Dengue hemorrhagic fever in a peripheral blood stem cell transplant recipient: the first case report

Jirayu Visuthranukul,1,2 Udomsae Bunworasate,3 Panisinee Lawasut,3 Chusana Suankratay,3

1Division of Infectious Diseases, Department of Medicine, Faculty of Medicine, Chulalongkorn University, Bangkok, Thailand; 2Division of Infectious Diseases, Department of Medicine, Faculty of Medicine, Chulalongkorn University, Bangkok, Thailand; 3Division of Hematology, Department of Medicine, Faculty of Medicine, Chulalongkorn University, Bangkok, Thailand

Abstract

Dengue infection, a mosquito-borne infectious disease in tropical and subtropical areas, has recently become an emerging global disease. The clinical course of dengue infection may be unfavorable in immunocompromised patients. In this report, we present a 16-year-old female patient with acute myeloid leukemia who received allogeneic peripheral blood stem cell transplant five months prior to presentation. She was hospitalized at King Chulalongkorn Memorial Hospital, Bangkok, Thailand, due to fever, headache, and myalgia for one day. During hospitalization, she developed capillary leakage syndrome and progressive thrombocytopenia. A diagnosis of dengue hemorrhagic fever was made and confirmed by positive dengue serology and polymerase chain reaction testing. She made a full recovery 14 days after hospitalization. Our case possibly acquired dengue virus from infected mosquitoes while visiting her relatives four days before her present illness. In conclusion, this is the first reported case of dengue hemorrhagic fever in a peripheral blood stem cell transplant recipient. In addition, we review all previous reports of dengue infection in organ transplant recipients.

Introduction

Dengue infection is an acute infectious disease caused by four dengue virus serotypes 1, 2, 3, and 4.1-7 The principal vector is Aedes aegypti, a mosquito with worldwide distribution in many tropical and subtropical areas. The clinical spectrum of dengue infection varies from asymptomatic to severe disease. All serotypes produce a similar clinical illness characterized by acute fever, headache, generalized myalgia, nausea, and vomiting, and induce a life-long immunity that is specific to the infecting serotype.1-7 A small proportion of infected individuals may develop a severe form of disease, dengue hemorrhagic fever (DHF), characterized by fever, thrombocytopenia, hemorrhagic manifestations, and excessive capillary leakage probably leading to dengue shock syndrome (DSS) and death.12 The clinical course of dengue infection may be unfavorable in immunocompromised patients. Bone marrow transplant recipients have an impaired cell-mediated immunity, placing them at increased risk of infections. We report a case of DHF in a peripheral blood stem cell recipient, and review all previous reports of dengue infection in organ transplant recipients.

Case Report

A 16-year-old female was hospitalized at King Chulalongkorn Memorial Hospital, Bangkok, Thailand, due to a high-grade fever without chills, bitemporal headache, generalized myalgia, and nausea one day prior to admission. She had returned four days before from visiting her relatives at Chon Buri, East Thailand. The patient had been diagnosed with acute myeloid leukemia (type M4) 14 months prior to the present illness when she noted acute fever, petechial rash, and bleeding of her gums and nose. She received induction and a consolidation course of chemotherapy. Allogeneic peripheral blood stem cell transplantation was performed five months before the present illness. She was in complete remission when last seen one month prior to her present illness. Complete blood count (CBC) showed hematocrit of 28%, white blood cell count of 8.84×109/L (neutrophil 79%, lymphocyte 5%, atypical lymphocyte 13%, and monocyte 3%), and platelet count of 161×109/L. DHF was suspected, and later confirmed by enzyme-linked immunosorbant assay10 and reverse transcriptase-polymerase chain reaction (PCR) testing.11 A diagnosis of primary dengue infection was made with dengue IgM of more than 40 U and IgM/IgG ratio of or more than 1:81 (dengue IgM rose from 86.65 to 121.03 U, and IgG rose from 49.18 to 134.01 U). Twelve days after hospitalization, she developed convalescent rash over her extremities. She eventually made a full recovery, and was discharged 14 days after hospitalization.

Discussion

This is the first reported case of dengue hemorrhagic fever in a peripheral blood stem cell transplant recipient. Our case had primary dengue infection, and possibly acquired dengue virus from infected mosquitoes while visiting her relatives at Chon Buri. Clinical manifestations of dengue infection in immunocompromised patients are usually similar to those noted in immunocompetent individuals. However, some patients experienced a longer duration (more than seven days) of illness than that in healthy individuals (4-7 days).13 A 23-year-old renal transplant recipient developed dengue hemorrhagic fever which lasted for 19 days.14 Our patient also had a 12-day course of dengue hemorrhagic fever.
In addition, an unusual presentation of dengue infection has been reported.12-13 A 25-year old patient with aplastic anemia with dengue infection developed polyserositis while he underwent conditioning for allogeneic stem cell transplant from his brother.12 Another case of unusual dengue infection was described in a renal transplant recipient who developed acute colitis one week after returning from Southeast Asia.13

Furthermore, a transmission of dengue infection from an organ transplant donor during transplantation has been reported.11,14 One case of possible transmission of dengue infection from a living donor to a renal transplant recipient was described.11 This patient developed acute fever five days after receiving a kidney transplantation from his mother who later had a positive PCR test in the blood for dengue virus serotype 1. Another case of dengue infection transmission was described in a bone marrow transplant recipient during a dengue epidemic in Puerto Rico in 1994.16 A 6-year old recipient developed an acute febrile episode four days after a transplant, and eventually died. Dengue serotype 4 was isolated from the blood and post-mortem tissues. The donor developed a dengue-like illness two days after a transplant. A diagnosis of dengue infection was confirmed by a positive dengue IgM from ELISA testing.

There are a few studies on the histopathology of bone marrow in dengue infection, and most of them are from post-mortem cases with severe dengue infection.15-17 Diffuse hypocellularity with and without hemophagocytosis is always described in bone marrow of patients with dengue hemorrhagic fever. In our case, examination of bone marrow revealed diffuse hypocellularity despite being performed early in the course of dengue infection. This is the first report describing dengue hemorrhagic fever in a peripheral blood stem cell recipient. A high index of suspicion of dengue infection should be given to every transplant patient living or returning from endemic areas who presents with an acute fever in association with viral syndrome.

References