



NHS Foundation Trust

Does the UK population suffer with symptoms related to **Gluten Sensitivity?**

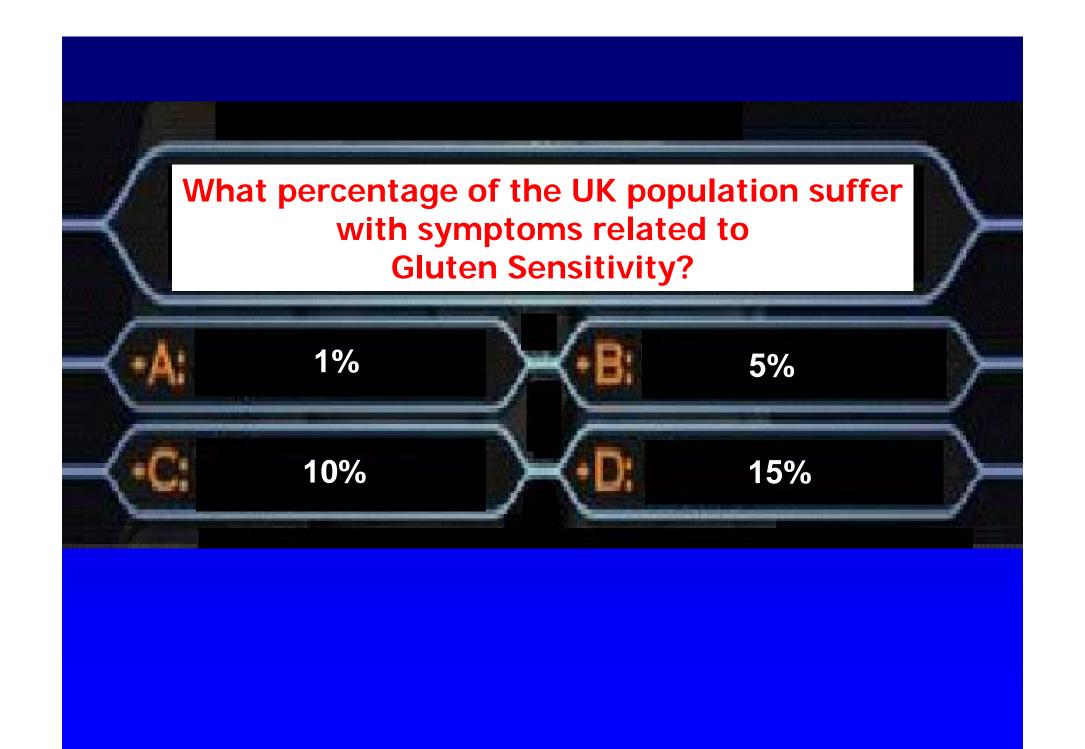
December 2012

Professor David S Sanders Consultant Gastroenterologist Royal Hallamshire Hospital & University of Sheffield





Jessica Ennes Gold Medal in Heptathlon



Stars' diet is free of gluten but full of fat

Celebrities claim to lose weight with wheat-free food, but it is often full of calories — and pricy, writes Kathryn Cooper

SUPPEMARKETS are cashing in on the celebrity trend for wheat free dies with products that can be up to three times the price of standard foods and contain more than three times the fat.

High-profile stars such as Gwyneth Paltrow, Rachel Weisz and Victoria Beckham beautie, which cut sout the protein found in wheat, basiley and rye. Many consumers are also adopting the diet, believing it to be a healthier option.

However, a Sunday Times survey of gluten-free products.

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survey of gluten-free products from four big supermarkets from four big supermarkets ably higher in fat than the

ably higher in fat than the quivalent standard product. A bandard equivalents, found bowl of glutten-free granula from Sainsbury's contained the standard equivalents, found that shoppers who believe glu-rife quitage the properties of the standard that shoppers who believe glu-rife quitage the properties of the standard saintbury's 'free from' multi-turing out glutten they are given to see weight. It's a myth, saintbury's 'free from' multi-said Tanya Thomas, a dietician and member of the British wholemeal pitta — 2478- more (glutten-free foods have none-free flowed from the standard wholemeal pitta — 2478- more more only granula that 248, of more only granula that 248, of more only granula that are the standard wholemeal pitta — 2478- more more only granula that are the standard properties of the standard wholemeal pitta — 2478- more more only granula that are the standard properties of the standard properties of

