



### DISCLOSERS

Consultant: Fody Foods Sponsored posts: Salix, Campbell soup, Casa de Sante Published Iow FODMAP books, app, e-books

# OBJECTIVES

 $\diamondsuit$  Distinguish between the causes and symptoms of food allergy and food intolerance

Detail factors involved with food intolerance/malabsorption and GI symptom induction

List potential IBS food triggers

### **IBS: ROME IV CRITERIA**

Recurrent abdominal pain x 3 month, at least 1 day per week associated with two or more of the following:

Related to defecation
 Onset associated with a change in the frequency of stool
 Onset associated with a change in the form of stool

Mearin et al. Gastroenterology. May 2016

### **ROME IV: ALTERATIONS TO NOTE**

 Motility disturbance: alteration in the movement of food and waste through the GI tract.
 Visceral hypersensitivity: heightened experience of pain in the internal organs.
 Altered mucosal + immune function: changes in the gut's immune defenses.
 Altered gut microbiota: changes in the <u>community of</u> <u>bacteria in the gut</u>
 Altered CNS processing: changes in how the brain sends and receives from the gut.

# 4 IBS SUBTYPES

Based on stool type and frequency. The subtypes have not changed, but Rome IV includes a slightly new way of identifying IBS subtypes. Subtypes are based frequency of lose or very hard stools.

	$\sim$	S
SUBTYPE	STOOL TYPE 1 & 2	STOOL TYPE 6 & 7
IBS with predominant constipation	More than 25%	Less than 25%
IBS with predominant diarrhea	Less than 25%	More than 25%
IBS with mixed bowel habits	More than 25%	More than 25%
IBS Unclassified: Patient wi whose bowel habits cannot three subtypes above. Source: Lacy BE, et al. Bowel Di	be accurately categori:	zed into one of the
35 IrritableBox		

### **IBS: GUT BRAIN AXIS**

Enteric nervous system and CNS derive from same nerves in fetus.

•Stimulation of the bowel can affect areas in the brain producing emotional distress, which in turn can affect bowel functioning.

Emotional conflict can lead to greater IBS symptoms; treatments directed at emotional distress, like hypnosis or relaxation methods and anti-depressants can reduce symptoms

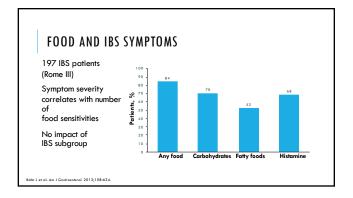


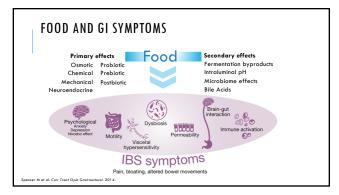
# **ALLERGY VS. INTOLERANCE**

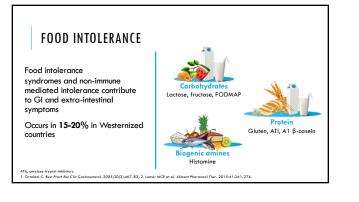
Food allergy: immune mediated reaction

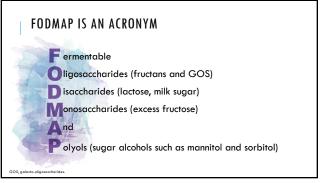
Food intolerance: non-immune, enzymatic defects such as lactose intolerance, transport defects such as fructose, pharmacological such as effects of vasoactive amines or undefined such as NCG/WS.

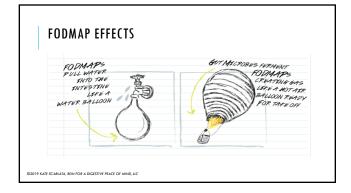
ALLERGY, AUTOIMMUNITY, AND INTOLERANCE					
Food reaction	Pathogenesis	Clinical Entity	Symptoms		
Allergy	IgE, non IgE, occasional IgE	Food allergy, F-PIES, EoE	Respiratory, GI, cardiovascular, ski anaphylaxis		
Autoimmune	Innate and adaptive immunity	Celiac disease (1% population)	Gl symptoms, fatigue, low iron osteoporosis, B12, folic acid, weight or gain, and more		
Food intolerance	Disorder of digestive/absorptive process, toxic or pharmacologic reactions	Lactose intolerance Sucrose-isomaltase deficiency,+ FODMAP Histamine	GI Gas, bloat, constipation/diarrhea, pain Other Hives, low blood pressure, headac pain, diarrhea		

