Fatal renal failure as the first manifestation of sarcoidosis diagnosed on necropsy in a young man: a case report

A Awasthi, R Nada, P Malhotra, R Goel, K Joshi

Renal involvement as the first manifestation of sarcoidosis is rare and has never been reported in India. This report describes a 35 year old man who was admitted to the emergency department with a clinical diagnosis of acute on chronic renal failure, secondary to obstructive uropathy. Postmortem examination unexpectedly revealed disseminated sarcoidosis.

A 35 year old man presented to the emergency department with a one week history of oliguria and altered sensorium. There was no history of renal colic, haematuria, or dysuria and there were no respiratory complaints in the past. On examination, he had bilateral "wet" crackles suggestive of fluid overload and a biphasic pericardial rub. There was no lymphadenopathy or hepatosplenomegaly.

Investigations revealed severe dimorphic anaemia (haemoglobin, 44 g/litre), leucocytosis (total leucocyte count, 14.4 x 109/litre), and a raised erythrocyte sedimentation rate (82 mm first hour). Blood urea and serum creatinine were raised at 400 mg% and 9.7 mg%, respectively. Urine examination revealed 1+ albuminuria, pyuria (8–10 pus cells/high power field), and microhaematuria (8–10 red blood cells/high power field). The kidneys were normal in size (right, 9.7 cm; left, 9.9 cm) on ultrasonography. However, there was an indistinct corticomedullary differentiation, along with bilateral hydroureteronephrosis and left sided nephrolithiasis involving the left renal pelvis and a 2 cm diameter stone in the lower part of the left ureter. Microscopic examination revealed 1+ albuminuria, pyuria (8–10 pus cells/high power field), and microhaematuria (8–10 red blood cells/high power field). The kidneys were normal in size (right, 9.7 cm; left, 9.9 cm) on ultrasonography. However, there was an indistinct corticomedullary differentiation, along with bilateral hydroureteronephrosis and left sided nephrolithiasis. Biochemical analysis revealed normal serum sodium (130 and 133 mg%, respectively) and potassium (3 and 5.2 mg%, respectively) values. Inorganic phosphate was raised (11.7 mg%), whereas serum calcium was low to normal (8.8 mg%). Liver function tests and total proteins (60 g/litre; albumin to globulin ratio of 1.0) were within normal limits. Arterial blood gases revealed metabolic acidosis and there was evidence of pulmonary oedema on chest x ray. Pus from a 2 x 1.5 cm sized perianal abscess grew *Escherichia coli* sensitive to amikacin. Blood and urine cultures were sterile.

The patient was treated with intravenous cloxacillin and amikacin at renal modified doses. Eighteen cycles of peritoneal dialysis were carried out; however, he had an episode of ventricular fibrillation and died.

A complete necropsy was performed after informed consent from the relatives. Both kidneys were normal in size (left, 9.7 cm; right, 9.5 cm) with evidence of bilateral hydroureteronephrosis. There was a 3 cm diameter stone in the left renal pelvis and a 2 cm diameter stone in the lower part of the left ureter. Microscopic examination revealed pronounced nephrocalcinosis in the form of calcification of the basement membranes and tubular epithelium, along with the interstitium (fig 1). In addition, there were focal masses of calcification surrounded by polymorphs. Calcification of the smooth muscles and the intima of medium sized blood vessels and glomeruli (fig 1, inset) was also present. The lungs were subcrepitant and there were enlarged hilar lymph nodes (2.5 x 4 cm in diameter). On microscopy of the lungs, typical non-caseating epithelioid cell granulomas were seen (fig 2). Metastatic calcification was seen in the alveolar septae and blood vessels. The hilar lymph nodes showed diffuse paracortical sclerosis and the presence of fibroed granulomas in the cortex. Giant cells in these granulomas revealed asteroid bodies. The liver was normal on gross examination. However, microscopically there were stellate scars connecting portal tracts and central veins. Some of these scars were composed of healed epithelioid cell granulomas with giant cells containing asteroid bodies (fig 3). Similar lesions were seen in the spleen. There was no evidence of necrosis in the granulomas. Metastatic calcification was also present in the duramater, myocardium, and in the vessels of the gastrointestinal tract and brain.

