by diabetic ketoacidosis, liver abscess, gut perforation, septic pulmonary emboli and necrotizing fasciitis in some refractory cases<sup>[1,10,30,31]</sup>.

Huang has suggested that most Classes 1 and 2 EPN can be managed by PCD combined with antibiotic therapy, and Classes 3 and 4 EPN with a fulminant course (more than two risk factors) require nephrectomy<sup>[18]</sup>. However, there is increasing evidence to show that the priority of a more conservative approach decreases the mortality rate from 80% to 20%<sup>[13,32]</sup>. With recent progress in medical care, most cases with extensive EPN can be successfully managed with PCD plus DJB stenting associated with antibiotic treatment<sup>[33-36]</sup>. Also, localized EPN responds well to antibiotic therapy alone with a good outcome<sup>[13,37]</sup>. A meta-analysis showed that emergency nephrectomy correlated with a higher mortality rate than a kidney-conserving therapeutic strategy<sup>[38]</sup>. Additionally, a standard management algorithm has been developed to optimize the treatment strategy to avoid aggressive nephrectomy<sup>[39]</sup>. Nephrectomy should be performed when there is no improvement with conservative therapy. As in our case, the patient with perinephric gas and abscess responds well to PCD plus aggressive antibiotic therapy. Prognostic factors for mortality in EPN include the need for hemodialysis, shock, altered mental status, thrombocytopenia, severe hypoalbuminemia and hyponatremia<sup>[36]</sup>.

Although CT is the gold standard for diagnosing EPN<sup>[13,40]</sup>, POCUS is portable and provides real-time information at the bedside without radiation exposure, and has become a promising tool facilitating rapid diagnosis in the past two decades<sup>[41]</sup>. The high acoustic impedance gradient between gas and renal tissues generates artifacts, which can be easily detected on POCUS at the bedside<sup>[42,43]</sup>. We performed a systematic literature search in PubMed using the key words "POCUS," "point-of-care ultrasound," "bedside ultrasound," "emergency ultrasound," "ultrasound," and "emphysematous pyelonephritis." A total of five other reports focusing on POCUS in EPN were identified<sup>[3,19,44-46]</sup> (Table 2). A hyperechoic focus with dirty acoustic shadowing is the most common sonographic feature on POCUS for the diagnosis of

