Eur J Pediatr (1998) 157: 693–694

23

M. Maghnie

Lymphocytic hypophysitis and central diabetes insipidus during adolescence: what are the criteria for diagnosis?

Received: 2 December 1997
Accepted: 9 February 1998

Sir: We read with interest the paper of Cemeroglu et al [1] describing a 14-year-old girl with acute central diabetes insipidus (DI), secondary amenorrhoea and normal neurological and visual examinations. MRI of the brain revealed a $10 \times 10 \times 10$ mm mass involving the pituitary stalk and hypothalamus which spontaneously reduced within 3 months and returned to $10 \times 8 \times 10$ mm 3 months later. At 6 months the T1-weighted posterior pituitary hyperintensity was absent (no mention of the posterior pituitary status on initial MRI). Endocrine evaluation revealed hypogonadotropic hypogonadism and low insulin-like-growth-factor-I (IGF-I) level. Histological examination showed signs of chronic inflammation with a predominant lymphocytic infiltrate compatible with lymphocytic hypophysitis.

We wish to make some comments and suggestions:

We have described an 8-year-old girl with acute onset central DI and acquired growth hormone (GH) insufficiency in whom the first MRI showed a thick pituitary stalk and undetectable posterior pituitary hyperintensity [2]. Serial MRI studies were unchanged for 5 years when a huge mass involving the pituitary stalk and hypothalamus was documented togeher with clinical and laboratory features of panhypopituitarism. Histopathology revealed perivascular inflammatory lymphoplasmatic infiltrates with absence of granulomatosis and necrosis and negative staining for S-100 protein. The patient was treated with high dose prednisolone (30 mg/kg per day, total dose of 2.4 g in 20 min infusion for 3 days). MRI performed 1 month later showed an approximately 50% decrease in the mass, with partial anterior pituitary recovery (thyroid and adrenal) maintained for 2 years after treatment began.

In our opinion, the features reported by Cemeroglu et al. [1] do not permit a convincing diagnosis of classical lymphocytic hypophysitis as traditionally conceived: the disease onset is unrelated to pregnancy, the posterior pituitary is involved, the mass on coronal MRI is confined to the pituitary stalk and hypothalamus while the anterior pituitary is spared, there is no evidence of lymphocyte infiltration and/or destruction of the anterior pituitary tissue and no other co-existing auto-immune disorders are associated [3]. We believe that the disease reported by Cemeroglu et al. [1] belong to a unique spectrum of inflammatory auto-immune vascular-mediated conditions variably affecting the hypothalamic-pituitary area. The statement that "the loss of the normal posterior pituitary T1-weighted hyperintensity may have been a clue to the diagnosis of lymphocytic hypophysitis before biopsy in this case" is rather misleading because the lack of posterior pituitary hyperintensity in central DI is a non-specific hallmark of a hypothalamic-neurohypophyseal axis lesion [2, 4].

The conservative management of these tumour-like conditions appears reasonable but in the patient reported by Cemeroglu et al., the chronic growth pattern and the size of the mass as well as the documentation of an inflammatory process after pituitary stalk biopsy require in our opinion a tentative treatment approach. Only few controversial data with different treatment modalities and outcome have been reported but the favourable response to glucocorticoids in our patient underlines the possible role of steroids in the management of such inflammatory masses. We believe that the low IGF-I level and patient weight increase (6.8 kg.) may have been due to acquired GH insufficiency which is frequently associated with such lesions. Thus, evaluation of GH secretion and clinical indication for GH treatment (metabolic and quality of life effects in adults) in case of GH deficiency merit consideration.

References

1. Cemeroglu AP, Blaivas M, Muraszko KM, Robertson PL, Vàzquez DM (1997) Lymphocytic hypophysitis presenting with diabetes insipidus in a 14-year-old-girl: case report and review of the literature. Eur J Pediatr 156:684–688
2. Maghnie M, Villa A, Aricò M, Larizza D, Pezzotta S, Beluffi G, Genovese E, Severi F (1992) Correlation between magnetic resonance imaging of posterior pituitary and neurohypophyseal function in children with diabetes insipidus. J Clin Endocrinol Metab 74:795–800
3. Ezzat S, Josse RG (1997) Autoimmune hypophysitis. Trends Endocrinol Metab 8:74–80
4. Maghnie M, Sommaruga MG, Beluffi G, Severi F (1993) Role of MR imaging in the evaluation of the functional status of the posterior pituitary gland: the view of a pediatric endocrinologist. Am J Neuroradiol 14:1443–1445

M. Maghnie
Department of Paediatrics,
University of Pavia,
IRCCS Policlinico S. Matteo,
I-27100 Pavia, Italy,
Fax: +39-382-527976

D.M. Vázquez
A. Pinar Cemeroglu

Reply

Received: 24 January 1998
Accepted: 9 February 1998

Sir: We are pleased to have the opportunity to update our report on each of the issues addressed in Dr. Maghnie's letter.

In the last few years, the broad spectrum of presentation of the condition known as lymphocytic hypophysitis (LYHY) has been established. Thus, LYHY can affect men and women and need not be related to pregnancy [1, 4, 5, 9, 10]. LYHY is not only confined to the anterior pituitary but can involve the posterior pituitary and the stalk [2, 14]. The term LYHY is used globally to describe a spectrum of pathology. Based on anatomical site and severity of the inflammatory process, LYHY can be subclassified as lymphocytic hypophysitis, lymphocytic infundibulo-neurohypophysitis and necrotizing infundibulo-hypophysitis. We elected to use the global term of LYHY in our report. We absolutely agree that the imaging studies performed in our patient point to pituitary stalk and hypothalamic involvement. As stated by Dr. Maghnie, this does not fit the classical diagnosis of lymphocytic hypophysitis strictly defined by anatomical features but, again, LYHY was used in the broader sense which encompases the whole spectrum of this condition. Because of the age of our patient, we did not originally