Therefore, the final postmortem diagnosis was sarcoidosis involving the lung, hilar lymph nodes, liver, and spleen with renal involvement in the form of nephrocalcinosis, hydroureteronephrosis secondary to nephrolithiasis, and chronic interstitial nephritis. There was also evidence of widespread metastatic calcification in the duramater, myocardium, and the blood vessels of the lung, gastrointestinal tract, and brain.

DISCUSSION

Sarcoidosis, a disease that is increasingly being recognised in India, primarily involves the lungs and lymphatic system.

Renal involvement is less frequent; the incidence in the West ranges from 7% to 22%, whereas two large numbers of cases have been reported, e.g. a case involving a 30 year old woman who presented with nephrocalcinosis and a 6.5 cm nephrolithiasis in the left kidney and a 3 cm stone in the right kidney.

In contrast, this report provides evidence of renal involvement in the form of interstitial and tubular calcification in two separate patients with sarcoidosis. The incidence of renal involvement in sarcoidosis is not well documented, although it is believed to be lower than in other case reports.

A complete necropsy was performed to confirm the final diagnosis of sarcoidosis in the form of interstitial and tubular calcification. The kidneys were normal in size (right, 9.7 cm; left, 9.9 cm) on ultrasonography. However, there was an indistinct corticomedullary differentiation, along with bilateral hydroureteronephrosis and left sided nephrolithiasis. The kidneys were normal in size (right, 9.7 cm; left, 9.5 cm) with evidence of bilateral hydroureteronephrosis. There was a 3 cm diameter stone in the left renal pelvis and a 2 cm diameter stone in the lower part of the left ureter. Microscopic examination revealed pronounced nephrocalcinosis in the form of calcification of the basement membranes and tubular epithelium, along with the interstitium (fig 1). In addition, there were focal masses of calcification surrounded by polymorphs. Calcification of the smooth muscles and the intima of medium sized blood vessels and glomeruli (fig 1, inset) was also present. The lungs were subcrepitant and there were enlarged hilar lymph nodes (2.5 x 4 cm in diameter). On microscopy of the lungs, typical non-caseating epithelioid cell granulomas were seen (fig 2). Metastatic calcification was seen in the alveolar septae and blood vessels. The hilar lymph nodes showed diffuse paracortical sclerosis and the presence of fibroed granulomas in the cortex. Giant cells in these granulomas revealed asteroid bodies. The liver was normal on gross examination. However, microscopically there were stellate scars connecting portal tracts and central veins. Some of these scars were composed of healed epithelioid cell granulomas with giant cells containing asteroid bodies (fig 3). Similar lesions were seen in the spleen. There was no evidence of necrosis in the granulomas. Metastatic calcification was also present in the duramater, myocardium, and in the vessels of the gastroinestinal tract and brain.

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25(OH)2 vitamin D3 deficiency also interferes with calcium skeletal responsiveness to parathyroid hormone. The 1,25(OH)2 vitamin D3 result in reduced renal failure, reduced excretion of phosphate and decreased calcium absorption from the gut, which is already impaired in patients with nephrolithiasis, nephrocalcinosis, and advanced renal failure. In fact, the presence of metastatic calcification usually correlates with serum calcium concentration over 13 mg%.10 The low to normal serum calcium in our patient is also attributable to the presence of advanced renal failure. In renal failure, reduced excretion of phosphate and decreased production of 1,25(OH)2 vitamin D3 result in reduced skeletal responsiveness to parathyroid hormone. The 1,25(OH)2 vitamin D3 deficiency also interferes with calcium absorption from the gut, which is already impaired in uremia.

In conclusion, our case shows that renal failure resulting in death can rarely be the initial manifestation of sarcoidosis. The presence of normal serum calcium values in a patient with nephrolithiasis, nephrocalcinosis, and advanced renal failure does not exclude sarcoidosis as a possible cause; calcium concentrations may be normalised as a result of secondary hyperparathyroidism of renal failure or widespread metastatic deposition of calcium.

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