INTRODUCTION

Brazilian spotted fever (BSF) is a rickettsial illness caused by the bacterium *Rickettsia rickettsii*. Because of its high case-fatality rate and apparent increase in areas of transmission, it is considered to be the rickettsial illness of primary public health interest. Cases of this disease have historically occurred in Southeastern Brazil. This article reports the first fatal case of BSF in Southern Brazil. This case highlights the importance of BSF to be considered as a differential diagnosis for acute hemorrhagic fever in areas where cases of BSF may not be expected.

**Keywords:** Rickettsial illness. *Rickettsia rickettsii*. Tick-borne diseases.

CASE REPORT

The patient was a 12-year-old girl from the City of São Carlos de Ivaí, located in the Northern part of Paraná State in Southern Brazil. On January 3, 2015, the patient presented with symptoms of hyporexia and weakness, followed by frontal headaches and a dry cough. She was medicated by family members with paracetamol, dipyrone, and diclofenac. On the seventh day, the patient presented with fever and abdominal pain. At this point, she sought medical attention. She was given paracetamol and discharged. On the ninth day of symptoms, the patient' s condition worsened and she was hospitalized with suspected dengue fever based on her symptoms of fever, myalgia, headache, vomiting, exanthema, and dehydration. Tests to detect the NS1 antigen for dengue were negative, and blood tests revealed the presence of leukopenia and thrombocytopenia. During hospitalization, physicians...
administered only symptomatic medication and parenteral hydration with crystalloids. On the 11th day, the patient presented with exanthema and a reduced level of consciousness, as well as psychomotor restlessness, anuria, worsened thrombocytopenia, and leukocytosis. Antibiotic therapy with ceftriaxone was initiated. The patient was transferred to the intensive care unit (ICU), where she had seizures. At this point, meningitis was suspected. The patient’s condition worsened, and hemodynamic and ventilatory support were initiated. These measures were provided by an ICU in a neighboring city, where the patient died on the 13th day of the illness.

Laboratory investigations included immunologic and molecular analyses of a serum sample collected on January 14 (the 11th day of the patient’s illness) for a diagnosis of spotted fever group rickettsiosis. An immunofluorescence antibody assay using R. rickettsii as the antigen yielded an immunoglobulin G (IgG) concentration of <64 mg/dL and an immunoglobulin M (IgM) concentration of 128 mg/dL. For a molecular analysis, a serum sample was submitted for deoxyribonucleic acid (DNA) extraction using the PureLink® Viral ribonucleic acid/deoxyribonucleic acid (RNA/DNA) Mini Kit (Invitrogen, Carlsbad, CA, USA) and tested with two real-time polymerase chain reaction (PCR) assays: one targeting the rickettsial gltA gene of any member of the Rickettsia family and another specific to R. rickettsii, targeting the hypothetical protein A1G_04230. Results of the patient’s serum DNA were positive for both assays.

In an epidemiological investigation performed post-mortem, it was found that the patient had a history of travel to a farm with a lake inhabited by capybaras (Hydrochoerus hydrochaeris) within Ribeirão Claro municipality, Paraná State. During her last trip, she remained on the farm for 16 days. Her symptoms began the day after she returned. Because capybaras are considered to be the most important source of R. rickettsii infection to A. scapulatum ticks in southeastern Brazil, it is possible that BSF would have been suspected during the first days of illness had the information of the contact with capybaras been shared with the physicians. On the other hand, the physicians’ unfamiliarity with spotted fever rickettsiosis could also have contributed to the lack of BSF suspicious. Finally, the patient’s city of residence (São Carlos do Ivaí) was experiencing a dengue fever epidemic at the time, which could have hampered the suspicious of other vector-borne diseases.

**DISCUSSION**

Brazilian spotted fever has been considered one of the main challenges to public health in Brazil, particularly because of its high case-fatality rates. Factors that may be associated with the low rate of suspicion of BSF include health care professionals’ lack of knowledge on the clinical features of the disease, the eco-epidemic aspects associated with the risk of infection, and the risk factors for exposure to the vector and to the transmission areas of the disease.

The current case reflects the potential for the development of serious forms of the disease with systemic bodily dysfunction, such as respiratory and renal failure, hemorrhagic manifestations, neurological changes, and shock. This report also demonstrates that BSF frequently goes unrecognized as a diagnostic hypothesis and is frequently confused with more prevalent illnesses, such as leptospirosis, dengue fever, and meningococcal disease. In fact, the latter two were considered by physicians in the present case.

Dengue fever is the vector-borne disease with the most substantial impact in Brazil. Recent arbovirus events (which also cause febrile symptoms) represent a significant challenge when it comes to appropriate medical interventions in cases of BSF. The case in question is particularly noteworthy because of two factors that contributed to the outcome: 1) epidemiological history was not considered as part of the patient's clinical history during her treatment process, and so it was not determined that she had been in an area that was eco-epidemiologically associated with the risk of infection because it was a rural area inhabited by capybaras; and 2) no one considered the possibility that the infection developed in a location different from the patient's city of residence that was endemic for the disease and where the patient traveled for leisure activities. If these possibilities had been considered during medical treatment, they would likely have contributed to a greater suspicion of BSF, making the use of appropriate antibiotic treatment more likely and, as a result, reducing the risk of death by BSF.

Current treatment protocols for BSF include doxycycline, an antibiotic indicated for the treatment of R. rickettsii infection and other tick-borne diseases. Indeed, doxycycline is the first choice for treating clinical rickettsioses, regardless of a patient’s age. However, without more knowledge of the disease and the risk factors for infection, and considering the importance of introducing early adequate antibiotic treatment, it is unlikely that the treatment protocol will reduce case-fatality rates. This report therefore provides indispensable information to the search for methods to decrease deaths associated with BSF in Brazil.

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**Conflicts of Interest**

The authors declare that they have no conflicts of interest.

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