Parvovirus infection associated with aplastic crisis in a patient with HEMPAS

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SUMMARY An aplastic crisis associated with parvovirus infection occurred in a patient suffering from hereditary multinuclearity associated with a positive acidified (Hams) test (HEMPAS). This case emphasises that any patient who has a shortened red cell survival is susceptible to an aplastic crisis induced by parvovirus.

In 1981 Serjeant in Jamaica¹ and Pattison in the United Kingdom² established an association between parvovirus infection and aplastic crisis in children suffering from sickle cell anaemia. Similar cases associated with parvovirus infection have been described in subjects with hereditary spherocytosis,³⁴ pyruvate kinase deficiency,⁶ and thalassaemia intermedia.⁷ Aplastic crises in haemolytic anaemias are probably due to a single transmissible agent, the human serum parvovirus.⁸

Between 25% and 70% of adults in Europe and the United States of America have evidence of past parvovirus infection,⁹ the acquisition of the antibody occurring most commonly between 4 and 10 years of age.¹⁰ The most common clinical manifestation of human serum parvovirus infection is erythema infectiousum, a mild illness, the common presenting symptom of which is an erythematous rash; and this is often complicated in adults by arthralgia or arthritis.¹¹¹²

Hereditary erythrocytic multinuclearity associated with a positive acidified serum (Hams) test (HEMPAS) is a rare congenital dyserythropoietic anaemia.¹³ It is characterised by anaemia, episodes of jaundice, and splenomegaly. The bone marrow examination shows severe erythroid hyperplasia, with many of the erythroblasts containing two or more nuclei. The circulating red cells show moderate anisocytosis, anisochromia, and poikilocytosis. Reticulocyte counts are only minimally raised. Red cell life span is reduced to a mean of 18 days¹³ (normal > 26 days), but ineffective erythropoiesis is clearly a major cause of the anaemia.¹⁴

Serological abnormalities consist of an increased susceptibility of the red blood cells to agglutination by anti-i and lysis by anti-i and anti-I. The susceptibility to lysis of the red blood cells by some normal sera acidified to pH 6.8 is a characteristic feature (emphasised in the name of the disorder).

Despite these abnormalities only about 25% of the patients described require transfusion support, most being only mildly anaemic. Aplastic crisis has not previously been associated with HEMPAS.

We report an acute human serum parvovirus infection associated with an abrupt fall in haemoglobin concentration and a lesser fall in white blood cells and platelets in a patient with HEMPAS.

Case history

HEMPAS was diagnosed in a woman aged 20. She had experienced intermittent attacks of jaundice for several years. The diagnosis was based on the characteristic appearance of the marrow, a positive Hams test with two of three normal sera (not her own serum), increased i antigen on her red blood cells, and increased susceptibility to lysis with anti-I. She had a moderately enlarged spleen and in 1984 underwent cholecystectomy for pigment gallstones.

Over eight years of observation her haemoglobin concentration remained between 10 and 12 g/dl without transfusion, except during pregnancy when transfusions were required. The white cell and platelet count were consistently within the normal range. Her reticulocyte count ranged from 2% to 4%. Her red cell life span (T½ ⁵¹Cr) was estimated on four separate occasions, giving results of 11, 12, 15, and 19 days (normal > 26 days).

In July of 1985 she was admitted to hospital as an
emergency. She had suffered abdominal pain, nausea, vomiting and diarrhoea for five days. She complained of progressive weakness.

On admission she had a fever of 38°C, her haemoglobin was 5·3 g/dl, platelets 86 × 10⁹/ℓ, and white cell count 1·7 × 10⁹/ℓ. Reticulocytes were absent. Her bone marrow was mildly hypocellular with left shift of erythroid precursors and a dearth of multinuclear cells, in sharp contrast to previous examinations of her marrow, which had shown severe hypercellularity, with increased erythropoiesis and 30% of the erythroblasts being multinuclear.

She was transfused and her fever settled. Reticulocytes appeared in her peripheral blood reaching 5% six days after admission. Her haemoglobin remained stable after this episode, and her platelet and white cell count returned to normal values.

Serum taken six and 31 days after admission was examined for human serum parvovirus specific IgM, according to the method of Cohen et al. Throat washings and urine collected nine days after admission were examined for the presence of human serum parvovirus DNA by dot blot DNA hybridisation using the method of Anderson et al.

The serum samples contained 30 and 34 arbitrary units of antihuman parvovirus specific IgM and IgG antibody, respectively, six days after admission and 35 IgM and 26 IgG arbitrary units at 31 days after admission, indicating recent infection. Human serum parvovirus DNA was found neither in the urine or throat washings collected nine days after admission, nor in either serum sample.

Discussion

It seems likely that human serum parvovirus infection caused an aplastic crisis in this patient. Aplastic crises have been reported in both acquired and hereditary haemolytic anaemias. These have in common a shortened red cell survival so that aplasia in the bone marrow is rapidly reflected by a fall in peripheral haemoglobin concentration. It has been suggested that although there may be a direct infection of the erythroid precursors with the virus, aplasia of short duration (seven to 10 days) would pass unnoticed unless there were also shortened red cell survival. Our case supports this hypothesis, because although a major component of the anaemia of HEMPAS is caused by ineffective erythropoiesis, our patient also had a shortened red cell survival.

As far as we know this has not been reported before in association with HEMPAS due to the rarity of this condition, and our case emphasises that anyone who has a shortened red cell survival may be susceptible to an aplastic crisis induced by parvovirus.

References


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