COMMENTARY: THE INTERNATIONAL FACE OF CELIAC DISEASE

Ravi N Sharaf MD

Elizabeth Verna ND

Peter HR Green MD

Columbia University, New York, NY

Correspondence: Peter HR Green MD

180 Fort Washington Ave

New York, NY

USA 10032

Celiac disease is a genetically determined chronic inflammatory intestinal disorder induced by an environmental precipitant, gluten(1). Patients with celiac disease are mount an immunologic

response to gluten, the main storage protein of wheat and similar proteins found in rye and barley. While much has been learnt recently about the genetic influence as well as the role of environmental factors, there are still many unsolved questions. The genetic influence is demonstrated by the ~10% occurrence in first degree relatives(2) and 70% concordance among identical twins(3). The HLA alleles that determine the presence of DQ2 or DQ8 are required for the disease. They however account for less than 50% of the genetic influence. Other genes have yet to be conclusively identified.

The environmental factors that are important for childhood celiac disease include both the amount of gluten given to children and the timing of gluten ingestion, as well as infections in childhood(4). The peak age of diagnosis in adults is in the fourth and fifth decades(5); however, factors that influence the age of onset in adults are unknown.

Formally considered a rare disease of childhood, limited to populations of European descent, recent studies have demonstrated the disease to be very common, affecting individuals of all ages from many diverse regions of the world. Several studies have demonstrated a prevalence of 1% among different European populations including Finnish school children(6), adults(7) and children in the United Kingdom(8).

The typical "face" of celiac disease is however changing. An example of this is recorded by Cataldo et al their paper describing celiac disease among immigrant children in Italy(). This highlights the international nature of celiac disease. They report cases of celiac disease in children from East Europe, Northern, West and East Africa, Southern Asia, and the Middle East. The disease is common in several countries from these regions of North Africa(9-12), the Middle East(13-15), and South Asia(16-18). Whites in South Africa are, not unexpectantly, diagnosed with celiac disease(19), though the report by Kaven includes a Zulu patient with celiac disease(19).

The potential for celiac disease to become a major health issue in developing countries is highlighted by the report of the high rate of celiac disease in a population of refugee children in North Africa. In this study Catassi et al found 5.6% of Saharawi children to have a positive endomysial antibody(9). The endomysial antibody has a virtual 100% specificity for celiac disease. This certainly raises important issues for the health of these refugee populations because the major grain supplied to refugee populations is wheat that is produced in excess by developed countries.

What about celiac disease in the Americas? Celiac disease is reported from Brazil(20), Argentina(21) and Chile, including native South American Indians in Chile(22). While there are no reports from Central America, there are reports from the Cuba(23, 24) and black children, of West Indian origin, resident in the United Kingdom(25).

In North America celiac disease is considered a rare disease by most practitioners, though the rate of diagnosis is increasing(26). Patients have a long duration of symptoms prior to diagnosis and see many physicians(5). Hopefully the awareness of the recent serologic studies demonstrating prevalence rates similar to Europe(27) and an NIH Consensus conference on celiac disease, planned for June 2004, will influence current medical practice and result in greater diagnostic awareness in the United States.

Currently there are few minorities diagnosed with celiac disease in the United States, though we have seen celiac disease among African Americans, people born in Cuba, Puerto Rico, the Dominican Republic and Asians from China and the Philippines. There is a low rate of diagnosis of celiac disease among minority populations in the United States because of lower frequency of HLA DQ2 or DQ8 and problems with access to health care services.

There are no studies that have specifically screened African Americans, Hispanic or native American populations in the United States. It is of interest that several studies in the United States have referred to celiac disease among non-whites and specifically African Americans. These include a blood donor serologic screening study(27, 28), a study of United States military veterans with celiac disease in which 7% were "non-white"(29) and in a study of people with iron deficiency(30) . A recent study from Canada described celiac disease among Asian-Canadians with origins from Northern India, Japan and China(31). In addition Freeman reported celiac disease in a North American native person(32).

The use of serologic screening tests has resulted in an increase in the rate of diagnosis of celiac disease in many diverse populations around the world, including groups not traditionally

3

considered to have celiac disease. Celiac disease is seen in many patients from developing countries, both in their native lands and as immigrants in developed countries. This has important implications for medical practitioners throughout the world as well as for authorities that will need to address the supply of gluten-free diets to those diagnosed with celiac disease.

REFERENCES

- 1. **Green PH, Jabri B.** Coeliac disease. *Lancet.* 2003;362(9381):383-91.
- 2. Bevan S, Popat S, Braegger CP, et al. Contribution of the MHC region to the familial risk of coeliac disease. *J Med Genet.* 1999;36(9):687-90.
- 3. **Greco L, Romino R, Coto I, et al.** The first large population based twin study of coeliac disease. *Gut.* 2002;50(5):624-8.
- Ivarsson A, Persson LA, Nystrom L, et al. Epidemic of coeliac disease in Swedish children. Acta Paediatr. 2000;89(2):165-71.
- 5. **Green PHR, Stavropoulos SN, Panagi SG, et al.** Characteristics of adult celiac disease in the USA: results of a national survey. *Am J Gastroenterol.* 2001;96(1):126-31.
- Maki M, Mustalahti K, Kokkonen J, et al. Prevalence of Celiac disease among children in Finland. N Engl J Med. 2003;348(25):2517-24.
- West J, Logan RF, Hill PG, et al. Seroprevalence, correlates, and characteristics of undetected coeliac disease in England. *Gut.* 2003;52(7):960-5.