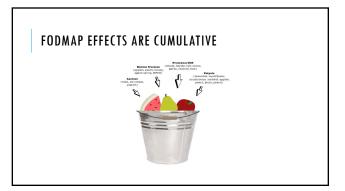


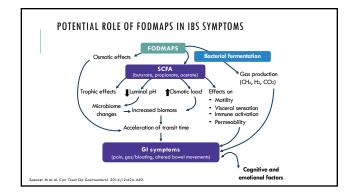


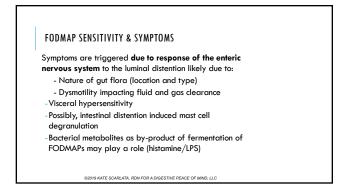












### WHY ARE FODMAPS MALABSORBED?

Lactose: Up to 70% of the world population has lactase non persistence, also secondary Ll observed in postinfectious IBS & SIBO

 Fructose: poor absorption due to it's slow, low-capacity transport mechanism across the epithelium & SIBO; FM occurs in 1 in 3 ppl

•Fructans/ GOS: humans lack digestive enzymes

Polyols: too large for passive diffusion; absorbed in pores in small intestine.

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**Purpose:** Investigate the action of fructose and inulin on the small bowel and colon in healthy subjects using MRI technique.

N=16 healthy subjects (no IBS) Randomized single-blind crossover study

Murray el al Am J Gastroenterol 2014

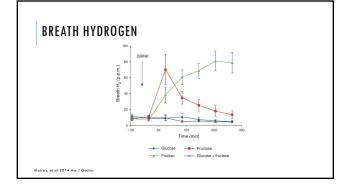


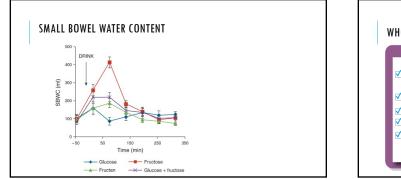
### METHODS

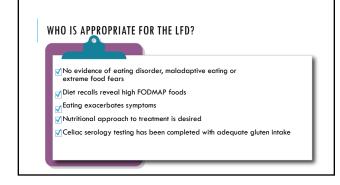
Volunteers underwent a baseline fasted scan 45 min before ingestion of the test meals.

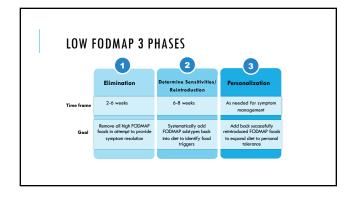
Test drinks=500 ml H\_20 w/ either 40 g gluc, fructose, inulin or mix of 40 g gluc + 40 g fructose

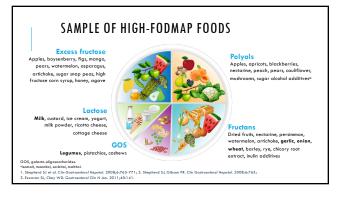
Followed by a scan every hour up to t=300 min. After each scan, breath hydrogen (H<sub>2</sub>) tests were performed using a portable hand-held breath H<sub>2</sub> meter



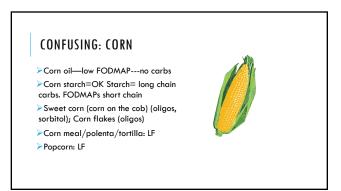








FODMAP S	WAPS	
Food	Choose	Lose
Garlic	Garlic infuse oil; Boyajian brand	Garlic flesh, garlic powder or salts
Onion	Shallot or onion infused oil or use chives, asafetida powder or green part of leeks or scallions	Onion, shallot, leek (Fructan in the bulb); onion powder or salts
Legumes	1/2 cup canned: rinsed and drained chickpeas or 1/2 cup canned lentils	Kidney beans or dried beans
Wheat flour	King Arthur GF flour blend, Trader's Joes GF flour, Bob's Red mill 1 for 1 cup GF	Wheat flour
Soybean	Firm tofu, edamame	Soymilk made w/ whole soybean, silken tofu
Milk alternatives	Lactose free milk, hemp, almond, canned coconut milk, rice milk	Cow's milk
Cheese	Most hard/aged cheese	Ricotta/ Cottage



### DAIRY

Most cheeses low lactose.

Lactose is in wet part that is drained off with whey.

