Idiopathic Retroperitoneal Fibrosis Presenting with Hypertension and Acute Renal Failure

Hipertansiyon ve Akut Böbrek Yetmezliği ile Prezente Olan İdiopatik Retroperitoneal Fibrozis

Engin Onan1, Saime Paydaş1, Merve Erkoç2, Tuba Korkmaz2, Hasan Bilen Onan1, Mustafa Balal1

Introduction

Retroperitoneal fibrosis is frequently idiopathic, and autoimmune diseases such as Hashimoto thyroiditis, surgeries, drugs, and malignant diseases can cause idiopathic retroperitoneal fibrosis (IRF) (1, 2). Abdominal aortic aneurysm can trigger IRF with the leakage of inflammatory cells from aortic plaques to the retroperitoneal area (3). Retroperitoneal fibrosis usually occurs with pain in the back, flank, or abdomen; vomiting; leg edema; or urinary obstruction induced by fibrosis in the retroperitoneum (1, 4).

Case Report

A 49-year-old-man was admitted to the Nephrology Clinic with complaints of hypertension and abdominal pain. He was previously admitted thrice to other emergency rooms with unrelenting flank pain and hypertension before arriving at our clinic. Parenteral analgesics and antihypertensives were administered. Nausea, weight loss, and anorexia were added to the complaints in the previous 2 months.

On examination, his blood pressure was 180/110 mmHg, and remaining physical examination findings were normal, as were chest X-ray and n electrocardiogram. He had a history of glaucoma and 30 pocket-year cigarette smoking and was administered 2 mg/day doxazosin for hypertension. Contrast agent was not administered. Laboratory test results are summarized in Table 1.

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after discharge, he was found asymptomatic. His blood pressure was 110/70 mmHg, and other physical examinations, including pulse rates, were found to be normal. BUN/serum creatinine levels were normal. Control abdominal computed tomography and serum IgG4 antibody titer were normal, and anti-GBM antibody was negative (Figure 1c). The patient continues to return for routine follow-ups. Informed consent was obtained from the patient.

Discussion

Retroperitoneal fibrosis is a rare condition, with an incidence of 1-2/200,000 individuals (1). There are no specific laboratory tests for IRF. C-reactive protein levels may be high or normal and is often used as an indicator of response to therapy (2). In our patient, there was no diagnostic test result. Erythrocyte sedimentation rate was slightly increased. Only anti-GBM antibody was positive. Radiological imaging is the gold standard for diagnosing abdominal aortic aneurysm and IRF. Ultrasonography may be useful for detecting ureteric obstructions but has a limited value for diagnosis. Computed tomography and magnetic resonance imaging with contrast agents are ideal imaging techniques and are effective in demonstrating the involvement of disease and fibrosis. In our patient, we first determined hydronephrosis using ultrasonography followed by computed tomography. We could not histopathologically confirm the diagnosis. Although there was no presence of hypocomplementemia, we could not rule out IgG4-related diseases.

There was no eosinophilia or eosinophiluria related to acute interstitial nephritis. Retroperitoneal involvement for IgG4-related disease is 20% and is frequently observed as a mass in the retroperitoneal area, lymphadenopathy, or inflammation of vessels, e.g., periaortitis/periarteritis. IgG4-related diseases are present in 55%-57% of patients with IRF. Retroperitoneal involvement in patients with IgG4-related IRF usually begins in the peri-aortic or per-iliac regions and may progress to ureteral involvement. Increasing evidences have shown that noninfectious aortitis is related to IgG4, and this process may also affect iliac/mesenteric arteries. Pathologic findings of tissue samples, obtained by re-evaluating archived and clinical data, were correlated with the IgG4-related disease. Diagnostic criteria include (i) increased levels of IgG4-positive plasma cells as >50 magnification x400 for lungs, (ii) IgG4/IgG ratio of >0.40 or >0.30 with in situ hybridization, and (iii) characteristic histological findings such as lymphocytic infiltrate, spindle fibrosis, and obliteratorive phlebitis in the tissue. Interventional procedures were refused by our patient; hence, we could not explore the potential association among aortitis, retroperitoneal fibrosis, hydronephrosis, and IgG4-related disease.

