

LETTERS TO THE EDITOR

Cytomegalovirus gastritis after rituximab treatment in a non-Hodgkın's lymphoma patient

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TO THE EDITOR

Rituximab is a chimeric monoclonal antibody against CD20 antigen expressed on most B cells and is used for the treatment of malignant lymphomas expressing the CD20 antigen^[1]. It temporarily eliminates normal B-lymphocytes without a substantial decrease in serum immunoglobulin levels^[2]. Recently, several serious viral infections have been reported in association with rituximab use. Cytomegalovirus (CMV) is one of the agents that may be a cause of morbidity and mortality in immunocompromised individuals^[2, 3]. In the face of exogenous or endogenous causes of immunosuppression, cytomegalovirus can result in retinitis, colitis, pneumonitis, or encephalitis. Presented here is a case of a patient with non-Hodgkin's lymphoma (NHL) who developed gastritis and enterocolitis due to CMV infection after treatment with rituximab.

A 65-year old female patient was admitted with complaints of epigastric pain and diarrhea. Prior to her admission she was treated for diffuse large cell type NHL, stage IIIB bulky disease with six courses of a regimen consisting of cyclophosphamide, doxorubicin, vincristine and prednisone (CHOP) and rituximab, and a good partial response was achieved. Due to the presence of bulky disease at presentation, she also received radiotherapy (RT) to the parailiac and inguinal regions at a dose of 3000 centigray

(cGy), one month after chemotherapy. Two weeks after radiotherapy, she presented with epigastric pain and diarrhea without blood and mucous. Microscopic examination of stool specimens was unremarkable. Her blood and stool cultures were negative. Her chest X-ray was also normal. Initial investigations revealed normal hemoglobin concentration and platelet count as well as mild leucopenia. The number of white blood cells was $3.6 \times 10^9 / L$ (3.0× 10⁹/L neutrophils). Serum levels of glucose, amylase, liver transaminases, urea and electrolytes were normal. Based on the preliminary diagnosis of radiation enteritis, antidiarrheal therapy was initiated. The patient did not benefit from this, and her epigastric pain aggravated. A computerized tomography scan of the abdomen revealed gastric wall thickening as well as findings consistent with colitis of the ascending and transverse colon. Subsequent upper gastrointestinal endoscopic evaluation demonstrated multiple linear exudative gastric ulcers, the largest being 4 cm in diameter, from which multiple biopsy specimens were obtained. Histological examination revealed hyperchromatic epithelial cells with nuclear viral inclusions (owl's eye). These cells were positive for monoclonal antibodies against CMV. However, peripheral blood CMV viral load was negative. She failed to respond to acid suppression therapy and her clinical condition deteriorated further. Repeat upper gastrointestinal endoscopy and histological examination of biopsy specimens were consistent with the previous results. Consequently, treatment with intravenous gancyclovir (2×5 mg/kg) was initiated with a diagnosis of CMV gastritis and enterocolitis. Dramatic clinical improvement was achieved within one week and treatment was continued for 3 weeks. Her symptoms resolved completely and a subsequent endoscopy only showed signs of healing ulcers.

Pelvic or abdominal RT is a known cause of acute enteritis characterized by abdominal cramping and diarrhea in approximately 50% of treated patients, and the incidence is higher with concomitant chemotherapy^[4]. Acute intestinal side effects of RT occur at doses of approximately 1000 cGy. Since the curative doses for most abdominal or pelvic tumors range between 5 000 and 7 000 cGy, enteritis is more often than not likely to occur^[5]. Our patient received 3000cGy of RT and probably developed RT-induced enteritis. However, antidiarrheal therapy with oral opiates failed to alleviate the patient's complaints, which only resolved after anti-CMV therapy.

Cytomegalovirus enterocolitis should be considered in the differential diagnosis in patients receiving rituximab whose symptoms fail to resolve after conventional antidiarrheal therapy, irrespective of peripheral blood CMV viral load.

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