SIBO – Everything You Need To Know About The #1 Cause of IBS



Dr. Tomah Phillips, ND

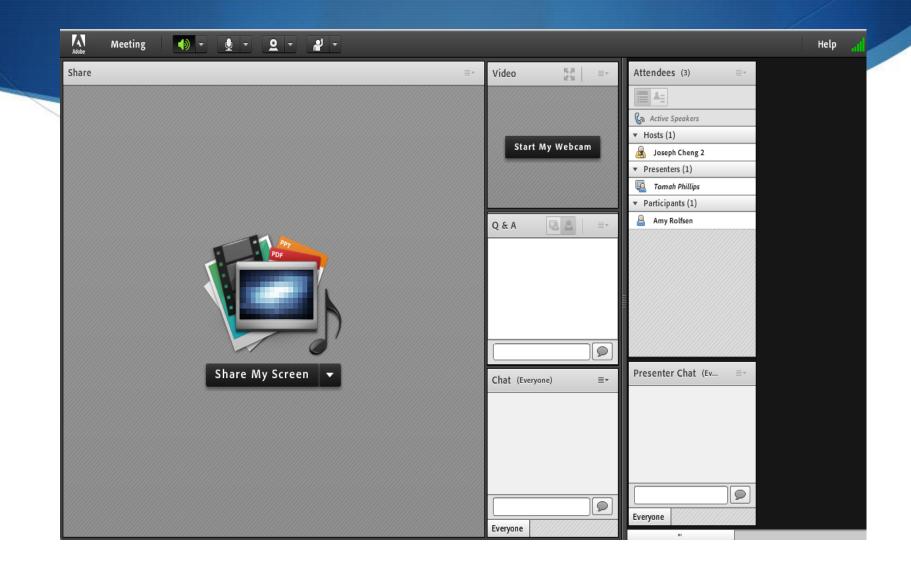


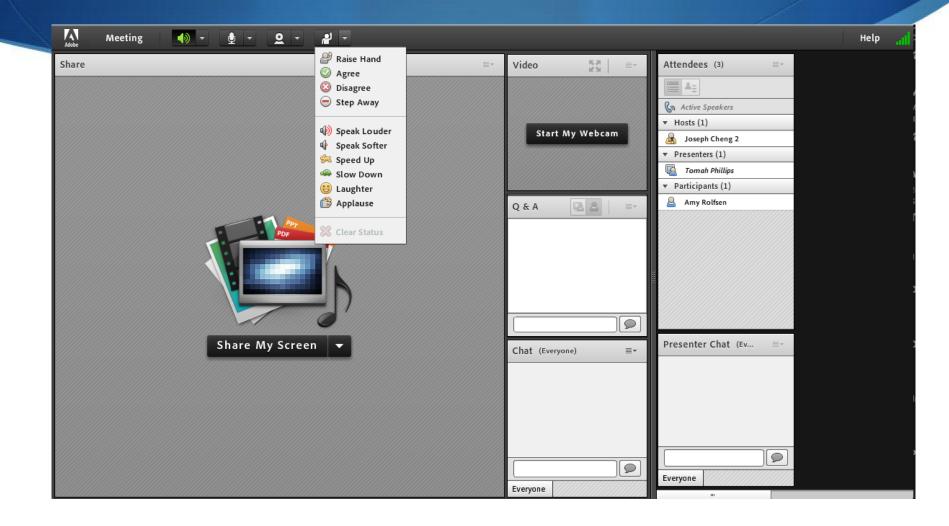
- Intro What is SIBO?
 - "Tolle causam" Underlying causes
- Clinical manifestations
- Pathophysiology
- Diagnosis
 - Best options, and test interpretation
- Treatment
 - Pharmaceutical, herbal and dietary considerations
- Tips for preventing relapse

Bio

- Graduated from Boucher Institute of Naturopathic Medicine
- Instructor of Biomedical Sciences at Boucher
- Instructor of Physiology and Pathology at the Canadian School of Natural Nutrition
- Practice at Evoke Integrative Medicine in downtown Vancouver
- Medical advisor for Vita Aid Professional Therapeutics







What IS SIBO?