ON THE MENU

Sainsbury's "free from" multi-seed pitta 5.2g Sainsbury's wholemeal pitta 1.5g

Dietetic Association. The supermarket's "free from" oaty granola has 27.4g of

Gluten-free foods have none-free less become big business. Harks & Spencer recently reported that sales of its "made without wheat" range had some without wheat" range had some without wheat "ange had some without wheat" range had some without wheat "ange had some without wheat" range had some without wheat "ange had some without wheat" range had some without wheat "ange of glutten-free leads where the some with stabilities of a plate from years ago and went not with Wimbledon and the Us you which he said was down to fail set required in free from products for technour that will reasons". This is at odds with the desired products a state of the wind with the wind with the single proportion the splitten from belief to make the products and this contains 15.7g of fat per 10g compared with 0.5g for two had contained that in order to product splitted to a plate the free realing. The tensities at a possible value of a product of a produ

Fat Per 100g

2.7g 0.9g

6.9g 7.7g 6.9g 17g

Beckham, top left, Rachel Weisz, inset, and Gwyneth Paltrow eat gluten-free food

House price rise gathers pace

Kathryn Cooper Economics Correspondent

HOUSE prices have increased at their fastes

increased at their fastest annual rate for almost two years, raising further hope that the economy has turned a corner. According to the Land Registry, the price of an annual rate for almost their control of th

year to £363,802. The southeast also saw buoyant growth, with the average property up 2.3% to £210,301. However, six out of 10 house-price regions suffered falls over the past year. The northeast had the biggest annual drop, with the average price down 3.2% to £39,163. In towns, Hartlepool experienced the biggest annual price rise in the annual price rise in the

year to September with an increase of 5.7%, while Blaenau Gwent in south Wales suffered the biggest fall, with a drop of 8.3%.

Sales of homes in the £1.5m-£2m band rose by 31%, but those priced over £2m recorded no change. The property data follows news that Britain emerged from its double-dip recession in the third quarter of this year when the economy grew by 1%.

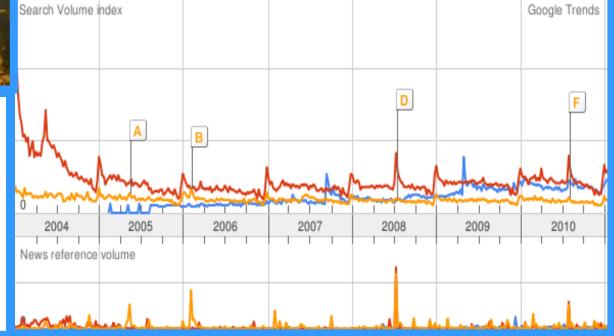
Mixed fortunes

Annual change to Sept 2012 Southeast +2.3 East +1.9 Southwest +0.9 East Midlands -0.1 West Midlands -1.6
Northwest -2.2 Yorkshire -3.0 Northeast -3.2



The U.S. market for gluten-free food and beverage products grew at a compound annual growth rate of 28 percent from 2004 to 2008, to finish with almost \$1.6 billion in retail sales

- Gluten Free diet
- Low Carb diet
- Low Fat diet



The UK retail market is worth £125 million*- when you consider the healthcare market is only worth around £37 million** and that there are only around 120,000 diagnosed coeliac patients in the UK then this may suggest other individuals opting for a gluten-free diet

^{*} Kantar Worldpanel, April 2011

^{**}IMS data



What is the prevalence of Gluten Sensitivity?

Any clues from secondary care?

- Center for Celiac Research University of Maryland
- 2004-2010
- 347/5896 ~ 6% or 1:7
- Secondary Care Referral Pattern
- Referral Bias or Ascertainment Bias?

Sapone A et al BMC 2012



What is the prevalence of Gluten Sensitivity?

Any clues from secondary care?