**Table 2** Cases of emphysematous pyelonephritis diagnosed by point-of care ultrasound

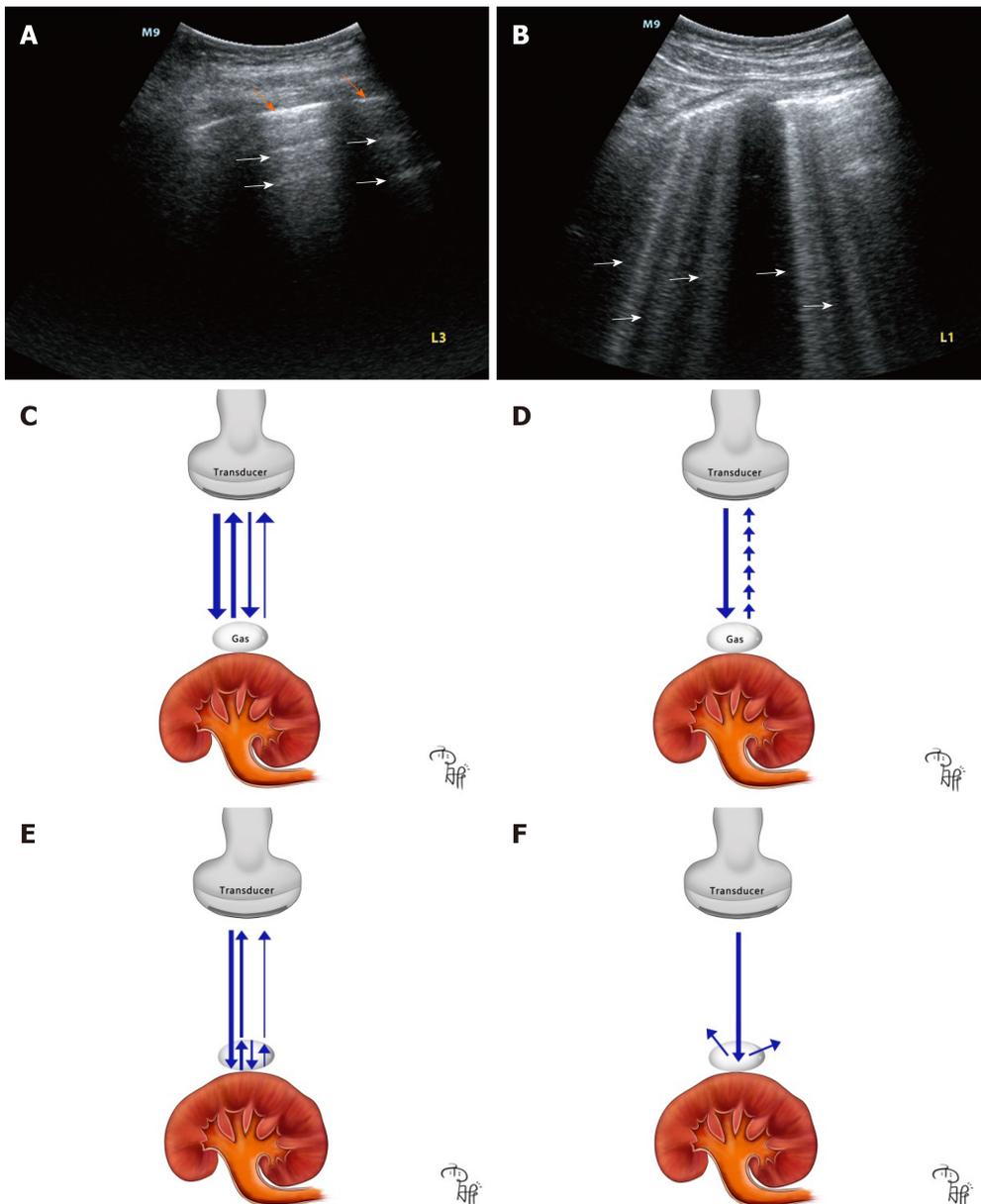
Ref.	Age in yr	Sex	Diabetes/comorbidities	Class of EPN	Treatment strategy	Outcome	Location of gas on ultrasound	POCUS features
McCafferty <i>et al</i> <sup>[3]</sup> , 2017	84	Woman	Diabetes/CKD/hypertension	Class 2	MM + nephrectomy	Recovered	Renal cortex	Hyperechoic focus/dirty shadowing
Stone <i>et al</i> <sup>[19]</sup> , 2005	47	Woman	Diabetes	Class 3A	MM + nephrectomy	Death	Renal parenchyma	Echogenic foci/dirty shadowing
Peng <i>et al</i> <sup>[44]</sup> , 2017	68	Woman	Diabetes	Class 3A	MM + nephrectomy	Recovered	Perirenal space	Poor delineation of the kidney
Koratata <i>et al</i> <sup>[45]</sup> , 2019	22	Woman	Diabetes	NM	NM	NM	Renal parenchyma	Hyperechoic focus/dirty shadowing/B-lines
Brown <i>et al</i> <sup>[46]</sup> , 2019	60	Man	Diabetes	Class 3A	MM + PCD + DBJ stenting	Recovered	Renal parenchyma/collecting system	A-lines

CKD: Chronic kidney disease; DBJ: Double-J catheter; EPN: Emphysematous pyelonephritis; MM: Medical management; NM: Not mentioned; PCD: Percutaneous drainage; POCUS: Point-of-care ultrasound.

EPN<sup>[3,19,45]</sup>. However, other imaging features have also been reported, including poor delineation of the kidney, A-lines and B-lines<sup>[44-46]</sup>. Additionally, we report that the comet-tail artifacts and the “falls” sign are also imaging features on POCUS in EPN. But physicians should keep in mind that these air-related artifacts on POCUS vary in different cases. The variation not only results from multiple effects of gas bubbles such as volume, shape, position, and orientation, but also correlates with a mismatch of acoustic impedance between the gas bubbles and its surrounding renal tissues<sup>[43]</sup>. Moreover, the utility of POCUS remains a challenge as a result of its dependence on the skills and experience of the operators, especially non-imaging professionals<sup>[41]</sup>. So, we suggest that the standardization of the air-related artifacts on POCUS in EPN should be implemented on the basis of sufficient faculty training.