- Overgrowth of bacteria in the small intestine that normally should not reside there in significant quantities
- Bacteria interfere with normal digestion and nutrient absorption, and produce gases that lead to common IBS symptoms, such as gas, bloating, diarrhea/constipation
- A number of causes can lead to development of SIBO, and SIBO is associated with a variety of conditions

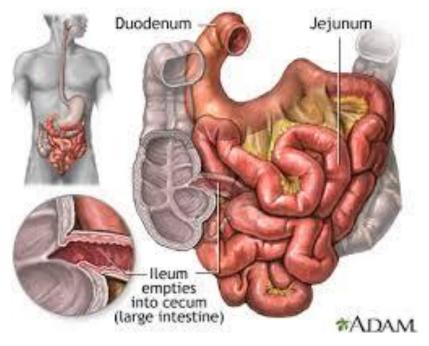


SIBO Defined

- No single accepted definition
- ♦ Previous gold standard = presence of ≥1 × 10⁵ bacteria per mL of proximal SI aspiration
 - Others estimate normal subjects rarely exceed 1×10^3 cfu/mL
- In research and clinical practice, diagnosis often made indirectly through hydrogen/methane breath testing

Anatomy Review

- Normal SI has low bacteria counts
 - Above SI → HCl in stomach prevents overgrowth
 - Below SI → Ileocecal valve separates colon from SI
 - Migrating motor complex (MMC) periodically sweeps through the 'cleanse' SI of bacteria
- SIBO characterized by high levels of normal colonic bacteria that ferment carbs → produce gas
 - Bacteria are not pathogenic!





Disease or Symptoms?

- For a long time patients with abdominal pain, gas, bloating, and diarrhea/constipation were diagnosed with IBS
- ND's have many tools for helping with IBS
 - Food allergy/sensitivity testing or elimination diet
 - Enzymes, probiotics, omega-3s, etc
 - Stress reduction

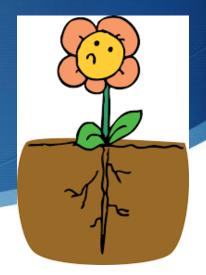


Many would get better – but what about those that don't improve with the basics...?

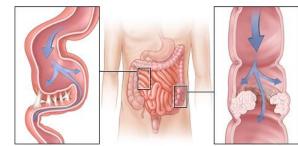
SIBO – Why Should You Care?

- GI complaints are one of the most common reasons for doctors visits (both MD and ND)
 - IBS is the most common of all GI disorders
- Some studies estimate SIBO to be the underlying cause in 84% of IBS cases
 - Other estimates as low as 30% of IBS cases likely somewhere in between (50-75% of cases)
- Related to MANY other common conditions
 - Prevalence in celiac disease estimated at 50%
- Elderly may be more susceptible
 - Low HCl, polypharmacy

Tolle Causam Treat the Cause



- Bacterial overgrowth occurs when intestinal stasis gives bacteria the opportunity to proliferate locally, such as due to mechanical stasis following bowel surgery
 - Other prominent causes of stasis include diabetes, scleroderma, intestinal diverticulosis, and intestinal obstruction caused by strictures, adhesions, cancer
- Certain medications may also predispose to SIBO, such as PPIs and opiates

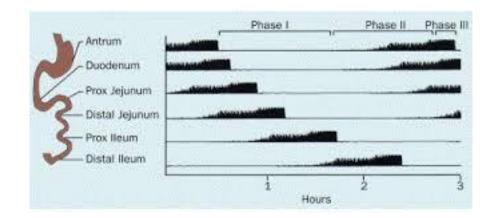


Scar tissue on small intestine

Cancer in colon



- Dysfunction of the Migrating Motor Complex (MMC)!
 - Due to **gastroenteritis**, diabetic neuropathy, hypothyroidism, sclerosis, nerve damage, opiates, surgery, stress