- Bingley PJ, Williams AJ, Norcross AJ, et al. Undiagnosed coeliac disease at age seven: population based prospective birth cohort study. *Bmj.* 2004;328(7435):322-3.
- Catassi C, Ratsch IM, Gandolfi L, et al. Why is coeliac disease endemic in the people of the Sahara? *Lancet*. 1999;354(9179):647-8.
- Ashabani A, Abushofa U, Abusrewill S, Abdelazez M, Tuckova L, Tlaskalova-Hogenova H. The prevalence of coeliac disease in Libyan children with type 1 diabetes mellitus. *Diabetes Metab Res Rev.* 2003;19(1):69-75.
- Boudraa G, Hachelaf W, Benbouabdellah M, Belkadi M, Benmansour FZ, Touhami
 M. Prevalence of coeliac disease in diabetic children and their first- degree relatives in west Algeria: screening with serological markers. *Acta Paediatr Suppl.* 1996;412:58-60.
- al-Tawaty Al, Elbargathy SM. Coeliac disease in north-eastern Libya. Ann Trop Paediatr. 1998;18(1):27-30.
- al-Hassany M. Coeliac disease in Iraqi children. J Trop Pediatr Environ Child Health. 1975;21(4):178-9.
- Shahbazkhani B, Malekzadeh R, Sotoudeh M, et al. High prevalence of coeliac disease in apparently healthy Iranian blood donors. *Eur J Gastroenterol Hepatol.* 2003;15(5):475-8.
- 15. Rawashdeh MO, Khalil B, Raweily E. Celiac disease in Arabs. *J Pediatr Gastroenterol Nutr.* 1996;23(4):415-8.
- Sood A, Midha V, Sood N, Malhotra V. Adult celiac disease in northern India. Indian J Gastroenterol. 2003;22(4):124-6.
- Sood A, Midha V, Sood N, Kaushal V, Puri H. Increasing incidence of celiac disease in India. *Am J Gastroenterol*. 2001;96(9):2804-5.
- Yachha S MSSA, Krishnani N, Saxena A. Effects of gluten-free diet on growth and small bowel hitology in children with celiac disease in India. *J Pediat Gastroenterol Nutr*. 2000;31(S23).
- Kavin H. Adult coeliac disease in South Africa. An analysis of 20 cases emphasizing atypical presentations. S Afr Med J. 1981;59(18):628-32.

- 20. Gandolfi L, Pratesi R, Cordoba JC, Tauil PL, Gasparin M, Catassi C. Prevalence of celiac disease among blood donors in Brazil. *Am J Gastroenterol.* 2000;95(3):689-92.
- Gomez JC, Selvaggio GS, Viola M, et al. Prevalence of celiac disease in Argentina: screening of an adult population in the La Plata area. *Am J Gastroenterol.* 2001;96(9):2700-4.
- 22. Araya M, Mondragon A, Perez-Bravo F, et al. Celiac disease in a Chilean population carrying Amerindian traits. *J Pediatr Gastroenterol Nutr*. 2000;31(4):381-6.
- Sagaro E, Jimenez N. Family studies of coeliac disease in Cuba. Arch Dis Child. 1981;56(2):132-3.
- Rabassa EB, Sagaro E, Fragoso T, Castaneda C, Gra B. Coeliac disease in Cuban children. Arch Dis Child. 1981;56(2):128-31.
- Hung JC, Phillips AD, Walker-Smith JA. Coeliac disease in children of West Indian origin. Arch Dis Child. 1995;73(2):166-7.
- Murray JA, Van Dyke C, Plevak MF, Dierkhising RA, Zinsmeister AR, Melton LJ,
 3rd. Trends in the identification and clinical features of celiac disease in a North
 American community, 1950-2001. *Clin Gastroenterol Hepatol.* 2003;1(1):19-27.
- Fasano A, Berti I, Gerarduzzi T, et al. Prevalence of celiac disease in at-risk and not-atrisk groups in the United States: a large multicenter study. *Arch Intern Med.* 2003;163(3):286-92.
- Not T, Horvath K, Hill ID, et al. Celiac disease risk in the USA: high prevalence of antiendomysium antibodies in healthy blood donors. *Scand J Gastroenterol.* 1998;33(5):494-8.
- 29. **Delco F, El-Serag HB, Sonnenberg A.** Celiac sprue among US military veterans: associated disorders and clinical manifestations. *Dig Dis Sci*. 1999;44(5):966-72.
- 30. **Karnam US, Felder LR, Raskin JB.** Prevalence of occult celiac disease in patients with iron-deficiency anemia: a prospective study. *South Med J.* 2004;97(1):30-4.
- Freeman HJ. Biopsy-defined adult celiac disease in Asian-Canadians. *Can J Gastroenterol.* 2003;17(7):433-6.

32. **Freeman HJ.** Celiac diseasse associated with primary biliary cirrhosisin a Coast Salish native. *Can J Gastroenterol.* 1994;8:105-7.