Membranous glomerulonephritis in a patient with syphilis

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To the Editor,

La glomerulonefritis membranosa Membranous glomerulonephritis (MGN) is the second most prevalent renal pathology to be identified in biopsies. One of the most common causes of nephrotic syndrome in the adult population, it is characterised by the formation of immune complexes, predominantly IgG and complement, on the subepithelial side of the glomerular capillaries, and this is associated with increased proteinuria.

In general, its aetiology is idiopathic or primary and, less frequently, secondary (immunological, infectious, drug and medication-related, or neoplastic).

Unfortunately, it is difficult to distinguish primary from secondary forms by historical means; so explicit clinical information, including the age of the patient, history of exposure to medicines or toxic substances, serological tests and suspect neoplasias which are linked to the pathology, is required.

The importance of serological tests lies in their ability to confirm the diagnosis. In the case of syphilis screening, non-treponemal tests are performed: the VDRL (Venereal Disease Research Laboratory) and RPR (rapid plasma reagin) tests. If the results are positive, the more specific treponemal tests are performed to confirm the diagnosis: FTA-ABS (absorption of fluorescent antibodies by Treponema) and MHA-TP (Treponema pallidum microhaemagglutination). They must be repeated three and six months later to ensure the response to treatment.

The case which concerns us is relevant, owing to the small number of publications on the association between syphilis and MGN.

The patient was a 27-year-old, white, Caucasian man with a history of cryptorchidism, adenoidectomy and amygdalectomy in childhood. He was an active smoker, a social drinker and a homosexual. Two months before being assessed by our department and, coinciding with a slight pharyngodynia, an induration had appeared in the patient’s right groin, as well as ulcerated serpiginous lesions on the penis and a whitish urethral discharge, which was initially treated with azithromycin. While waiting for the serological results, maculopapular lesions were observed in the surrounding area on the thighs and trunk. They spread to the patient’s feet and hands, progressing through different phases with no signs of fever, and accompanied by oedema of the lower limbs and genitals, with a slight increase in the abdominal perimeter and a decrease in diuresis, which is why the case was reported to us. The patient’s urine was normal in colour, with no evidence of dysuria or blood in the urine. Blood pressure (BP) was within normal limits.

The analytical findings of note were as follows: urea: 61mg/dl; creatinine: 1.73mg/dl; normal ions; total protein: 4.4g/dl; albumin: 1.8g/dl; total cholesterol: 295mg/dl, HDL: 61mg/dl, LDL: 206mg/dl, triglycerides: 140mg/dl and normal hepatic enzyme levels. Significant findings in the urine analysis included proteinuria: 13.4g at 24h, 250 red blood cells per microlitre and a negative leukocyte count. The haemogram and coagulation were normal, except for an FTP of 762g/l. Autoimmunity assays: anti-nuclear antibodies (ANA) and anti-neutrophil cytoplasmic antibodies (ANCA) negative; complement and protein tests were normal. Serology tests for hepatitis B (HBV), hepatitis C (HCV) and human immunodeficiency (HIV) viruses were negative. Positive 1/32 titre RPR (rapid plasma reagin) and FTA (anti-Treponema antibody) results.

Renal ultrasound showed the kidneys to be normal in size. The echocardiogram was within normal limits and no lung parenchyma changes were detected in the chest X-ray.
Syphilis appears months or years after infection if it has not been properly treated. In developed countries, largely due to the discovery of penicillin, syphilis was practically wiped out in the 1950s. In the 1980s, owing to the concern about the AIDS epidemic, sexual behaviour changed and an even greater decrease in its incidence was observed. In recent years we have been witnessing a resurgence of this disease in Spain, with an increase in its incidence from 2.57 cases per 100 000 inhabitants in 1995 to 5.70 per 100 000 in 2008, and this is also happening in other European countries and the United States. The new cases occur predominantly in young homosexual men and a large portion of them present coinfection with HIV (20%-70%, depending on the area in question). Perhaps this is due to a relaxation in sexual behaviour as a result of a reduction in protective measures following the appearance of highly active antiretroviral therapy (HAART) against HIV.

Although the association between syphilis and renal disease has been known for over 100 years, there are few cases reported in Spain in reviews on the subject, which makes diagnosis more difficult, as it is seldom suspected in clinical practice.

Syphilis can cause a wide variety of clinical and pathological forms of renal disease. In addition to MGN, rapidly progressive GN, diffuse endocapillary GN with or without extracapillary formation or minimal change GN have been described. Proteinuria is the most common clinical manifestation. The definitive diagnosis is confirmed by renal biopsy.

In our case, the patient had been diagnosed with syphilis before and its association with nephropathy facilitated the aetiological diagnosis of MGN. After starting specific treatment (penicillin G benzathine) to eliminate the triggering factor, the nephrotic syndrome remitted.

This experience has made us see that it is of vital importance to conduct a detailed assessment when dealing with a case of nephrotic syndrome. Once we have an exact result and diagnosis, this will enable us to adopt an economic, effective and, above all, curative approach.