Greek yogurt lower in lactose than traditional.

Cheeses such as American, Cheddar, feta, Brie, semi-soft goat cheese low enough for low FODMAP diet

Whipped cream, half and half, cream cheese (2 TB) in small amount well tolerated and low enough in lactose.



# TOMATO Diced tomatoes (avoid canned w/ onion/garlic)=low Sun dried tomatoes: limit to 3 pieces (8 grams) Marinara: Choose products made without onion/garlic (Prego Sensitive Recipe, Rao's Sensitive Formula, Fody foods)

### SOY

#### High FODMAP

Soy flour, whole mature soybeans, silken tofu

#### Low FODMAP

Edamame (1 cup), firm tofu, soy milk made with soy protein (8<sup>th</sup> Continent ®), soy sauce, soy lecithin



#### SOURDOUGH BREAD

✓ FODMAP testing at Monash University has shown sourdough wheat/spelt bread to be lower in FODMAP than regular wheat and spelt breads

 $\checkmark$  Choose breads leavened with sourdough culture only-as slow leavened—less FODMAPs

✓ If product contains baker's yeast—rising was accomplished faster – and likely more FODMAPs remain.

### LABEL READING TIPS

#### Highlight of some <u>High FODMAP ingredients</u>:

 Agave, honey, HFCS, chicory root, inulin, FOS, fructose, dates, molasses

Wheat as primary ingredient—unlike GF diet, traces are okay

Soy flour or whole soybean, other bean flours

Apple and pear juice

 Natural flavors-can denote onion/garlic in savory USDA regulated foods (meat products/ broth)

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### BASIC GUIDELINES For the reintroduction phase

 Test one FODMAP group (lactose, excess fructose etc.) at a time & choose foods that contain only one FODMAP.
 Consume a food amount that represents a normal intake (not excessive amounts).

Continue to restrict all FODMAPs (maintain a low FODMAP diet) except the food that is being tested until tolerance or intolerance is confirmed.

Record symptoms experienced for each challenge.

>Use the same food for each of the 3 challenge days.

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#### DETERMINE SENSITIVITIES WITH THE FOLLOWING FOODS

- >Lactose: 1/2 -1 cup milk
- Fructose: 1-2 TB. honey or 1/2 mango
- Fructans: 2 slice wheat bread, 1 TB onion, ½ garlic clove
- ►GOS: 1/2 cup beans

Polyols: ½ cup mushrooms, 1/3 cup cauliflower (mannitol) or 1 peach , 5 blackberries (sorbitol)

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#### WHAT IS A FAILED CHALLENGE?

- A failed challenge should be a noticeable & significant change in symptoms.
- > Symptoms may resemble an IBS flare: diarrhea, cramping, return of constipation, bloating.
- > = Undesirable outcome.

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#### WHY REINTRODUCE FODMAPS?

- Research has shown that the low FODMAP diet reduces bifidobacteria and other problotic gut bacteria (butyrate-producing Clostridium cluster XIVa and mucus-associated Akkermansia muciniphila (Halmos, Gut 2015)
- Stool pH increases slightly on the low FODMAP diet---this may allow pathogenic microbes to grow. (Halmos, Gut 2015)
- The low FODMAP has been shown in 2 studies to increase gut microbial diversity—a good thing. (Halmos Gut 2015, McIntosh Gut 2016)

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#### NON-RESPONDER

#### Assess symptoms:

- >Bloating & post prandial fullness: r/o SIBO, gastroparesis
- Constipation: assess for slow transit constipation and/or dyssynergic defecation, high colonic stool burden, methane + SIBO
- Diarrhea: parasitic infection, bile acid malabsorption, SIBO
- >Other food intolerance/sensitivities: gluten, fat, food chemicals-
- histamine, milk protein (A1 vs A2)
- Consider probiotics, gut-directed hypnotherapy + other gut-brain directed therapies.