Table 1. Laboratory data during hospitalization

<table>
<thead>
<tr>
<th>Parameters</th>
<th>Day 1</th>
<th>Day 5</th>
<th>Day 10</th>
<th>Day 18</th>
<th>Month 6</th>
</tr>
</thead>
<tbody>
<tr>
<td>Hematocrit, % (39.5-50.3)</td>
<td>35.7</td>
<td>38</td>
<td>34.6</td>
<td>36.7</td>
<td>41.2</td>
</tr>
<tr>
<td>Hemoglobin, g/dL (16-17.2)</td>
<td>10.9</td>
<td>12.1</td>
<td>11</td>
<td>11.6</td>
<td>14</td>
</tr>
<tr>
<td>White blood cells, /mm$^3$ (4.5-10.3)</td>
<td>7.00</td>
<td>6.30</td>
<td>6.50</td>
<td>8.20</td>
<td>9.80</td>
</tr>
<tr>
<td>Platelets, x10$^3$/μL (156-373)</td>
<td>225</td>
<td>219</td>
<td>193</td>
<td>210</td>
<td>183</td>
</tr>
<tr>
<td>BUN, mg/dL (8-20)</td>
<td>40</td>
<td>37</td>
<td>36</td>
<td>32</td>
<td>14</td>
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<tr>
<td>Creatinine, mg/dL (0.4-1)</td>
<td>2.94</td>
<td>2.43</td>
<td>1.85</td>
<td>1.65</td>
<td>0.93</td>
</tr>
<tr>
<td>Na$^+$, mmol/L (136-144)</td>
<td>134</td>
<td>134</td>
<td>136</td>
<td>137</td>
<td>139</td>
</tr>
<tr>
<td>K$^+$, mmol/L (3.6-5.1)</td>
<td>4.8</td>
<td>4.6</td>
<td>4.5</td>
<td>4.2</td>
<td>3.9</td>
</tr>
<tr>
<td>Ca$^{2+}$, mg/dL (8.9-10.3)</td>
<td>9.2</td>
<td>10.1</td>
<td>9.7</td>
<td>9.2</td>
<td>9.8</td>
</tr>
<tr>
<td>P, mg/dL (2.4-4.7)</td>
<td>4.31</td>
<td>4.0</td>
<td>3.7</td>
<td>4.1</td>
<td>3.6</td>
</tr>
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<td>Uric acid, mg/dL (4.8-8.7)</td>
<td>6.4</td>
<td>-</td>
<td>-</td>
<td>-</td>
<td>5.5</td>
</tr>
<tr>
<td>Total protein, g/dL (6.1-7.9)</td>
<td>6.4</td>
<td>7.6</td>
<td>6.9</td>
<td>7</td>
<td>6.4</td>
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<td>Albumin, mg/dL (3.5-4.8)</td>
<td>3.6</td>
<td>4.3</td>
<td>3.9</td>
<td>4</td>
<td>4.0</td>
</tr>
<tr>
<td>Anti-GBM</td>
<td>++</td>
<td>Negative</td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

BUN: blood urine nitrogen; Na$^+$: sodium; K$^+$: potassium; Ca$^{2+}$: calcium; P: phosphorus; anti-GBM: anti-glomerular basal membrane

Figure 1. a-c. Bilateral hydronephrosis (white arrowheads) and extended ureters (white arrows) (a). Aortic calcification and aortitis (white arrow) (b). Abdominal tomography at 6 months after discharge showed no hydronephrosis and aortitis (c)
Anti-GBM antibody was positive before treatment, but after six months of treatment, anti-GBM antibody was negative, with the regression of hydronephrosis. Takeuchi et al. reported that anti-GBM antibody was positive with hydronephrosis, and with the regression of hydronephrosis, it was negative. They also observed the development of crescentic glomerulonephritis if the positivity for the antibody persisted (5). These findings were similar to those of our case.

IRF-related ureteric obstruction can be treated with steroids, invasive stent, or open ureterolysis. Prednisolone inhibits fibrosis by suppressing inflammation. Our patient had nausea, vomiting, and bilateral hydronephrosis. Despite intravenous fluids and urethral catheter, there was minimal decrease in BUN/serum creatinine levels. Using radiological examinations, abdominal aneurysm and retroperitoneal fibrosis were detected; however, because our patient refused interventional procedures, 32 mg/day oral methylprednisolone was administered. In the following days, BUN/serum creatinine levels rapidly improved. We also radiologically detected an improvement in the process of fibrosis, and the patient’s kidneys were functioning normally 6 months after diagnosis.

The recommended treatment duration is minimum 1 year for IRF. Treatment options include methotrexate, azathioprine, and cyclophosphamide. Immunosuppressive treatments can be used with ureterolysis or invasive stents. Open ureterolysis is also effective, but laparoscopic ureterolysis is a better option for the geriatric population because of its minimal complication rates (6, 7).

Conclusion

Idiopathic retroperitoneal fibrosis is a rare cause of post-renal acute kidney injury. With or without abdominal aortic aneurysm, IRF may cause hydronephrosis and acute kidney injury. Along with abdominal pain, severe hypertension, acute renal dysfunction, hydronephrosis, and abdominal aortic aneurysm, the possibility of retroperitoneal fibrosis should also be considered.

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References