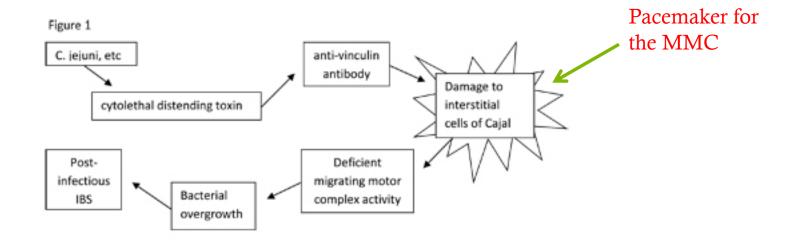


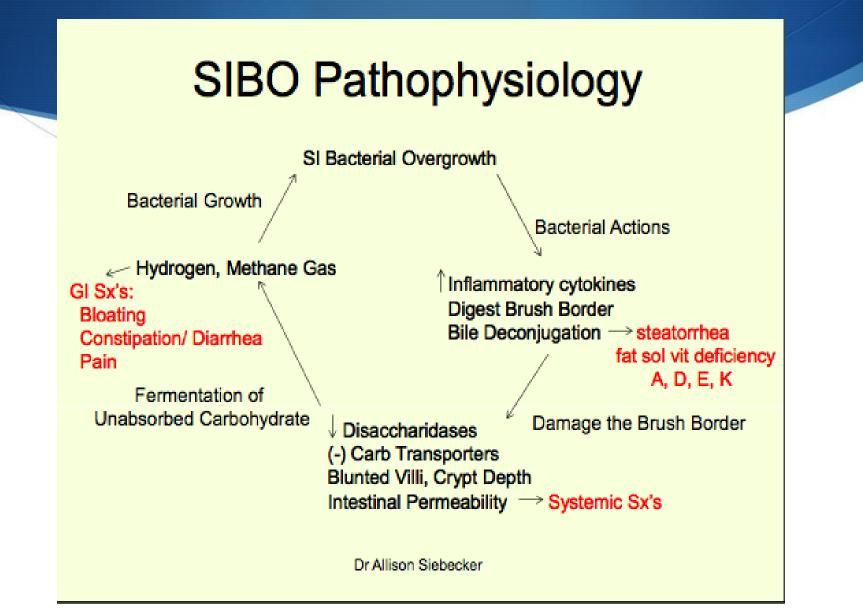


- SIBO often develops following bout of gastroenteritis
 - Estimated 7-31% of GI infxn will develop PI-IBS (SIBO)
- Bacteria (i.e. Campylobacteri jejuni) secrete cytolethal distending toxins (CDTs) that impair muscle & nerve connections and inactivate the MMC
- ♦ CDTb through molecular mimicry forms Abs to a cytoskeletal protein called vinculin → our immune system damages SI nerves and pacemaker cells while trying to attack the CDTb



• Organisms that can trigger PI-IBS include *Campylobacter, Salmonella, Shigella, E. coli, Giardia*, and certain viruses





Clinical Manifestations

• Non-specific/IBS Sx:

• Distension, flatulence, abd pain/discomfort, diarrhea, constipation

Malabsorption

- More in severe cases
- Def. of fat soluble vits, B12, iron

• Systemic Sx:

• Can result from increased intestinal permeability ("leaky gut")

Note: Symptoms can be distorted by underlying cause (i.e. scleroderma, diabetes, hypothyroid)

Passing Gas



- Hydrogen and methane gases are not normally produced by human cells
 - Hydrogen \rightarrow diarrhea
 - Methane (produced by archaea) \rightarrow constipation
- Treatment will differ depending on presence and levels of gases

A Third Gas...?

- ♦ Hydrogen sulfide gas can also be produced by sulfate-reducing bacteria → convert H₂ to H₂S
- Not as well characterized as Hydrogen and Methane gas, as it is not detected on standard breath tests
- Clues for H₂S overgrowth:
 - "rotten egg" smelling flatulence
 - sensitivity to sulfur containing foods



• SIBO sx but "flat-line" hydrogen and methane levels on breath test

Associated Conditions

- Acne Rosacea
- Acne Vulgaris
- Acromegaly
- Age
- Alcohol Consumption (moderate intake),
- Anemia
- Atrophic Gastritis
- Autism
- Celiac Disease
- Cystic Fibrosis
- Chronic Fatigue Syndrome
- Diabetes
- Diverticulitis
- Dyspepsia
- Fibromyalgia
- Fructose Malabsorption
- Gallstones
- Gastroparesis
- GERD
- ♦ HIV
- Hepatic Encephalopathy