- Department of Clinical Medicine of St Orsola-Malpighi Hospital (Italy)
- January 2009 and June 2011
- N=78 included in present study (symptoms on ingestion of gluten)
- negative for EMA and TTG
- Negative for Wheat (specific IgE and skin prick tests)
- Normal duodenal mucosa or 33/78 (42%) IEL's
- 36/78 (46%) HLA DQ2 Or DQ8

Volta U et al *J Clin Gastroenterol* 2012



SELF REPORTING OF GLUTEN SENSITIVITY: SECONDARY CARE EXPERIENCE IN THE UK

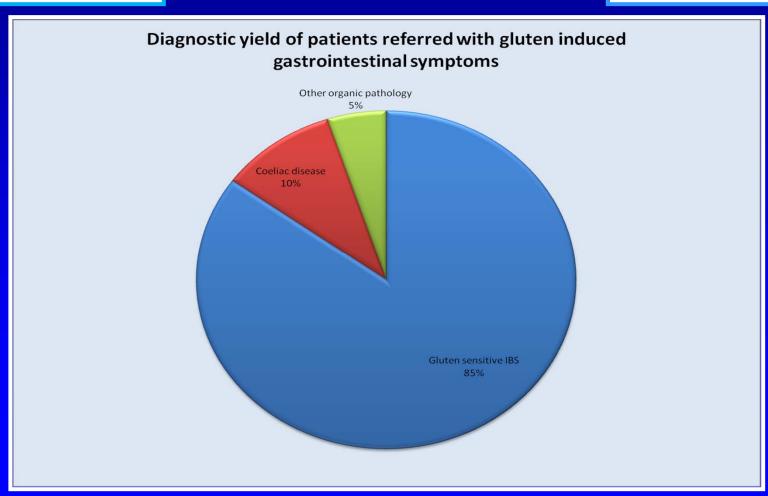


- All patients were referred to a dedicated coeliac or gluten sensitivity clinic by GPs
- The referral criteria were "GI symptoms attributed to gluten ingestion"
- 140 patients were investigated over a 5 year period. 80% were women and the median age of presentation was 37 yrs (range 16-88).



SELF REPORTING OF GLUTEN SENSITIVITY: SECONDARY CARE EXPERIENCE IN THE UK







SELF REPORTING OF GLUTEN SENSITIVITY: SECONDARY CARE EXPERIENCE IN THE UK



- A positive coeliac serology (p < 0.0001) was significantly associated with coeliac disease
- All patients with coeliac disease were HLA DQ2 or DQ 8 compared to 44% of GS-IBS cases
- There was statistically no significant difference in gender, clinical symptoms or baseline bloods (haemoglobin, vitamin B12, folate, ferritin, calcium or albumin) between the groups, (p > 0.05)



Meadowhall

8th largest
Shopping
centre in the
UK

1,500,000 sq ft (139,355 m2)

280 shopping outlets



Medical Questionnaire On Gastrointestinal Symptoms

Dear Sir/Madam

We would be grateful if you would kindly complete this 1-5 minute questionnaire survey as part of a research project being undertaken by the Gastroenterology department at the Royal Hallamshire Hospital, Sheffield. There are two parts to this questionnaire and although it asks about your bowel symptoms and past medical history, it is anonymous and the results are confidential and will be used only for research purposes. This questionnaire has been registered with the Sheffield Teaching Hospitals. You may find parts of this questionnaire repetitive but please try and answer all the appropriate questions tailored for you. Should you have any queries or difficulties completing this survey, please ask our helpful young student doctors!

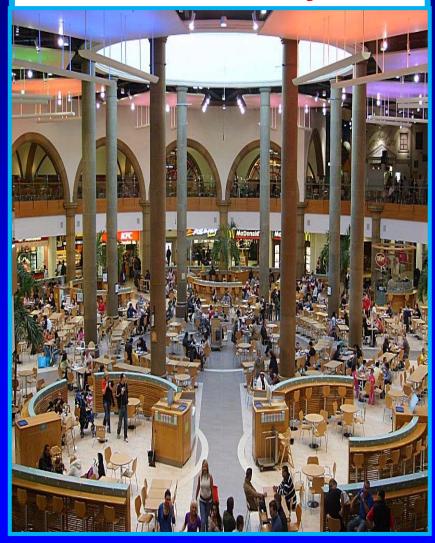
Thank you for your co-operation and time in completing this questionnaire