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### THE SCIENCE: RCTS EVALUATING THE LOW-FODMAP DIET FOR IBS

- 7 RCTs compared a low FODMAP diet with various controls in 397 participants A low FODMAP diet was associated with reduced overall symptoms compared to controls (RR 0.69; 95% CI 0.54, 0.88, I2 25%)
- The 3 RCTs that compared low FODMAP diet with rigorous control diets had the least heterogeneity between studies but also the least magnitude of effect
- The overall quality of the data was "very low" according to GRADE criteria Most studies were high risk of bias
- Heterogeneity between study designs

JC et al. Presented at DDW 2018. Washington, DC: June 4, 2018; Abst

### LOW FODMAP VS. TYPICAL AUSTRALIAN DIET

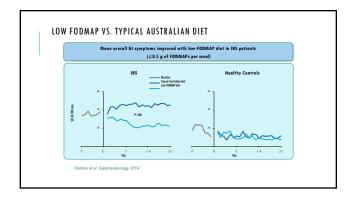
 $\succ$  This RCT compared GI sx. x 3 weeks of low FODMAP diet w/ the moderate FODMAP intake of a typical Australian diet

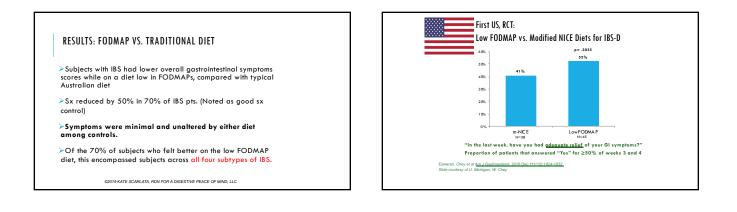
Crossover interventional study

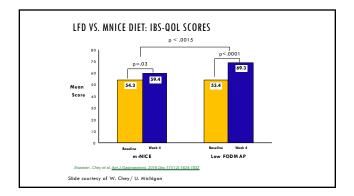
>38 participants: 30 IBS & 8 healthy controls

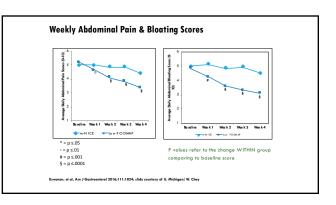
Almost all food, comprising 3 main meals & 3 snacks daily provided.

GI symptoms were measured daily during the baseline week and interventional diet periods using a 100 mm visual analogue scale (VAS). 82019 Kate Scantala, RDN For a Digestive Peace of Mind, LLC Helmon, E et al Gastmenterology (2014)









### LFD & THE METABOLOME

>N=37 IBS (19 LFD; 18 HFD) x 3-week diet.

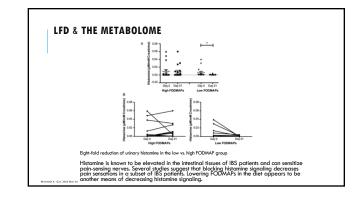
>LFD increased bacterial richness.

Metabolic profiling of urine differed after the diet (p<0.01), with 3 metabolites (histamine, p-hydroxybenzoic acid, azelaic acid)</p>

Histamine, a measure of immune activation, was reduced eightfold in the LFD group (p<0.05) and increased in the HFD group in subsets of patients. Strong evidence that histamine is an important signaling molecule linked IBS symptoms.

≻P-Hydroxybenzoic & azeliac acid increased on LFD—associated with potential anti-inflammatory effects.

Malntosh K. Gut. 2016



#### LFD & METABOLOME

Do FODMAPs modulate visceral sensitivity due to changes in gut microbiome and gut permeability?

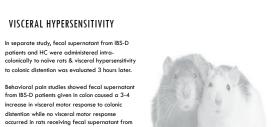
N=12 ; 6 IBS and 6 HC

Fecal samples obtained before & after LFD

Fecal LPS was 2 fold higher in IBS-D patients compared to HC

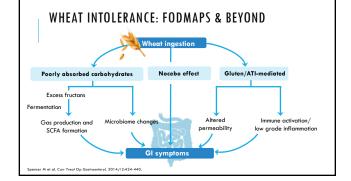
4 week treatment of LFD resulted in significant improvement of IBS symptoms and normalized fecal LPS to level similar of HC  $\,$ 

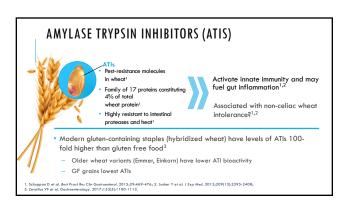
Zhou SY, Gillilland M 3rd, Wu X, et al. FODMAP diet modulates visceral nociception by lipopolysaccharide-mediated intestinal inflammation and barrier dysfunction. *J Clin Invest*. 2018;128(1):267-280.

