Cyclic thrombocytopenia: A case report

Siklik trombositopeni: Olgu sununu

Abdullah Şumnu1, Reyhan Diz-Küçükkaya2
1Department of Internal Medicine, Istanbul University, Istanbul Medical Faculty, Istanbul, Turkey
2Division of Hematology, Department of Internal Medicine, Istanbul University, Istanbul Medical Faculty, Istanbul, Turkey

Abstract

Cyclic thrombocytopenia (CTP) is a rare disorder characterized by periodic decreases and increases in platelet levels. Each cycle usually spans a period of 3-5 weeks. Clinical features are similar to those of idiopathic thrombocytopenic purpura (ITP), so patients are frequently misdiagnosed as having ITP. However, CTP usually does not respond to most treatments used in ITP such as corticosteroids, splenectomy and intravenous immunoglobulin. In this case report, we present a 33-year-old woman with CTP misdiagnosed as ITP.

Received: November 4, 2008  Accepted: April 8, 2009

Key words: Cyclic thrombocytopenia, idiopathic thrombocytopenic purpura, hepatitis B, case report

Özet


Anahtar kelimeler: Siklik trombositopeni, idiyopatik trombositopenik purpura, hepatit B, olgu sunusu

Geliş tarihi: 4 Kasım 2008  Kabul tarihi: 8 Nisan 2009
Introduction

Cyclic thrombocytopenia (CTP) is a rare disorder characterized by periodic decreases and increases in platelet levels [1]. Each cycle usually spans a period of 3-5 weeks. The platelet counts fluctuate by a large scale on a periodic basis. In the reported cases, the median nadir and peak platelet counts are 10x10^9/L (range 1-90 x10^9/L) and 330x10^9/L (range 72-2300 x10^9/L) [2]. Rebound thrombocytosis (> 500x10^9/L) without therapy is an important and distinctive feature for CTP. Women are affected more than men. Although some cases have been reported in myeloproliferative diseases, most cases are idiopathic [3,4]. The pathophysiology is unclear. Some potential mechanisms are autoimmune platelet destruction, megakaryocytic hypoplasia/aplasia, and infectious and hormonal etiology. Clinical presentation is not different from that of idiopathic thrombocytopenic purpura (ITP). Some patients are asymptomatic. Patients may experience easy bruising, gingival bleeding, recurrent epistaxis, menorrhagia, and hematuria. They may present with more serious bleeding like gastrointestinal or central nervous system hemorrhages. In most cases, CTP is not considered as a differential diagnosis so patients are diagnosed and treated as having ITP. CTP usually does not respond to most treatments used in ITP such as corticosteroids, splenectomy and intravenous immunoglobulin (IVIG). In this condition, a diagnosis of CTP should be considered. Close observation without treatment seem to be the best management if clinically permissible. There are some case reports with complete remission after hormonotherapy (danazol and sex hormones) [5-8].

Case Report

A 33-year-old woman was admitted to the hospital in April 2006 with purpura. Initial complete blood cell count showed thrombocytopenia (platelet count: 6x10^9/L), a mild iron deficiency anemia and normal white blood cell count. After making an initial diagnosis of ITP, the patient received 1 mg/kg/day methylprednisolone for 1 week, 1 g/day methylprednisolone for 3 days and finally 1 g/kg IVIG therapy. There was no response. Then, the platelet count began to increase spontaneously and reached 750x10^9/L in the third week. After 45 days, when she was admitted to the hospital for the second time with purpura, she was administered 2 mg/kg/day methylprednisolone. No response was achieved. The platelet count reached 800x10^9/L again without therapy. Three months later, when the third purpuric attack occurred, splenectomy was suggested, and the patient was admitted to our Hematology Department. Oral informed consent was obtained from the patient.

She was married, with no children, and was working in a hair salon. Her hepatitis B surface antigen was positive. Her physical examination revealed petechiae and ecchymoses on her body and hemorrhagic bullae in the oral mucosa. There was no hepatomegaly, splenomegaly or lymphadenomegaly. Signs of Cushing syndrome were seen and she was still taking 48 mg methylprednisolone per day orally.

Laboratory investigations revealed: platelet: 180x10^9/L, hemoglobin: 10.7 g/dl, hematocrit: 32%, MCV: 74 fl, ferritin: 3 ng/ml, SGOT: 240 IU/ml, SGPT: 280 IU/ml, hepatitis B surface antigen (HBsAg): +, and HBV-DNA: +. Peripheral blood smear showed hypochromia, microcytosis, and normal leukocyte differentiation with no abnormal cells, and the platelets had normal morphology. We tapered and then stopped the corticosteroid dose, administered lamivudine for hepatitis B, and added oral iron therapy for iron deficiency anemia. We followed the patient every week with complete blood count. The
platelet count and also neutrophil and lymphocyte count showed cyclic decreases over a period of five weeks (Figure 1). Anemia as a consequence of thrombocytopenia and menorrhagia contributed to the clinic picture. The diagnosis of CTP was made. Medroxyprogesterone acetate was initiated to control hypermenorrhea after gynecological consultation. Tranexamic acid was also added to control bleeding symptoms when the patient had nadir platelet count. After we accumulated data revealing that the patient had thrombocytopenia every five weeks, we started to use oral contraceptives and tranexamic acid to manage the patient’s menstruation. With this therapy, hypermenorrhea was controlled, and the patient did not experience any major bleeding during the two-year follow-up.

Discussion

In this case report, a 33-year-old woman with CTP misdiagnosed as ITP was presented. She had severe thrombocytopenia with bleeding symptoms followed by thrombocytosis. She received corticosteroids and IVIG therapy with no response. Because she had rebound thrombocytosis, we suspected the diagnosis of CTP. The observation of this patient with weekly platelet counts showed cyclic fluctuation, proving the CTP diagnosis.

The pathophysiology of CTP is unclear. Some potential mechanisms are autoimmune platelet destruction, megakaryocytic hypoplasia/aplasia, and infectious and hormonal etiology. There are some reported cases that respond well to hormonotherapy, especially to danazol. It has been shown that most female CTP patients had low platelet counts during menstruation. It has also been shown that hysterectomy and bilateral salpingo-oophorectomy did not affect platelet fluctuation in those patients [2].

In our case, platelet cycles occurred in synchrony with menstruation (Figure 1). Because our patient had hypermenorrhea causing anemia, we used oral contraceptives for controlling menstruation. Although the oral contraceptive drug did not affect platelet fluctuations, it helped to manage the time of menstruation. We also used tranexamic acid to decrease bleeding symptoms.

The association between CTP and infections has not been described. To date, CTP had been reported only in dogs via *Ehrlichia platys* [9,10] and it was documented to resolve after antibiotherapy [11]. However, there is not yet any data about human ehrlichiosis causing CTP.

Considering that HBsAg in our patient was positive, the association between CTP and hepatitis B needs to be studied. There are some case reports about isolated thrombocytopenia in the course of acute or chronic hepatitis B [12,13]; however, a CTP-hepatitis B association is unknown.

In conclusion, CTP is a rare disorder, but in patients with ITP who have not responded to several treatments and have rebound thrombocytosis, a diagnosis of CTP should be considered.

Conflict of Interest

No author of this paper has a conflict of interest, including specific financial interests, relationships, and/or affiliations relevant to the subject matter or materials included in this manuscript.

References