Hepatic Injury

۵

۵

٨

۸

- H pylori Infection
- Hypochlorhydria
- Hypothyroid/Hashimoto's Throiditis
- IBD (Inflammatory Bowel Disease)
 - -Crohn's
- -Ulcerative Colitis
- IBS (Irritable Bowel Syndrome)
- Interstitial Cystitis
- Lactose Intolerance
- Leaky Gut (Intestinal Permeability)
- Liver Cirrhosis
- Lyme
- Malabsorption Syndrome
- Medications: Proton Pump Inhibitors, Narcotics/Opioids, NSAIDS
- Muscular Dystrophy (myotonic Type 1)
- Myelomeningocele (spina bifida)

- NASH/ NAFLD (non-alcoholic steatohepatitis/non-alcoholic fatty liver
- Obesity
- Pancreatitis
- Parasites
- Parkinson's
- Pernicious Anemia
- Prostatitis (chronic)
- Radiation Enteropathy
- Restless Leg Syndrome
 - Rheumatoid Arthritis
 - Scleroderma (Systemic Sclerosis)
- Short Bowel Syndrome
- Surgery: Abdominal, Post Gastrectomy, Post Esophageal and Gastric Cancer, Post-Cholecystectomy
- Tropical Sprue
- Whipple's Disease



- IBS develops following acute infectious gastroenteritis
- IBS sx improve with antibiotic use
- Worsening of IBS sx with prebiotics (FOS) or fiber intake
 - *Constipation that is worse with fiber*
- Celiac pt that does not improve on gluten-free diet
- IBS sx and chronic low ferritin with no other known cause

SIBO Diagnosis

- Stool testing is of little/no value
- Endoscopy/aspiration expensive and invasive, only samples proximal SI, contamination
- Best and most widely used test is hydrogen/methane breath test
 - Detects presence of hydrogen and/or methane gas produced in response to lactulose solution





- Lactulose vs Glucose
 - Glucose rapidly absorbed in duodenum so may not detect distal SIBO
 - Avg transit through SI is ~120 minutes, so lactulose gives indication of distal SI
- Samples taken q20 minutes after baseline
- 3 Hour time should show second peak when lactulose reaches the LI and is fermented



Breath Test - Preparation

Prep Diet

- Meat/fish/poultry
- White rice
- Eggs
- Hard Cheese
- Clear beef or chicken broth
- Oil
- Salt and pepper
- *12 hour fasting before test

What about vegetarian/vegan patients?







Breath Test

<u>https//sibocenter.com/faqs/</u>

Instructions for taking the test.





Positive test considered:

- a rise over baseline in hydrogen production of 20 parts per million (PPM) or greater within 120 minutes after ingesting the test substrate
- a rise over baseline in methane production of 12 ppm or greater within 120 minutes after ingesting the test substrate
- a rise over baseline in the sum of hydrogen and methane production of 15 ppm or greater within 120 minutes after ingesting the test substrate

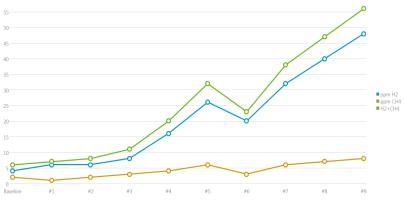
Breath Test Example

Sample	Time	ppm H ₂	ppm CH ₄	Total H2 + CH4	CO ₂ Check	50 45	<u> </u>
1. Baseline	10:30	_	4	5	OK	40	
2. 20 min	10:52		4	5	OK		
3. 40 min	11:12		3	4	OK	35	
4. 60 min	11:32		2	10	OK	ਤੱ ³⁰	
5. 80 min	11:52		1	10	OK	P 25	
6. 100 min	12:12		3	19	OK	ā	
7. 120 min	12:32	23	5	28	OK	1 ^{° 20}	
8. 140 min	12:52	24	4	28	OK	15	
9. 160 min	13:12	33	4	37	OK	10	
10. 180min	13:32	40	6	46	OK		
						5	
	-	lotes				0 1 2 3 4 5 6 7 8 9 Sample Number	10
11:47 mild stoma							
12:24 mild stoma							
13:07 mild stoma	ch pain and norn	nal stool				Total H2 + CH4	
						Result Flag	Normal
						Combined baseline total = 5	<20PPM
						Greatest H ₂ increase within first 120 minutes = 22 H	<20PPM
						Greatest H ₂ level within first 120 minutes = 23 H	<20PPM
						Greatest CH ₄ increase within first 120 minutes = 4	<12PPM
						Greatest CH ₄ level within first 120 minutes = 5	<12PPM
						Greatest combined H ₂ & CH ₄ increase within first 120 mins = 24 H	<15PPM
						Methane Producer = 6 H	<2PPM