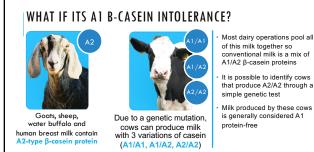


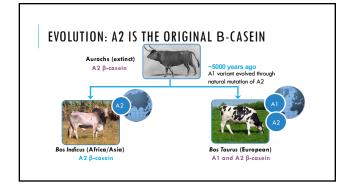
HC.

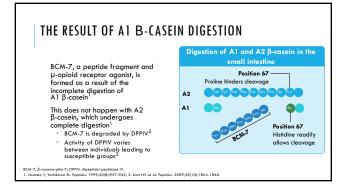
DDW 2016 Zhou, SY et al; U. Michigan

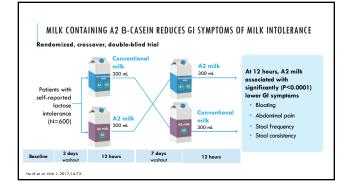


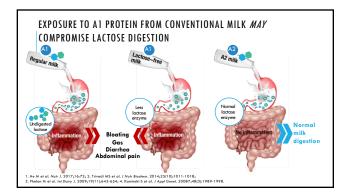


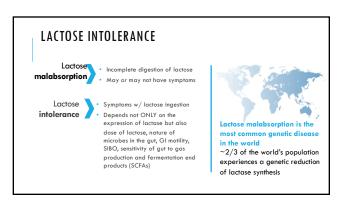


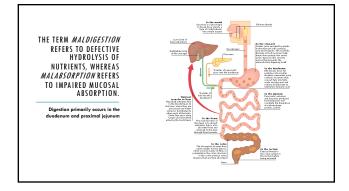












CAUSES OF MALDIGESTION	Inactivation of pepsin by achlorhydria; PPIs, H. pylori, autoimmune disease, etc.
	Improper mixing (gastric surgery, gastroparesis, neuroendocrine imbalance)
	Pancreatic insufficiency (output issue)
	Inactivation of lipase by low pH (dumping syndrome)
	Rapid transit causing malabsorption (hyperthyroidism)

	Mechanism	Malabsorbed Substrate	Causes
MECHANISMS	Conjugated Bile Acid Deficiency	Fat, fat-soluble vitamins, calcium, magnesium	Liver disease, biliary obstruction, SIBO, ileal disease
OF MALDIGESTION	Pancreatic insufficiency	Fat, protein, CHO, fat- soluble vitamins, B12	Congenital, chronic pancreatitis, pancreatic tumors, hyperacidity (inactivating pancreatic enzymes)
	Reduced mucosal digestion	CHO, protein	Mucosal disease (i.e. Crohn's, Celiac) , brush border enzyme deficiency (i.e. lactase),
	Intraluminal consumption of nutrients	B12, macronutrients	SIBO, parasitic infection

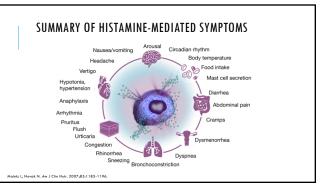
### WHEN TO SUSPECT MALABSORPTION

Fats: light-colored, foul smelling stools which are bulky and soft; difficult to flush. Proteins: fluid retention (edema) Sugars: explosive diarrhea, flatulence, or bloating Vitamins/minerals: wasting, malnutrition, low blood pressure, weight-loss, and anemia

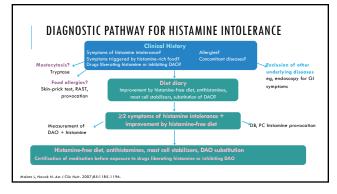


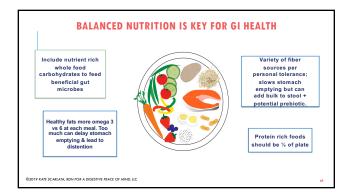
- Histamine: chemical released from our immune cells, in foods, created by gut microbes from amino acid.
- Histamine intolerance results from a disequilibrium of accumulated histamine and the capacity for histamine degradation
- In healthy persons, dietary histamine can be rapidly detoxified by DAO, whereas
  persons with low amine oxidase activity are at risk of histamine toxicity
- Some drugs can impair DAO activity
- DAO is synthesized by mature apical enterocytes-located on upper intestinal villi
- Mucosal damage (gastroenteritis, SBS) may reduce DAO and lactase activity

DAO, diamine oxidase. 1. Maintz I, Novak J. Am J Clim Nutr. 2007;85(5):1185-1196; 2. Enko D et al. Can J Gastroenterol Hepatol. 2016;Article IC 4893501.