Part 1: This asks for basic information about yourself, any abdominal symptoms and your general state of health

Carlo Car				
Q1) Age		D	О.В	
Q2) Male 🗌	or	Female		
Q3) Employed 🔲	, Unemployed	☐, Disabled	☐, Retired	
Q4) Single	, In a relationshi	p 🗆 , Divorced	□, Widow	ed 🗆
Q5) Race: Whi	te 🗆, Black 📮	Asian □, O	ther 🔲 (please state	
Q6) Post code				
Q7) Have you suffe more?	red with episodes Yes		s or discomfort for t - if No, please go to	
Q8) If yes, <u>how mar</u> discomfort?	ny days in a month	<u>i</u> do you approxima	ntely experience these	e abdominal pains or
One day a month			Four days a mo	nth 🗆
Two days a month			5-10 days a mo	nth 🗌
Three days a month		M	ore than 10 days a mo	onth 🗆
Q9) Do you suffer fro	om abdominal bloa	nting (feeling full of	gas)? Yes 🗆	No 🗆



Q10) Do you feel an improvement in your abdon your bowels? Yes	ninal pai No		j			
Q11) Was the start of your abdominal pains or di frequency? Yes	iscomfort No		: I			
Q12) If yes to Q11, how would you best describe the Diarrhoea, Constipation, Al-						
Q13) Was the start of your abdominal pains o consistency? Yes	or discom No	_	ıl			
Q14) If yes to Q13, how best would you describe your stool motions? Loose, watery or sloppy Hard, pellet like Combination of loose at times and hard other times						
Q15) Do your abdominal and bowel symptoms get Q16) Are you known to suffer from any of the follo						
		Chronic headaches				
Anxiety						
Depression		Nut allergy				
Bipolar disorder		Egg allergy				
Schizophrenia		Dairy product intolerance				
Schizophrenia Thyroid disease		Dairy product intolerance Bowel cancer				
Thyroid disease		Bowel cancer				
Thyroid disease Young onset diabetes (childhood/early adulthood)		Bowel cancer Stomach cancer				
Thyroid disease Young onset diabetes (childhood/early adulthood) Pernicious anaemia (low vitamin B12)		Bowel cancer Stomach cancer Irritable bowel syndrome				



Part 2 – Gluten related symptoms

This part of the questionnaire focuses on whether you develop problems when you eat gluten. Gluten is a product found in wheat, barley or rye. Therefore, it is found in common everyday diets such as <u>cereal, bread, cakes, biscuits, pasta, pizza</u> etc. Should you have any queries or difficulties completing this survey, please ask our helpful young student doctors!

Bloating (feel full of air) Headaches	ì						
Abdominal Pain Mental confusion							
Abdominal discomfort Lack of co-ordination							
Diarrhoea Numbness/pins & needles							
Constipation Lack of energy							
Belching Skin Rash							
Flatulence Joint Pains —							
Sickness Anaemia							
Others (please specify)							
If you do not suffer from any gluten related symptoms there are no further questions — than you. Q2) If yes, how often do you experience symptoms after eating gluten products?							
Every time I eat gluten products Few times a month							
On most occasions/days Few times a year							
Few days a week	Ì						
Q3) How soon after eating gluten products do you develop symptoms?							
Almost immediately (less than one hour) The next day							
1-6 hours later A few days later							
6-24 hrs later							
Q4) How long do your symptoms generally last for?							