Breath Test Example #2

Small Intestinal Bacterial Overgrowth (Lactulose) Analytical Record

Patient:	DoeJane						
Patient ID:	SAMRE-1						
DOB	02/04/1995						
Weight (at collection)	:						
Substrate given:	Lactulose						
Samples collected:	03/04/2016						
Samplesanalyzed:	03/07/2016						
Nurse/Technician:							
Referring physician:							
Notes							
Pre-test notes							
SWELERFORT1							
Pre-test symptoms Neussa, Vomiting, Weight Loss, Diamhea, Boating							
In-test notes							



Diagnosis/recommendation:

Sample	Time	ppmH2	ppm CH4	ppmH2+CH4	%002	Correction	Symptoms
#0-0	8:00 AM	4	2	6	32	1.71	
#1 - 20	8:20 AM	6	1	7	3.3	1.66	
#2-40	8:40 AM	6	2	8	3.1	1.77	
#3-60	9:00 AM	8	3	11	3.4	1.61	
#4-80	920 AM	16	4	20	3.1	1.77	
#5 - 100	9:40 AM	26	6	32	2.9	1.89	
#6-120	10:00 AM	20	3	23	4.1	1.34	
#7 - 140	10:20 AM	32	6	38	3.5	1.57	
#8-160	10:40 AM	40	7	47	3.9	1.41	
#9-180	11:00 AM	48	8	56	4.3	1.27	

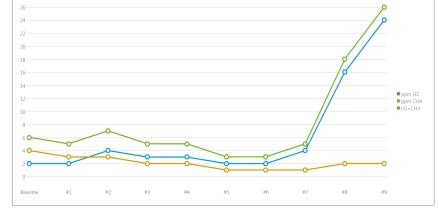
Physician Signature:

Date:

Breath Test Example #3

Small Intestinal Bacterial Overgrowth (Lactulose) Analytical Record

Patient:	John Doe
Patient ID:	SAMPLE-2
DOB	01/10/1970
Weight (at collection):	
Substrate given:	Lactulose
Samplescollected:	03/02/2016
Samplesanalyzed:	03/07/2016
Nurse/Technician:	
Referring physician:	
Notes	
Pre-test notes SAMPLEREPORT2	
Pre-test symptoms Nausea, Weight Gain, O	onstipation, Bloating



Diagnosis/recommendation:

Sample	Time	ppmH2	ppmCH4	ppmH2+CH4	%002	Correction	Symptoms
#0-0	7:20 AM	2	4	6	3.9	1.41	
#1 - 20	7:40 AM	2	3	5	4.5	1.22	
#2-40	8:00 AM	4	3	7	4.7	1.17	
#3-60	8:20 AM	3	2	5	3.4	1.61	
#4-80	8:40 AM	3	2	5	4.8	1.14	
#5-100	9:00 AM	2	1	3	4.6	1.19	
#6-120	9:20 AM	2	1	3	3.8	1.44	
#7 - 140	9:40 AM	4	1	5	4.5	1.22	
#8-160	10:00 AM	16	2	18	4.4	1.25	
#9-180	10:20 AM	24	2	26	27	2.03	

Physician Signature:

In-test notes

Date:



- Important to first identify if hydrogen and/or methane gases present
- May include liver support preparation stage (1-2 weeks) to reduce die-off reaction
- Then begin eradication phase (2-4 weeks) using pharmaceutical/herbal antimicrobials
- Then SIBO diet following eradication
 - Also include prokinetic agents

1. Eradication Phase – Rifaximin (Xifaxan)

- Most commonly used antibiotic
 - 550mg bid/tid x 14 days
- **Pros**:
 - Effective at eradicating SIBO
 - Not absorbed, so little/no s/e
 - Can be used on it's own for hydrogen(+)
- **Cons**:
 - Needs to be combined with Neomycin (or other a/b, or allicin) for Methane(+) SIBO (constipation)
 - Expensive, may be difficult to find