# FOOD INTOLERANCE: RECAP

Often portion driven

Lactose intolerance: many ppl can tolerate 4 grams per serving of lactose; can use over the counter lactase enzymes to aid digestion.

Histamine: stress management, reduce histamine in diet, mast cell stabilizers and antihistamines to manage symptoms

FODMAPs: reduce, re-challenge and personalize to least restrictive diet.

Beta-casein: choose goat or sheep milk/cheese; trial A2 milk Assess level of food fear in IBS and/or maladaptive eating

#### WHEN HEALTHY EATING GOES TOO FAR.

There is some evidence that people with gastrointestinal disorders who undergo dietary change may be at increased risk for **disordered eating behaviors.** 

 $\ensuremath{\textbf{Orthorexia}}$  nervosa: a condition in which people restrict their diet based upon its quality.

This condition is associated with symptoms such as an "obsessive focus on food choice, planning, purchase, preparation, and consumption, food regarded primarily as a source of health rather than pleasure; and exaggerated faith that inclusion or elimination of particular kinds of **food can prevent or cure disease** or affect daily well-being.

Hill P, Muir JG, Gibson PR. Controversies and Recent Developments of the Low-FODMAP Diet. Gastroenterology & Hepatology. 2017;13(1):36-45.

#### ORTHOREXIA

Bratman (1997) coined the term "orthorexia nervosa" to describe people whose extreme diets - intended for health reasons - are in fact leading to malnutrition and/or impairment of daily functioning.

> Dunn TM, Bratman S. On orthorexia nervosa: a review of the literature and proposed diagnostic criteria. Eat Behav. 2016;21:11–17.

#### **ORTHOREXIA & FOOD FEARS**

When eating leads to guilt & shame...or a false sense of control. RDNs help individuals work on developing peace with food –not fear.

Reminding individuals that "managing digestive health is about creating a fuller life, not a more limited one".

### ARFID

Avoidant/restrictive food intake disorder

ARFID was introduced in the DSM-5 as a diagnosis of eating or feeding disturbance due to lack of interest in eating, avoidance of sensory characteristics of food, and/or fear of adverse eating consequence (eg, choking, vomiting, or digestive distress).

To meet diagnostic criteria, one doesn't have a distorted body image and the food disturbance must lead to one or more of the following: nutritional deficiency, weight loss, psychosocial impairment, or dependence on oral nutritional supplements or tube feedings.

ARFID can't be diagnosed if the eating disturbance is attributable to a concurrent medical or psychiatric condition.

American Psychiatric Association. Diagnostic and statistical manual of mental disorders (DSM-5). USA: American Psychiatric Association; 2013.

### ARFID

Preliminary research by Zia and colleagues found that approximately 28% of their functional gastrointestinal disorder (FGID) patient sample met criteria for ARFID.

Interpret with caution as we don't want to be too quick to assign an eating disorder to GI patients given the individuality of one's sensitive gut to potential food triggers and associated behaviors.

ARFID and ED screening tools not validated in IBS.

Zia J, Riddle M, DeCou CR et al. Frevalence of Eating Disorders, especially DSM-5's Avoidant Restrictive Food Intake Disorder, in Patients with Functional Gastrointestinal Disorders: A Cross-Sectional Online Survey DDW 2017, Abstract Mo1551

### **ROLE OF DIETITIAN**

Self-guided elimination place individual at nutritional risk.

Dietitians with expertise in food intolerance are essential to help guide individual Important to rule out serious health conditions and screen for celiac serology test PRIOR to diet change.

Diet is a factor –but stress can exacerbate symptoms. Gut and brain linked via bidirectional pathway, GBA. Stress management should be part of your plan. Dietitian can and should screen for maladaptive eating or ED prior to instructing patient on elimination diet which could trigger ED behaviors.

### RESOURCES

KATE SCARLATA BLOG AND WEBSITE: RECIPES, FREE DOWNLOADS, DIGESTIVE TIPS www.katescarlata.com CHECK OUT MY MMM CHANNEL: FOR A DIGESTIVE PEACE OF MIND Monash University Iow FODMAP diet app

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#IBELIEVEINYOURSTORY VIDEO

https://www.youtube.com/watch?v=7HQAIraW50s