LETTERS


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Human Rickettsia sibirica mongolitimonae Infection, Spain

To the Editor: Rickettsia sibirica mongolitimonae has been recently reported as a subspecies of R. sibirica (1). The first evidence of R. sibirica mongolitimonae pathogenicity in humans was documented in France in 1996 (2). Since then, 11 more cases in France, Algeria, South Africa, Greece, and Portugal have been reported (3–6). Because the main clinical manifestations include lymphangitis, the acronym LAR (lymphangitis-associated rickettsiosis) has been proposed (3). We report a case from Spain that confirms the broad distribution of this agent in southern Europe.

A 41-year-old man was admitted on June 19, 2007, to the Hospital de Cruces (Baracaldo, Spain) with fever (39°C), malaise for a week, sweating, lumbar and knee pain, disseminated myalgias, and headache. He reported that 20 days before admission he had removed an engorged tick from his right leg while working as a topographer in the Balmaseda Mountains, 30 km from Bilbao. He had also removed several ticks from his body 4 days before the onset of symptoms. Physical examination did not demonstrate relevant findings. There was no inoculation eschar at the tick-bite sites. Rash, lymphadenopathies, and lymphangitis were not observed.

Chest radiograph did not show consolidation or other abnormality. Initial laboratory examination, on June 21, 2007, showed a leukocyte count 5.2 × 10⁹/μL, hemoglobin 14.1 g/dL, platelet count 190,000/μL, erythrocyte sedimentation rate 9 mm/h, urea 38 mg/dL, creatinine 0.9 mg/dL, aspartate aminotransferase 229 IU/L, alanine aminotransferase 111 IU/L, gamma-glutamyl-transpeptidase 158 IU/L, alkaline phosphatase 170 IU/L, gamma-glutamyl-转肽酶 111 IU/L, total bilirubin 1.3 mg/dL, and C-reactive protein 4.3 mg/dL. Because the patient had been bitten by a tick, acute-phase serum and EDTA-treated blood samples were sent to the Special Pathogens Laboratory (Área de Enfermedades Infecciosas – Hospital San Pedro from La Rioja), where a presumptive diagnosis of rickettsiosis was made. On June 22, 2007, treatment with doxycycline was begun (100 mg/day for 12 days), and his condition rapidly improved.

The early-phase serum yielded low immunoglobulin (Ig) G titer (<64) against Rickettsia conorii and Anaplasma phagocytophilum antigens, and results of ELISA and Western blotting for Lyme borreliosis were negative. A convalescent-phase serum sample collected 7 weeks later did not contain IgG antibodies against spotted fever group Rickettsia species when R. conorii antigen was used.

DNA was extracted from the early whole-blood specimen by using QIAamp DNA Blood minikit (QIAGEN, Hilden, Germany) according to the manufacturer’s instructions. This DNA extract was used as template in nested PCR assays targeting the spotted fever group rickettsial ompB (420 bp) and gltA (337 bp) genes (7). Quality control included both positive (with R. conorii Malish #7 grown in Vero cells) and negative controls that were extracted and PCR amplified in parallel with the specimens. Negative controls consisted of sterile water instead of template DNA. Amplification products of the expected size were obtained. The sequences of these amplicons allowed the identification of R. sibirica mongolitimonae with 99.5% and 100% similarity for ompB and gltA, respectively (GenBank accession nos. DQ097083 and DQ097081).

To our knowledge, Rickettsia species have never been detected in ticks or human specimens in Spain. The host ticks of this rickettsia are likely Hyalomma species, which are more prevalent in southern Spain. In our region in northern Spain, Hyalomma marginatum represented 8% of ticks that fed on humans during 2001–2005, although an increase in this number was recorded last year (data not shown).

In our patient, Rickettsia’s pathogenic role was demonstrated by PCR, a technique that has previously enabled us to identify other arthropod-borne Rickettsia species (8,9). This case suggests that R. sibirica mongolitimonae infection should be considered in the differential diagnosis of rickettsiosis and tick-bite febrile patients in Spain and confirms the distribution of this rickettsia in southern Europe. According to the literature (3), some patients in whom R. sibirica mongoli-
Rickettsia infection is diagnosed have >1 eschar, which raises the suspicion that some cases of Mediterranean spotted fever with multiple eschars reported in Spain could be caused by this rickettsial species. More studies about the vectors of this bacteria are needed because studies of Hyalomma and Rhipicephalus ticks (the suspected hosts) conducted in our area have not demonstrated the presence of this Rickettsia species.

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Lymphangitis in a Portuguese Patient Infected with Rickettsia sibirica

To the Editor: We report a case of Rickettsia sibirica mongolotimonae strain infection associated with lymphangitis (1). A 44-year-old man was admitted to São Bernardo Hospital in Setubal, Portugal, on August 21, 2006. Twelve days previously while on vacation at Troia Peninsula, he noted malaise, insomnia, and dry buccal mucosa. The next day he observed a small erythematous pruritic papulocerebral lesion associated with lymphangitis on the right forearm (Figure). Admission examination showed platelets 117,000/μL, total bilirubin 0.42 mg/dL, albumin 3.42 g/dL, creatinine 1.1 mg/dL, alanine aminotransferase 244 U/L, aspartate aminotransferase 54 U/L, alkaline phosphatase 1061 U/L, creatine phosphokinase 87 U/L, lactate dehydrogenase 784 U/L, C-reactive protein 7.1 mg/dL, radiographic pulmonary diffuse reticular pattern, arterial pO2 68 mm Hg, O2 saturation 94%, pCO2 22 mm Hg, and arterial blood pH 7.35. The differential diagnoses included rickettsiosis, pneumonia, and cellulitis. Treatment with vancomycin, ceftriaxone, and 100 mg of doxycycline twice a day was begun. On the day after hospitalization, a heparinized blood sample and 2 skin biopsy samples were collected. Vancomycin and ceftriaxone were discontinued at 48 hours when rickettsial infection was confirmed by PCR on skin biopsy; 48 hours later, the patient was afebrile.

Immunofluorescence assay for antibodies that used R. sibirica mon- golotimonae strain as antigen demonstrated seroconversion with no antibodies in the serum sample collected on August 21 and immunoglobulin G (IgG) and IgM antibodies at a titer of 1:256 in serum collected on August 20. DNA was extracted from 1 skin biopsy sample by using a DNeasy Tis- sue Kit (QIAGEN, Hilden, Germany). The products of nested PCR showed 100% similarity with gltA (353/353) and ompA (350/350) nucleotide sequences of R. sibirica mongolotimonae strain (GenBank accession nos. DQ423368.1 and DQ423367.1) (1).

Cutaneous biopsy indicated epidermal and dermal necrosis with extensive lymphocyte- and macro-