1. Eradication Phase – Herbal



- Many consider herbal antimicrobials as effective as Rifaximin
- However, needs to be used for longer time (4 weeks vs. 2 weeks)
- And may see longer or more severe die-off reaction

Options include:

- Berberine-containing herbs (Coptis chinesis. Berberis sp., Hydrastis)
- ♦ Allium sativum*



- Oregano
- Clove
- Neem
- Cinnamon

May be best to rotate (i.e. herb combo for 2 weeks, then switch to different herbs for 2 weeks)

2. Dietary Phase

- Pearl: Do NOT start SCD/SIBO diet during the eradication phase!
 - Research shows that SIBO eradication is improved when a/b tx combined with fiber in diet

"Happy bacteria as easier to kill"

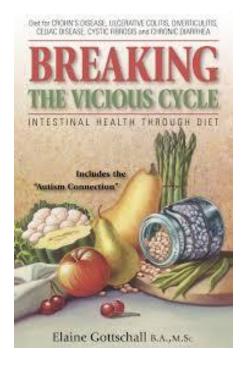


• Wait until after eradication phase (2-4 weeks) to begin diet

Tx – Diet Options

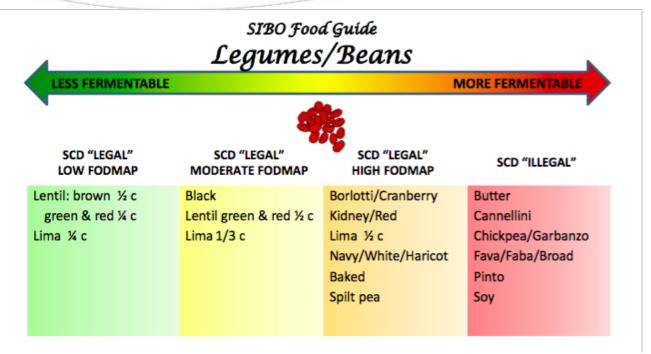
- There is no one-size fits all diet for SIBO
- Most common are SCD, GAPS, low fodmap, or combination diet
- May be able to introduce foods after a period of time (i.e. 2-3 months) and see how patient reacts
- Elemental diet
 - Effective, but costly and poor taste

Specific Carbohydrate Diet



- Developed by Elaine Gottschall
 - "Breaking the Vicious Cycle" book and website are good resources
- Aim is to cut out specific carbs (disaccharides) to starve bacteria
- Avoids grains, starches, dairy

SIBO Diet



www.siboinfo.com/diet.html

Tx - Prokinetics

- Pearl: Key to preventing recurrence is stimulating migrating motor complex (MMC)
 - Begin after eradication phase (use along with diet)
- Pharma options include:
 - Low dose erythromycin (50mg QD hs)
 - Low dose naltrexone (2.5 mg q.h.s. for IBS-D or 2.5 mg b.i.d. for IBS-C)
 - Prucalopride may be better for constipation than diarrhea
- Other options:
 - ♦ 5-HTP
 - D-limonene (consider if GERD also present)
 - Probiotics: *Bifidobacterum lactis, Lactobacillus rhamnosus*
 - Iberogast
 - Fasting in between meals / overnight stimulates MMC



Be sure to treat the cause (i.e. stimulate MMC, HCl, dec stress, etc)

- Betaine HCl
 - If hypochlorhydria suspected as part of underlying cause
 - Careful if methane(+) though
- Gut healing
 - L-Glutamine
 - Zinc carnosine
- Probiotics
 - Make sure without FOS

Thank You

This concludes the CE portion of the webinar



- Prep Phase
- Eradication Phase
- **Maintenance Phase: Prokinetics & Recurrence Prevention**
- Product Information
- ♦ Q&A

Protocol

- **Prep Phase** (1-2 weeks prior to Eradication Phase)
 - Hepasylin
 - Liver Support to minimize die-off reaction
 - Take 2 capsules QD AC
 - Glass of lemon water in the morning
 - Castor oil packs daily