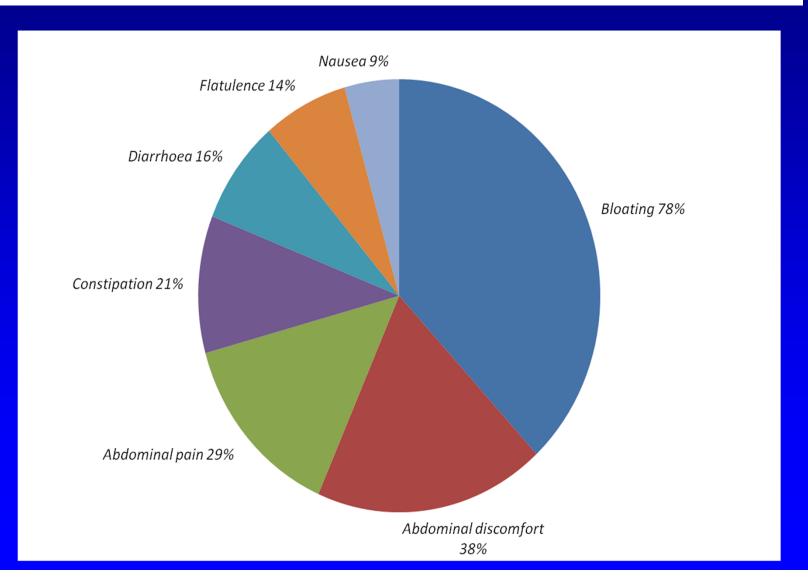
Q5) Which gluten product(s) se	ems to cause proble	ms? (tick as mar	y that apply)	
Bread	Pizz	a 🗆		Cakes
Cereal	Pas			Biscuits
Porridge	Oth	ers (please state	:)	
Q6) How long have you had a p	roblem related to glu	ıten?		
(state approximate nur	mber) months	oryears		
Q7) Have you ever seen a healt	hcare professional d	ue to problems r	elated to gluten?	Yes □ No □
				165 110
Q8) If yes, please state whom y				
GP □ , Hospital do	ctor 🗆, dietician	□, other □	(please state)	
Q9) If yes, have you undergon	e any of the followin	g tests to look	pecifically for a	cause as to why
you have problems related to g	luten? (tick as many	that apply)		
Coeliac blood test	Yes 🗌	No		Not sure
Skin prick allergy test	Yes	No		Not sure
Endoscopy (camera into stoma	ch) Yes 🗆	No		Not sure
You have had no tests at all	Yes			
Other tests (please state)				
010) # +- 07 00 00 b-		£ eb - £-U-		fulcasa sala eka
Q10) If yes to Q7, Q8, Q9 ha student doctor to explain the d	_		wing diagnosis?	(piease ask the
Coeliac disease□, Wheat	allergy □,	Coeliac disc	ease has been ex	cluded,
No explanation given 🔲 ,	You are not sure	, other		
Q11) Have your ever tried a glu	iten free diet?		Yes 📙	No 📙
Q12) If yes, was it beneficial for	your symptoms?	Yes	No 🗆	Not sure
O13) If yes to O11. Are you still	on a gluten free diet	7	Yes 🗆	No 🗆

Overall Demographics:

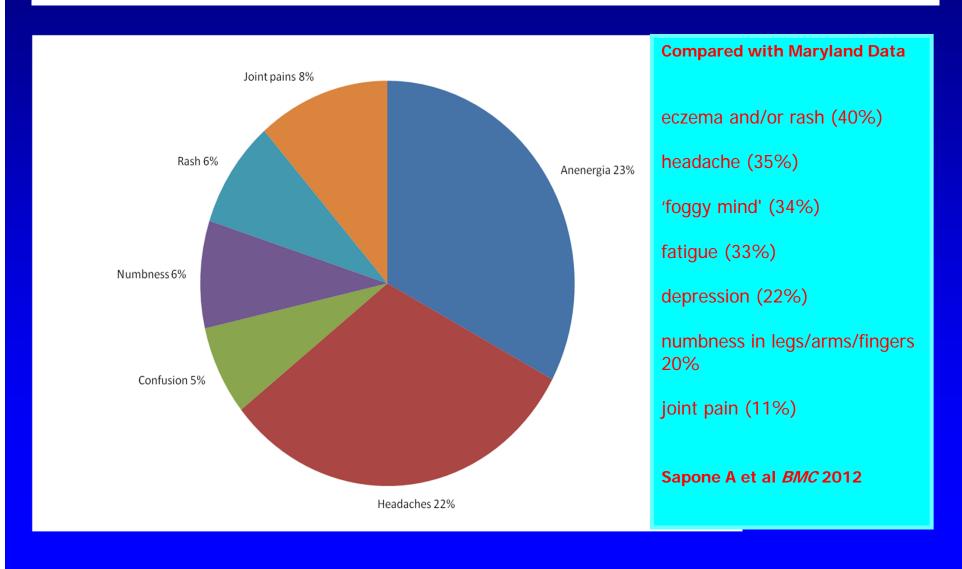
- 1002 adults completed the population based questionnaire
- 55% female, age range 16-93, mean age 39 years
- Prevalence of individuals fulfilling the ROME III criteria for IBS was 6% with up to 80% being female (p<0.0001)

