Spontaneous Massive Bladder Hematoma in the Late Period After Kidney Transplantation in Patients with AA Amyloidosis: Two Cases

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Abstract

Massive bladder hematoma can occur following a trauma, neoplastic states, or as a longterm side effect of a pelvic irradiation therapy. It is also reported, albeit rarely, in patients with systemic primary (AL) or secondary (AA) amyloidosis which can affect the bladder and can manifest itself as macroscopic hematuria, bladder hematoma, or bladder perforation. We presented two cases of spontaneous massive bladder hematoma in the late period after the transplantation in kidney recipients with systemic AA amyloidosis.

Keywords: Amyloidosis; hematoma; urinary bladder; familial Mediterranean fever; kidney transplantation

Systemic amyloidosis which is caused by the deposition of circulating misfolded proteins can occur due to plasma cell abnormalities [light chain (AL) or heavy chain (AH) amyloidosis], chronic inflammatory diseases (deposition of serum amyloid A, AA amyloidosis), in the presence of genetic mutations (deposition of transthyretin, ATTR amyloidosis) and in some other rare conditions. Massive bladder hematoma can occur following a trauma, neoplastic states, or as a longterm side effect of a pelvic irradiation therapy. It is also reported, albeit rarely, in patients with systemic primary (AL) or secondary (AA) amyloidosis which can affect the bladder and can manifest itself as macroscopic hematuria, bladder hematoma, or bladder perforation. We presented two cases of spontaneous massive bladder hematoma in the late period after the transplantation in kidney recipients with systemic AA amyloidosis.

Case Reports

Case 1

A 31-year-old male with end stage chronic kidney disease went through kidney transplantation from his sister. His primary kidney disease was chronic glomerulonephritis. He had stable functional allograft with triple immunosuppressive treatment during the post-transplant period, but he developed frequent episodes of lung infection in a timespan of seven years. Later on, he developed proteinuria which was increased to a nephrotic level in a short period of 6

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months. When the patient’s history was studied, it was found out that he had been suffering from general fatigue, syncope attacks and weight lost for a few months. A thoracoabdominopelvic computed tomography (CT) was performed and it showed bronchiectasis of the lower and middle lobes of the lungs bilaterally. There were frequent episodes of non-infectious diarrhea, which resulted in his becoming malnourished and cachectic. Therefore, gastroscopy and colonoscopy were performed and biopsies were taken from gastric and duodenal mucosa. The pathologic features were consistent with amyloidosis, the staining with amyloid A and P-component were positive on the entire vascular structural walls of the stomach and duodenum. His gene analysis showed no mutation in favor of Familial Mediterranean Fever. Thus, the patient was diagnosed as AA amyloidosis secondary to bronchiectasis. The patient was treated with 0.5-1.0 mg of colchicine daily. After two years, the patient was admitted to the hospital with severe macroscopic hematuria. There were no history of trauma and his coagulation parameters were in the reference range. Urinary ultrasonography showed hematoma in the bladder. Emergent cystoscopy revealed hematoma and perforation of bladder wall without any malignancy. Cystography showed that there was extravasation into the abdominal cavity. After cystography, the patient got septic, he needed broad-spectrum antibiotics and fluid resuscitation. Afterwards, his condition got worse and he died.

CASE 2

A 56 year-old male with secondary amyloidosis due to familial Mediterranean fever (FMF) developed end stage renal disease and had kidney transplantation from his wife. He had stable allograft function under the triple immunosuppressive and 1.0 mg of daily colchicine without an FMF attack. In the second year after transplantation, the patient was admitted to the hospital because of macroscopic hematuria without any history of trauma. Ultrasonography showed a 9.5×6.5-cm in size hematoma in the bladder. Cystoscopy revealed a large hematoma that completely filled the bladder. Urinary cytology showed no signs of malignancy. In a timespan of nine months, he had two more episodes of macroscopic hematuria. There were not any abnormality about coagulation parameters. Later on, he was admitted to emergency department with dizziness, lower urinary tract symptoms and fever. While the diagnostic work-up was still ongoing, he had sudden cardiac arrest, was resuscitated and admitted to intensive care unit with the diagnosis of urinary sepsis. Despite all efforts, he died because of urinary sepsis and heart failure.

The legal representatives of both of the patients have given their informed consents for these case reports.

DISCUSSION

Amyloid deposition may occur in the condition of an abnormal protein presence like light chains, in association with excess exposure to a normal protein like serum amyloid A (SAA) and for undiscovered reasons increased with age.\(^1\) The clinical presentation of amyloidoses depends upon the affected organs that include mainly the kidney, heart, nervous system, liver, and gastrointestinal and urinary tracts. Secondary amyloidosis occurs as a consequence of longstanding inflammatory diseases that lead to overproduction of serum amyloid A in the liver. Rheumatoid arthritis, familial Mediterranean fever (FMF), ankylosing spondylitis and Chronic’s disease are commonly associated with AA amyloidosis.\(^4\)

Secondary amyloidosis tends to accumulate within the vascular structures which cause amyloid angiopathy with increased fragility in blood vessels and impaired vasoconstriction which may explain why secondary amyloidosis of bladder has a high potential of massive hematuria and why renal biopsy in amyloidosis is known to be a hazardous procedure because of increased risks of bleeding.\(^5,6\) Although it is seen rarely, bladder amyloidosis mainly presents itself with painless macroscopic hematuria. Additionally, secondary bladder amyloidosis has a lower incidence than primary bladder amyloidosis.\(^3\)

In renal transplant recipients, macroscopic hematuria occurs with the prevalence of 12% after transplantation mostly because of infections, surgical complications, malignancies, graft rejections, recurrences of primary disease and calculus formation.\(^8\) Although there is a case of secondary bladder amy-
loidosis that presented itself with macroscopic hematuria in an FMF patient treated with renal transplantation, bladder hematoma or perforation without an obvious reason like trauma, clotting disorder, history of radiation or surgery is an extreme case and, to our knowledge, these are the very first cases with AA amyloidosis in renal transplant recipients. Although we had no chance to confirm bladder amyloidosis by biopsy, in both of these cases, there weren’t any other reasons about the patients to be prone to bleeding except from amyloidosis.

The control of underlying inflammatory disease is the main strategy for the treatment. Amyloidosis resulting from bronchiectasis has become rarer with the development of antimicrobial agents, but in immunosuppressive patients like renal transplantation recipients, it can be a problematic situation just like in our first case. Continuous colchicine use is recommended to prevent the involvement of other organs, even for end-stage renal disease patients with FMF, so its importance is greater in renal transplantation to prevent graft kidney amyloidosis.

CONCLUSION

We presented two cases of AA amyloidosis involving bladder caused hematoma and perforation, one of which being after renal transplantation. Both patients died because of the systemic involvements of amyloidosis. We believe that, in kidney recipients with systemic amyloidosis, bladder hematoma should be considered as an adverse prognostic factor.

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