Celiac disease-like enteropathy due to antihypertensive therapy with the angiotensin-ll receptor type 1 inhibitor eprosartan

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SUMMARY

An 83-year-old woman with hypertension received the angiotensin-II receptor type 1 blocker (ARB) eprosartan for more than 10 years. Six months ago, the dosage of the drug was doubled, and the patient reported a sudden onset of diarrhea. Duodenal biopsies showed a celiac disease-like pathology with flattened mucosa and an increase of intraepithelial lymphocytes and eosinophils, but serology of celiac disease remained negative. Celiac disease-like changes have been previously reported to be associated with other ARBs. This is the first case following eprosartan medication. In celiac-disease-like pathology of the small bowel with negative serology, drug-induced changes, for example due to ARBs, should be excluded.

Keywords: celiac disease – negative serology – ARB – eprosartan

Enteropatie podobná celiakii způsobená antihypertenzivní terapií inhibitorem receptoru typu 1 angiotenzinu-II – eprosartanem

SOUHRN

83leté pacientka s hypertenzí byla více než 10 let léčena inhibitorem receptoru typu 1 angiotenzinu-II – eprosartanem. Před šesti měsíci byla dávka léků zdvojnásobena a u pacientky došlo k náhlému vzniku průjmu. Biopsie duodena prokázala změny podobné celikakii s oploštěním sliznice a zmnožením intraepiteliálních lymfocytů a eozinofilů. Sérologická vyšetření na celiakii však byla negativní. Změny podobné celikakii byly už dříve spojeny s léčbou inhibitory receptoru typu 1 angiotenzinu-II. Tento případ poprvé vznikl při léčbě eprosartanem. Při patologickém nálezu v tenkém střevu podobnému celiakii s negativní sérologií by měly být vyloučeny terapií indukované změny, např. terapií inhibitorem receptoru typu 1 angiotenzinu-II.

Klíčová slova: celiakie – negativní erologie – ACE inhibitory – eprosartan

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Duodenal changes caused by drugs mimicking the classical pathology of coeliac disease are a well-known pitfall in pathology. They have been recently described in patients treated with olmesartan, a newer member of angiotensin-II receptor type 1 inhibitors (ARBs) with a prolonged degradation time and higher effect on hypertension. It has been documented in one study on 22 patients (1) and two case reports (2,3). We report here similar changes during therapy with eprosartan, another drug of the ARB family.

CLINICAL HISTORY AND METHODS

An 83-year-old woman reported a sudden onset of diarrhea. A gastroscopy was performed, and biopsies from the duodenum were taken. After fixation with 4% buffered formaldehyde and embedding in paraffin, slices of 5 µm thickness were stained

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Hans Maier M.D. Institute of Pathology, Klinikum Bayreuth, Preuschwitzer Strasse 101, D-95445 Bayreuth, Germany tel.: +49 921 400 5602, fax: +49 921 400 5609 e-mail: hans.maier@klinikum-bayreuth.de with hematoxylin/eosin (H&E). Histology revealed a severe attenuation of the duodenal villi (Fig. 1). The amount of intraepithelial lymphocytes was over 25/100 epithelial cells, accompanied by eosinophilic granulocytes (Fig. 2). Subepithelial collagen deposits were not noticed. No cells with expression of IgG4 were present.

The serology for celiac disease was negative in our patient. Our inquiry of more detailed information on the clinical history and on the possible use of medications disclosed a long-standing history of systemic hypertension and an antihypertensive therapy with eprosartan for the last ten years. Six months ago, the daily dosage of 300 mg (half of a tablet) had been doubled to 600mg, apparently causing the gastrointestinal symptoms. There was no history of celiac disease in previous biopsies. After our diagnosis of eprosartan-induced diarrhea and concomitant mucosal changes of the small bowel, eprosartan was replaced through an application of the calcium channel blocker amlodipin with cessation of diarrhea. Control biopsies three and six months after the diagnosis showed only slightly improved mucosal architecture, indicating a delayed regeneration of the duodenal mucosa after application of ARBs like eprosartan.

DISCUSSION

Eprosartan belongs to the first generation of ARBs recommended for the therapy of hypertension, chronic heart failure,