The prevalence of gluten related symptoms was 13% (129/1002), female 80% (P < 0.0001)

Gastrointestinal gluten sensitive symptoms reported (Sheffield, UK)



Extraintestinal gluten sensitive symptoms reported (Sheffield, UK)



Basic information	Never tried GFD (n 92)	Tried GFD (n 37)	P-value
Age	Mean 40 Median 39 Range 18-75	Mean 37 Median 36 Range 19-69	0.4
Females	76 % (70F, 22M)	86% (32F, 5M)	0.2
Employed	55%	57%	1.0
Duration of symptoms	Mean 54 months, Median 36 Range 2-240	Mean 96 months, Median 60 Range 3-480	0.01
Seen doctor	6.5% (6 yes, 86 no)	78% (29 yes, 8 no)	< 0.0001 OR 52

Patients with IBS were more likely to report sensitivity to gluten based products than non-IBS patients (35% versus 11%, p<0.0001)

Symptoms due to gluten ingestion	Never tried GFD (n 92)	Tried GFD (n 37)	P- value
Bloating	75% (69y, 23n)	86% (32y, 5n)	0.2368
Abdominal pain	15% (14y,78n)	64% (24y, 13n)	< 0.0001*
Abdo discomfort	30% (28y, 64n)	56% (21y, 16n)	0.0086*
Diarrhoea	4% (4y,88n)	43% (16y, 21n)	<0.0001*
Constipation	15% (14y, 78n)	35% (13y, 24n)	0.0168*
Belching	6.5% (6y, 86n)	24% (9y, 28n)	0.0116*
Flatulence	13% (12y, 80n)	24% (9y,28n)	0.1233
Sickness	4% (4y, 88n)	22% (8y, 29y)	0.0048
Headaches	15% (14y, 78n)	38% (14y,23n)	0.0085*
Confusion	2% (2y, 90n)	13.5% (5y, 32n)	0.0205*
In-coordination	0% (92n)	8% (3y, 34n)	0.0222*
Numbness	4% (4y, 88n)	11% (4y, 33n)	0.2251
Anenergia	17% (16y, 76n)	38% (14y,23n)	0.0202*
Rash	4% (4y, 88n)	11% (4y, 33n)	0.2251
Joint pains	6.5% (6y, 86n)	11% (4y, 33n)	0.4711

Confusion, Confusion!



Isolated gliadin antibodies are prevalent in the general population but more prevalent in IBS

Table 2 Coe	iac antibod	y results
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	Antibody negative	lg G antigliadin antibody only	IgA antigliadin	IgG and IgA antigliadin antibodies	EMA only	EMA and IgG antigliadin antibody	All three antibodies
Volunteers (n = 1200) Biopsy-proven coeliac disease Normal histology Intraepithelial lymphocytes	1038	139 1**	10 1 6 3	4 (1 declined*) 1 1	3 3	4	2 2

^{*}A 79-year-old female declined biopsy due to recent stroke.

^{**}Also one case of coeliac disease diagnosed coincidentally having been referred to secondary care for investigation of irritable bowel syndrome.

	Antik nega	lgG antigliadin only	lgA antigliadin only	lgG and IgA antigliadin	FMA only	EMA and IgG antigliadin	EMA and IgA antigliadin	All three antibodies
Study group (n=300)	234	49	4	1	6	4	1	1
Coeliac disease (n=14)*		1	1	1	5	4	1	1
Controls (n=300)	256	41	1			2		
Coeliac disease (n=2)			**			2		