Air surrounding the perirenal space prevents the transduction of sound waves resulting in artifacts, decreased visualization of deeper structures and an obscure outline of the kidney<sup>[43]</sup>. A-lines (Figure 4A) and B-lines (Figure 4B) are basic signs on lung ultrasound for the diagnosis of acute respiratory failure<sup>[47]</sup>. Both are artifacts generated when air is struck by ultrasound beams. A-lines are repetitive horizontal artifacts derived by repetitive reflection from the tissue-gas interface to the transducer (Figure 4C)<sup>[48]</sup>. B-lines are well defined, vertical, laser-like artifacts, and are generated by a ring down effect when the sound waves pass through gas bubbles associated with fluid collection, and provokes resonance within the air-fluid interface, emitting continuous waves back to the transducer (Figure 4D)<sup>[49]</sup>. A comet-tail artifact is produced when ultrasound beams are repeatedly reflecting on the shallow and deep sides of gas bubbles (Figure 4E)<sup>[50]</sup>, which usually looks like an inverted triangular hyperechoic lesion with reduced thickness and strength (Figure 1A).

Acoustic shadowing is a significantly reduced posterior echo, and it occurs when ultrasound waves pass through strongly reflecting or attenuating structures such as gas, bone, needles, calcifications and stones<sup>[51]</sup>. The “falls” sign should be differentiated between perirenal gas and perirenal calcification or renal wall calcification, which is non-specific pathology in renal wall tuberculosis<sup>[52]</sup>, perirenal tumors, polycystic kidney disease and very rare diseases such as Erdheim-Chester disease and tumoral calcinosis<sup>[53-56]</sup>. In most cases, perirenal calcification and urinary stones present with clean shadowing which is an absolute anechoic band. However, gas in EPN generates dirty shadowing which is a heterogeneous echoic band with reduced signal intensity<sup>[45]</sup> (Figure 1A). Previously, it was thought that clean shadowing was associated with sound-absorbing materials, such as stones, and dirty shadowing results from sound-reflecting materials, such as gas. However, studies have indicated that clean shadowing and dirty shadowing in essence correlate with the properties of the surface of the subjects, curvature and roughness, rather than the inner nature<sup>[57]</sup>. Dirty shadowing is considered the hallmark of ultrasound in EPN, and it is generated by reflection of ultrasound waves in multiple directions into the gas bubbles<sup>[49]</sup> (Figure 4F). We suggest that knowledge of the sonographic features of air-related artifacts in EPN plays an important role in physicians making an early diagnosis.



**Figure 4 A-lines and B-lines in pulmonary ultrasound in our clinical practice and cartoon illustrating how different air-related artifacts in emphysematous pyelonephritis are produced.** A: Point-of-care ultrasound (POCUS) of a healthy lung showing gradually diminished A-lines (the white arrows) and pleura lines (the orange arrows), and the equidistance between the lines; B: POCUS of lung edema showing B-lines (the white arrows); C: Cartoon showing how A-lines are produced. The ultrasound beams (the blue arrows) are repetitively reflecting between gas and the transducer with strength degradation; D: Cartoon showing how B-lines are produced. The ultrasound beam (the blue arrow) provokes resonance in the gas-fluid interface, emitting continuous waves back to the transducer (the small blue arrows); E: Cartoon showing how comet-tail artifacts are produced. The ultrasound beam is repetitively reflecting between the shallow and deep sides (the blue arrows) of gas bubbles with gradually diminished ultrasound beams returning to the transducer; F: Cartoon showing how dirty shadowing is produced. The ultrasound beam is reflecting in multiple directions (the blue arrows) deep into the gas.

Given the limitation of the case report, further cohort studies are needed to assess the diagnostic accuracy of air-related artifacts on POCUS *vs* CT imaging for EPN.

## CONCLUSION

EPN is a lethal gas-forming infection of the kidney. POCUS facilitates the timely diagnosis of EPN by the easily recognized hyperechoic focus associated with gas-related artifacts including A-lines, B-lines, comet-tail artifacts, dirty shadowing as well as a “falls” sign in our case. PCD plus antibiotic therapy can provide good clinical outcomes for most EPN cases.

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