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Clinical Case

Acute Lyme Disease in Costa Rica. Description of the first autochthonous case.

Enfermedad de Lyme aguda en Costa Rica.

Descripción del primer caso autóctono

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Abstract

This is the case of a 44-year-old patient, who lives in Santa Teresa, Puntarenas; with acute onset of fever, chills, severe arthralgias and malaise, 15 days after a tick bite. The indirect fluorescent antibody test for Lyme was positive and the patient improved with doxycicline for 14 days. Other infectious and noninfectious diseases were ruled out; we consider this case as the first autochtonous case of Lyme disease in Costa Rica.

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Abbreviations: LD, Lyme Disease; USA, United States of America

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Lyme's disease (LD) is typically a northern hemisphere pathology, particularly of the United States of America (US) and Europe, where it constitutes one the most frequent zoonoses. This distribution obeys principally to the location of the diseases' vector, the *Ixodes* gender tick, also known as hard tick.

LD in humans is associated with acute and chronic manifestations with a variable degree, affecting the osteomuscular, cardiac and nervous systems.

In Costa Rica there are no known autochthonous cases of LD, and the previously seen cases in the country were "imported" cases from the US. We present the first case of autochthonous LD described in Costa Rica.

Case report

A 40 year-old male patient, born in Italy but living in Santa Teresa of Puntarenas for the last 16 years, business man without any past history of illness. The patient's work activities required him to go horseback-riding very often, thus, exposing himself permanently to insect bites, like mosquito and tick bites.

The patient left Costa Rica for the last time 14 months prior to the symptom onset, visiting the US east coast and Milan (Italy) for a 30 day period, not leaving the Santa Teresa and Cóbano regions afterwards.

On early September 2009, he found a tick sticked to his leg's skin for an unknown period of time. He did not remember having any other cutaneous lesions. Fifteen days later he started with chills, fever of up to 39-40 degrees Celsius, malaise, intense joint pain and cramps. After 7 days of persisting symptoms

he consulted at the local clinic where he was prescribed with doxycycline 100 mg twice a day, without a precise diagnosis. Two days after, because of the symptom persistence, he arrived at México Hospital. There, an acutely ill, feverish, without tachycardia, mildly dehydrated patient was found, without pathologic findings at the physical examination.

The peripheral blood smears and serology (IgG and IgM) were negative for ehrlichiosis, serology (IgM) for Rickettsia rickettsii and leptospirosis were negative, as so were the VDRL, HIV ELISA and the **Epstein** monotest. Indirect Barr virus (Scimedex®) immunofluorescence for Borrelia burgdorferi was performed and tested positive for both IgM and IgG. It was controlled three times with a 15 to 22 day interval, showing a progressive decrease in the IgM titer with increase in IgG (figure 1). The patient presented full recovery of clinical signs and symptoms within 5 days of initiating doxycycline, which was prescribed for 14 days. With this information the diagnosis of Lyme Disease or acute borreliosis was made, being this the first autochthonous case in Costa Rica. There were no cardiovascular acute complications, ruled out with a normal electrocardiogram.

Discussion

LD was initially described in Europe, but it was not until the late 70's early 80's (in the city of Old Lyme, Connecticut, USA), that the etiologic agent was discovered to be the *Borrelia burgdorferi* spirochete.

For the USA, the hard tick *Ixodes scapularis* is the vector, *Ixodes ricinus* in Europe and *Ixodes persulcatus* for Asia, which has a 2 year life cycle with four evolutive stages: egg, larva, nymph and adult. Larva feed of blood of small vertebrates like rodents and birds (considered reservoirs for the spirochete),

2009.

Figure 1. Positive indirect immunofluorescense for IgM antibodies against Borrelia burgdorferi. The photo shows the first serum sample from the patient with Lyme disease, processed in the Bacteriology Laboratory of Hospital México in

and in superior stages, they feed from superior mammals like deers and men, with the possibility of transmitting Borrelia. For transmission to be effective the tick has to be adhered to its host at least 36 hours.

