Thrombotic thrombocytopenic purpura associated with pneumococcal sepsis

JEFFREY R SCHRIBER, MD, JOHN J FREEDMAN, MD, JOSEPH M BRANDWEIN, MD

JR SCHRIBER, JJ FREEDMAN, JM BRANDWEIN. Thrombotic thrombocytopenic purpura associated with pneumococcal sepsis. Can J Infect Dis 1993;4(3):145-147. The first documented case of thrombotic thrombocytopenic purpura (TTP) associated with pneumococcal septicemia is reported. This association has been previously demonstrated with hemolytic uremic syndrome. The patient presented with recurrent seizures, oliguric renal failure, fever, thrombocytopenia and microangiopathic hemolytic anemia; coagulation studies were normal. Blood and sputum cultures were positive for Streptococcus pneumoniae. The patient responded to therapy with plasmapheresis and antiplatelet agents as well as antibiotics. Coincident infection should be searched for in all cases of TTP.

Key Words: Plasmapheresis, Pneumococcal infections, Thrombotic thrombocytopenic purpura

Purpura thrombocytopénique thrombotique associé à une septicémie pneumococcique

RÉSUMÉ: Le premier cas documenté de purpura thrombocytopénique thrombotique associé à une septicémie pneumococcique a été rapporté. Cette association avait déjà été observée dans un syndrome urémique hémolytique. Le patient s'est présenté avec convulsions récurrentes, insuffisance rénale marquée par de l'oligurie, de la fièvre, de la thrombocytopénie, et anémie hémolytique microangiopathique. Les épreuves hémostatiques étaient normales. Les cultures de sang et d'expectorations étaient positives à l'égard de Streptococcus pneumoniae. Le patient a bien répondu au traitement par plasmaphérèse et agent antiplaquettaire de même qu'aux antibiotiques. Il faut toujours considérer la possibilité d'une infection concomitante dans tous les cas de purpura thrombocytopénique thrombotique.

Thrombotic thrombocytopenic purpura (TTP), initially described by Moschovitz (1), is characterized by the presence of consumptive thrombocytopenia, microangiopathic hemolytic anemia and neurological symptoms, frequently accompanied by renal failure and fever (2). The pathogenesis of this disease is unclear, although abnormalities of prostacycline and von Willebrand's factor have been described (2,3).

Various infectious agents have been implicated in the etiology of this disorder as well as in the related hemolytic uremic syndrome (4,5). Hemolytic uremic syndrome has been described in association with pneumococcal septicemia (6-10), including one adult (7); we report the first case of full-blown adult TTP in association with this infection.

CASE PRESENTATION

A 28-year-old woman with an uneventful past medical history and a one-week history of nausea, vomiting and diarrhea presented to the emergency room follow-
Antinuclear antibody, rheumatoid factor and was admitted to the intensive care unit and treated with x-ray body with peanut lectin and red cell antibody screen were negative. Red cells formed to rule out associated pregnancy, was negative. Eclampsia, with associated renal failure, fever and thrombocytopenia (12-10), although associations with Shigella dysenteriae (12) and enteroviruses (13) have been described. Of the 13 patients previously reported to have S pneumoniae infections in association with hemolytic uremic syndrome, nine presented with pneumonia, two with meningitis and two with sepsis of undetermined origin.

All patients required dialysis and three received plasma infusion or plasma exchange. Twelve of the
reported cases have been young children (age range five to 27 months). In these children there was a 50% mortality, whereas for hemolytic uremic syndrome in general the mortality is in the 5 to 10% range (4). Institution of appropriate antibiotics in the patients appeared to have been prompt, although in two fatal cases antibiotic use was not mentioned (10). The one adult previously reported had presented with fever, renal failure requiring hemodialysis and microangiopathy with thrombocytopenia (7). However, that patient, unlike ours, also had serological features suggestive of disseminated intravascular coagulation, including elevated fibrin degradation products and partial thromboplastin time, thus complicating the diagnosis.

_S. pneumoniae_ may cause hemolytic uremic syndrome via the action of circulating neuraminidase produced by this organism (8,10). Removal of n-acetylneuraminic acid from cell surface glycoprotein by neuraminidase exposes the normally hidden T-antigen (Thomsen-Friedenreich) present on red blood cells, platelets and glomerular membranes. This results in deposition of naturally occurring IgM anti-T with subsequent injury to blood cells and glomeruli (8). The presence of exposed T-antigen can be demonstrated by agglutination using peanut lectin. Of the 13 patients reported, 10 had evidence for exposed T-antigen using this method; this was not mentioned in the one adult reported (7). In our patient we were unable to demonstrate the presence of exposed T-antigen on red blood cells. Whether this indicates another mechanism for pneumococcus-induced thrombotic microangiopathy in the present case, or that the pneumococcal septicaemia was not an etiologic factor in this case, is unclear.

The treatment of choice for TTP is plasmapheresis or plasma exchange; antiplatelet agents may also be of benefit (2). Because of the small number of cases and the different varieties of therapy employed, it is difficult to determine the ideal treatment for _S. pneumoniae_-associated TTP/hemolytic uremic syndrome. The early use of aggressive plasmapheresis with antiplatelet agents, in addition to antibiotic treatment for the underlying infection, was effective therapy in this patient. This case also emphasizes the importance of searching for an underlying infectious cause in patients with TTP.

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**REFERENCES**
