Probable case of spotted fever group rickettsial infection in a new suspected endemic area, Colombia

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Received 15 June 2016; received in revised form 10 August 2016; accepted 19 August 2016

KEYWORDS
Rickettsia; Rickettsia infections; Spotted fever group; Colombia

Summary Spotted fever group (SFG) rickettsioses are actually considered as emerging and re-emerging zoonotic diseases, caused by pathogenic bacteria of the spotted fever group rickettsiae (SFGR). Recently, serologic studies in human and animals conducted in Colombian Orinoquia, showed a high seroprevalence against SFGR. In June 2015, a 50-year-old male was admitted to a hospital in Bogotá, Colombia, with two days of malaise and temperature of 39°C, associated to generalized rash 24h after the onset of fever. He referred a work visit and outdoor activities in rural area of the Department of Meta 15 days prior the onset of symptoms. The patient was transferred to the intensive care unit with supplementary oxygen, inotropic support and was assessed by the infectious diseases department, indicating the addition of Doxycycline. After seven days of antibiotic treatment the patient was discharged with no evidence of new symptoms or sequels. Retrospectively, two serum samples collected during the acute and convalescent phase were evaluated; there was four fold rise in titer against SFGR. With the foregoing, associated with the
Case report

Spotted fever group (SFG) rickettsioses are actually considered as emerging and re-emerging zoonotic diseases, caused by pathogenic bacteria of the spotted fever group rickettsiae (SFG), transmitted to human beings through tick bites [1]. In Colombia, to date, Rickettsia rickettsii has been the only species of the SFG identified in human cases as well as in Amblyomma patinoi ticks [2,3], being the central region (Department of Cundinamarca) and northwestern Colombia (Departments of Cordoba and Antioquia) known as endemic areas for such rickettsiosis [4–6]. Recently, serologic studies in human and animals conducted in Colombian Orinoquia (Departments of Arauca, Casanare, Guaviare, Meta and Vichada), showed a high seroprevalence against SFGR [7,8].

In June 2015, a 50-year-old male without previous medical history, was admitted to a hospital in Bogotá, Colombia, with two days of malaise and temperature of 39°C, associated to generalized rash 24 h after the onset of fever. He referred a work visit and outdoor activities in rural area of the municipality of Puerto Gaitán (Department of Meta, Eastern center region of Colombia) 15 days prior the onset of symptoms. He denied contact with animals, insect bites or contact with stagnant waters. Physical examination revealed good general condition, temperature of 38°C, heart rate of 95 beats/min, respiratory rate of 18 breaths/min, no conjunctival suffusion, oropharynx with enanthem, no lymphadenopathy, cardio-pulmonary exam without alterations, abdomen without visceromegalies, no meningeval signs, maculopapular rash on the trunk and extremities with involvement of palms and soles (Fig. 1, Panel A and B) and absence of inoculation eschar.

Initial blood analysis showed leukocytosis (13,970 cells/μL), neutrophilia (92.8%), thrombocytopenia (142,000 platelets/μL) and elevated transaminases (AST [99 U/L], ALT [118 U/L]), PCR (16 mg/L) and procalcitonin (2.1 ng/mL). The patient was hospitalized and started Ceftriaxone

Figure 1 Probable case of spotted fever group rickettsial infection in a Colombian patient. Maculopapular rash on the trunk (A) and extremities with involvement of palms (B) and soles.
(2 g/day) after taking blood cultures and serologic tests for Dengue, Chikungunya, HIV, Syphilis (VDRL), CMV, EBV, Toxoplasma and viral hepatitis (A, B and C). On the second day of hospitalization the patient showed clinical deterioration and persistence of fever, associated to onset of dry cough, mild respiratory distress, abdominal pain and petechial lesions in lower limbs. Serology and blood cultures were negative. An increase of leukocytosis (21,540 cells/µL), neutrophilia (97%), transaminases (AST [138 U/L], ALT [134 U/L]), PCR (26 mg/L) and procalcitonin (9.9 ng/mL) was observed, with persistence of thrombocytopenia (98,000 cells/µL). The patient was transferred to the intensive care unit with supplementary oxygen, inotropic support and was assessed by the infectious diseases department, indicating the addition of Doxycycline (100 mg/every 12 h) and additional serologic tests on the suspicion of rickettsial infection. On the third day of hospitalization a biopsy of the skin lesions was performed, with histopathologic findings suggestive of nonspecific lymphohistiocytic vasculitis of small vessels. Seventy-two hours later the patient showed significant improvement in the respiratory pattern, clinical stability and absence of new fever peaks. After seven days of antibiotic treatment with Ceftriaxone plus Doxycycline, the patient was discharged with no evidence of new symptoms or sequels. Retrospectively, two serum samples collected during the acute phase (second day of hospitalization) and convalescent phase (15 days after the first sample) were evaluated in order to determine total antibodies by the Microagglutination Test against 27 pathogenic serovars of *Leptospira* spp. and IgM antibodies (fluorescein isothiocyanate–labeled goat anti-human IgM [Sigma, St. Louis, MO]) by indirect immunofluorescence assay (IFA) against SFGR (*R. rickettsii* antigen). Both serum samples were negative for *Leptospira* spp., however, there was a four-fold rise in titer against SFGR with titers of 1/128 and 1/512 for acute and convalescent phase samples, respectively. Additionally, in order to identify the responsible *Rickettsia* species by molecular methods, a PCR assay was performed for amplification of the gltA rickettsial gene from DNA extracted from paraffin samples of the patient’s skin biopsy (QiAamp DNA FFPE Tissue Kit [Qiagen, CA, USA] was used according to the manufacturer’s instruction); unfortunately the result was negative.

This case presents several features both clinical and paraclinical compatible with probable infection by SFGR [9]: (I) unspecific acute febrile syndrome associated to maculopapular rash with involvement of palms and soles and subsequent presence of purpuric lesions associated with rapid clinical deterioration, with favorable evolution 72 h after the early treatment with Doxycycline; (II) leukocytosis with neutrophilia associated with thrombocytopenia and elevated transaminases; and (III) four fold rise in antibody titer against SFGR in paired samples by IFA test and presence of lymphohistiocytic vasculitis in the histopathologic findings. Regarding the latter, although skin biopsy is considered as an ideal sample for detection and molecular confirmation of the *Rickettsia* species, early treatment with Doxycycline prior to sampling can decrease the sensitivity of molecular methods [9]; situation that can explain the negative results of PCR in our case.

On the other hand, although there have not yet been published studies about molecular characterization of *Rickettsia* species in ticks of Meta department (likely area of the patient’s infection), it was recently described the presence of *Amblyomma mixtum* ticks (known vector of *R. rickettsii* in Central America) in neighboring Departments of Meta [10]. With the foregoing, associated with the recent serological evidence that suggests the circulation of SFGR species in the Colombian Orinoquia [7,8], we consider to recognize this region as a new endemic area for SFG Rickettsioses.

**Funding**

No funding sources.

**Competing interests**

None declared.

**Ethical approval**

Not required.

**References**