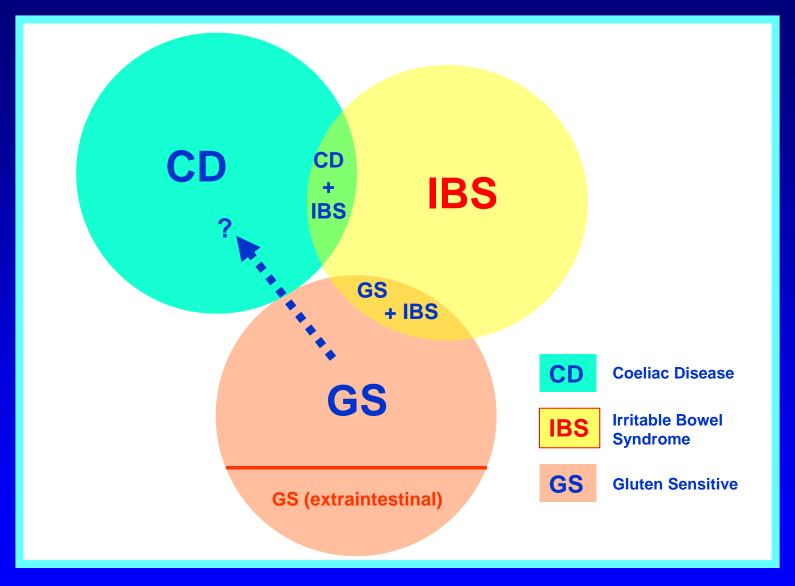
EMA=endomysial antibodies. *One patient refused to undergo biopsy.

Table 2: Coellac antibody results

1506

THE LANCET • Vol 358 • November 3, 2001

Sanders DS EJGH 2003 & Lancet 2001



A Model for the relationship between coeliac disease, IBS and gluten sensitivity?

Ball A and Sanders DS Am J Gastroenterol 2010;105:222-3



Non-Celiac Wheat Sensitivity Diagnosed by Double-Blind Placebo-Controlled Challenge: Exploring a New Clinical Entity

Antonio Carroccio, MD¹, Pasquale Mansueto, MD², Giuseppe Iacono, MD³, Maurizio Soresi, MD², Alberto D'Alcamo, MD², Francesca Cavataio, MD³, Ignazio Brusca, MD⁴, Ada M. Florena, MD⁵, Giuseppe Ambrosiano, MD², Aurelio Seidita, MD², Giuseppe Pirrone, MD² and Giovanni Battista Rini, MD²





Conclusion

This is the first UK population prevalence survey of gluten related symptoms

The prevalence self-reported gluten related symptoms was ~13% (129/1002), female 80% (P <0.0001)

3.7% are on a GFD for gluten related symptoms

Patients with IBS were more likely to report sensitivity to gluten based products than non-IBS patients (35% versus 11%, p<0.0001)

Gluten sensitivity was more common if the symptom duration was long or the individual had consulted a doctor



BMJ 2012;345:e7907 doi: 10.1136/bmj.e7907

PRACTICE

UNCERTAINTIES PAGE

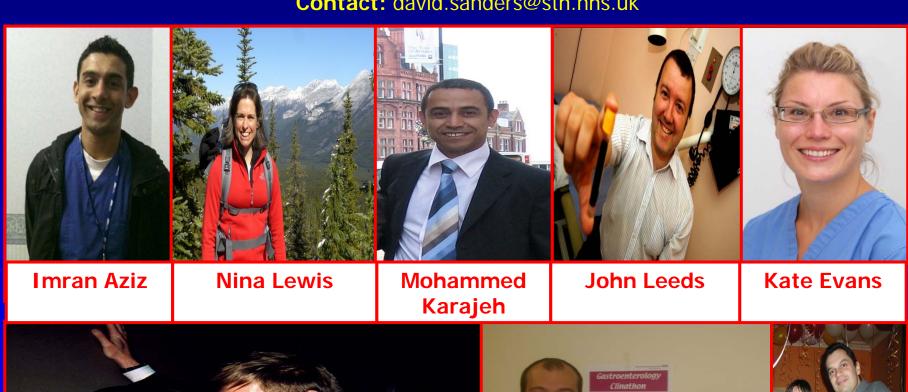
Does gluten sensitivity in the absence of coeliac disease exist?

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On-going Research at the Royal Hallamshire Hospital & International GI and Endoscopy Fellowships

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