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Dohan proposed that an inherited defect interacting with an environmental trigger of gluten precipitated schizophrenia in some individuals, and provided supportive epidemiological evidence. Some clinical trials and case studies showed that a cereal free diet improved remission of symptoms of schizophrenia. The most important genetic marker found in the study of coeliac disease (6p23-p22.3) is very close to the dysbindin locus, which has been implicated in schizophrenia.2

Participants, methods, and results

The case sample comprised 7997 people older than 15 who were admitted to a Danish psychiatric facility for the first time between 1981 and 1998 with a diagnosis of schizophrenia and known maternal identity. For each case we randomly selected 25 controls from a unselected patients to self manage warfarin and as such may give a real indication of expected uptake. For patients keen to undertake self management three quarters were able to complete training. These patients considered it a convenient and valuable method of controlling their own health and most were enthusiastic to continue after the trial. If self management by patients is to become established standardisation and dissemination of training are needed, accompanied by practical guidelines to encourage back up from clinicians.

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Coeliac disease is presumably underascertained in this study. Only coeliac disease occurring before onset of schizophrenia in the case was considered, and only cases in hospitals or specialty clinics are included. Clinical symptoms of coeliac disease occur in only a fraction (about one in seven) of people with the pathognomic antibody.

Ascertainment bias does not explain the results because it would exist with equal strength for Crohn's disease and ulcerative colitis, and the relative risk for them is weak. Ascertainment bias would not affect the relation of schizophrenia and coeliac disease in parents, which had similar strength to that in cases.

Removal of gluten from the diet is not dangerous or expensive and is an effective treatment for coeliac disease. Failure of replication in earlier clinical trials of gluten withdrawal may have been the result of sampling fluctuation since coeliac disease is rare. New screening tests for coeliac disease are inexpensive and carry minimal risk and discomfort. An important question is the degree to which removal of gluten from the diet will alleviate symptoms in the small proportion of people with schizophrenia who screen positively for coeliac disease but do not show its classical symptoms.

Supporting data for potential gluten withdrawal in schizophrenia


data table

S L Thomas, J G Wheeler, Andrew J Hall

Herpes zoster results from reactivation of latent varicella zoster virus infection. Risk of zoster increases with age and with depressed cell mediated immunity, but relatively little is known about other factors.1-3 Case reports and case series have indicated that mechanical trauma may be a risk factor.4 But these studies had no control group, and physical mishaps and surgical trauma are common in older people. We investigated whether trauma is associated with increased risk of zoster using a case-control design.

Methods and results

This investigation was part of a study based in general practice in London of the determinants of zoster in adults without underlying immunosuppression. Elsewhere, we describe recruitment and definitions of cases (patients with incident zoster) and controls (patients with no previous zoster, individually matched to cases by age, sex, and general practice) elsewhere.5 Participants gave informed consent.

We hypothesised that trauma increases the risk of zoster at the trauma site within one month of the trauma, as indicated by a previous case series.1 We asked participants about all physical trauma severe enough to cause bruising (without prompting as to the site of trauma) and about surgical procedures in the six months before interview. We compared occurrence of trauma among cases and their matched controls in the month before the case developed zoster, evaluating both trauma at the site of the case's rash and trauma occurring elsewhere. We used matched comparisons of the timing and site of trauma because the risk of trauma varies seasonally, and trauma occurs at certain body sites more often. We used multivariable conditional logistic regression to determine the independent effects of trauma on risk of zoster.

We got information on trauma for 243/244 cases (median age 57.2 years; range 16.5-91.2 years) and 483 matched controls. In the six months before interview, cases and controls had a similar frequency of trauma at body sites other than the site of the cases’ zoster. But cases more often reported prior trauma at the site of their rash—this was associated with an eightfold increased risk of zoster as determined by multivariable analysis (table). Fourteen of the 22 participants who experienced trauma at the same site as subsequent zoster (mostly the trunk or head) did so in the month before the rash started (see table A on bmj.com). This recent trauma was associated with an adjusted 12-fold increased risk of zoster (table). Again, cases and controls had similar frequency of trauma to other body sites.

### Table A on bmj.com gives details of 14 people who had recent trauma at the site of subsequent rash in the case.