There are 12 species of this Gram negative spirochete, forming the Borrelia burgdorferi sensu lato (s.l.) complex, of which Borrelia burgdorferi sensu stricto (s.s.), Borrelia garinii and Borrelia afzelii are the pathogenic species for men. When this microorganism is inoculated, it has a very fast hematogenous dissemination, with predilection for nervous, cardiac, skins and joint tissues. This apparent tissue tropism depends on the infective species, which explains the wide spectrum of manifestations attributed to the disease and the relative geographical predominance of some of them. The disease has three presentations: early localized, early disseminated and late persistent. The most common early form of presentation is the disseminated form that produces signs and symptoms of systemic affection, such as fever, headache, myalgias, arthralgias and malaise, which in 80% of the cases is accompanied by a red papule or spot where the tick bit the patient, which then expands leaving a pale central circle, also known as erythema migrans. These initial symptoms appear after a variable incubation period of some days up to 8 weeks. If the acute presentation lacks systemic symptoms and has only cutaneous manifestations, it is known as early localized.

Early disseminated LD can affect the heart, leading to carditis, which manifests itself most frequently with an atrioventricular block, although other conduction anomalies may occur, like myocarditis, pericarditis and even miocardiopathy in the late persistent form.

Persistent late LD is the chronic form and also shows cutaneous manifestations such as chronic erythema migrans, lymphocytoma cutis and acrodermatitis chronica atroficans. Other chronic manifestations of interest are joint manifestations, presenting generally as big joint mono arthritis, affecting mostly the knee joint, and in some cases

even temporomandibular affection has been reported.

In Europe, neurologic pathology is the most it can lead to acute seen, meningoradiculoneuritis, with involvement of cranial nerves, lymphocytic meningitis and intense radicular pain; also known as Garin-Boujadoux-Bannwarth syndrome. In the chronic phase, it produces meningoencephalitis, cranial nerve involvement, peripheral neuropathy and variable cognitive impairment. It is important to mention that neurologic manifestations are the least frequent amongst the mentioned. Recently, LD has been associated with depression, anxiety, chronic fatigue syndrome and fibromyalgia; all of them predominantly affecting women.

The diagnosis of the early form requires an elevated clinical suspicion, plus a compatible epidemiological history and a suggestive clinical scenario; and in the chronic form the suspicion index is lower, especially in low-incidence countries like Costa Rica. In high-incidence countries the diagnosis can be made only with clinical data, but in our environment, the infection has to be documented with complementary laboratory tests. The spirochete can be isolated from cutaneous lesions and rarely from cerebrospinal or joint fluids with the Barbour-Stoenner-Kelly media culture; but it is a complex process that could last up to 12 weeks. Diagnosis can also be made by detecting nucleic acids with the Polymerase Chain Reaction, although the test is expensive and of variable sensitivity. This explains why the indirect confirmation method of antibody detection is the gold standard, also being less expensive, less complex and with higher sensibility than the other tests mentioned. For this purpose, IgG and IgM antibodies against highly immunogenic bacterial proteins are detected with indirect immunofluorescence or ELISA tests, and then confirmed with a Western Blot, according to international recommendations. In the early phase of LD there could be no IgM antibodies, leading to a false negative, nevertheless, other pathologies like mononucleosis, syphilis, **leptospirosis** and

autoimmune diseases can lead to false positive results because of crossed reactivity of the antigens.

The antibiotic treatment of LD is indicated in every clinical stage of the disease, although it is more effective during the early stages, preferring doxycicline 100 mg twice a day oraly during 14 days. Alternative regimens include amoxicillin for 14 days and azithromycin for 5 days, both in the usual doses. In every case, it is enough to extend the oral regimen for 14 days, except for when there is central nervous system or cardiovascular compromise, in which case the recommendation is to extend the oral administration for 14 to 22 days. In case intravenous therapy is required, sodium penicillin, ceftriaxone or cefotaxime are recommended.

In the case previously presented, the differential diagnoses includes acute febrile pathologies that can lead to joint compromise, but the history of the tick bite guides us into a group of zoonoses that include ehrlichiosis and rickettsiosis, both ruled out.

Other spirochete-borne diseases could generate false positive results because of cross reactivity with the immunofluorescence for LD, like syphilis and leptospirosis, both were ruled out; also, autoimmune pathologies are less likely because of the clinical findings and the patient's evolution. Although the traditional geographic distribution of LD makes it less probable in our country, both the clinical findings

and the laboratory data confirm the diagnosis of the first autochthonous case of LD in Costa Rica. This case must widen the diagnostic horizon of local infectology, in one hand with acute febrile pathologies with an epidemiologic antecedent suggestive of zoonoses, and in the other hand with chronic cardiovascular, neurologic and joint pathologies that lack of a better explanation. The emergence of this type of new diseases must accompany itself with a wide research work that justifies it. Field research must be made to find the vector and the reservoir implied, completing the cycle. It is of our interest to motivate research in this field.

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