Protocol

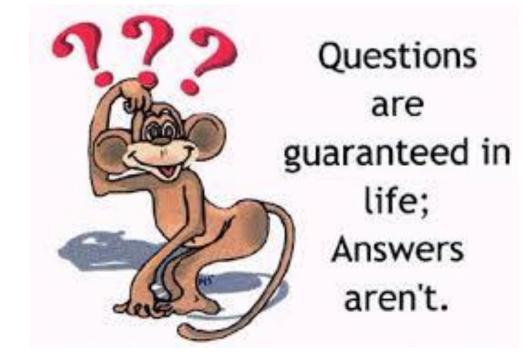
• **Eradication Phase** (4 weeks)

- **Microcidin** 1 capsule TID with food.
- Supreme-PB30+ DF (without FOS) Start with 1 capsule HS for 1 week.
 - If tolerated well, increase to 2 capsules HS from Weeks 2-4.
 - If NOT tolerated well, use **S. boulardii**, 2 capsules HS
- **Hepasylin** (Liver Support to alleviate die-off reaction):
 - Weeks 1-2 of protocol take 3 capsules QD AC
 - Weeks 3-4 of protocol take 2 capsules QD AC

Protocol

- Maintenance Phase:
 Prokinetic & Recurrence Prevention (3 months +)
 - Diet:
 - If diet compliance is an issue, incorporate digestive enzymes.
 - **Zyme-Aid Carbo Fort** 1 capsule CC, TID.
 - **5-HTP** 1 cap (100 mg) 1 hour after each meal, and 1 cap hs
 - D-Limonene for patients with GERD 12 drops on empty stomach BID or 24 drops HS.
 - ▲ L-Glutamine Plus 1 teaspoon QD (3 g L-glutamine + Vitamins A/C/E)
 - Supreme-PB30+ DF (without FOS) 1 capsule 2 hours after lunch, 1 capsule HS.





Microcidin

- Contains **multiple antimicrobial ingredients** with different mechanisms to inhibit and destroy the pathogenic microbials, such as bacteria, virus, fungi (eg. Candida albican) and parasites.
- Coptis (Huang Lian) contains high berberine. Coptis is the king herb for detoxification in Traditional Chinese Medicine, especially during infections and Damp Heat. In modern medicine, coptis itself or combined with clove extract have been proven to inhibit fungal growth and candidiasis.
- Contains lab-standardized high allicin content from freeze-dried garlic concentrate. Allicin is a potent natural antibiotic that does not cause dysbiosis.
- Synergized with highly concentrated **oregano** extract, **undecylenic acid** and **caprylic acid** for broad spectrum microcidal effect in prevention and eradication of microbial infections.

Supreme-PB30+ DF (without FOS)

- Dairy-Free probiotic formula that contains 55 billion viable cells comprised of 6 species of human gut-anchoring probiotics to ensure successful establishment in each of their particular niche.
- Includes Bifidobacterium lactis Bl-04 & Lactobacillus rhamnosus Lr-32
- Carefully selected strains with complete resistance tests (22 antibiotics)
- All strains are **acid- and bile-resistant**, ensuring their passage through the entire GI tract while helping inhibit H. pylori infections.
- Clinically proven to anchor and colonize at the human gut linings to improve dysbiosisassociated symptoms (eg. constipation, diarrhea, bloating, and allergic reactions) caused by antibiotics and other GI disorders.

Limonen-E (d-limonene liquid)

• D-limonene has been shown to be effective in relieving occasional heartburn and gastroesophageal reflux disorder (GERD).

• Ultra-potent D-Limonene (>99.5%)

 1 g of D-limonene (~24 drops) daily or every other day has been clinically shown to achieve complete relief of symptoms

Hepasylin

- Formulated with milk thistle, dandelion, artichoke and alpha lipoic acid to protect the hepatocytes and maintain healthy liver function.
- Contains milk thistle extract standardized by not only silymarin (80%) but also silybin (30%) - the most active compound of silymarin group - to guarantee the maximum liver-protecting effect.
- Increases the reduced glutathione concentration in the liver, promotes bile flow, provides powerful antioxidants for cell protection, and enhances the liver's ability to detoxify

Webinar Promotion

- Vita Aid will be offering 10% off the following products until March 31st 2016:
 - Microcidin
 - ♦ 5-HTP
 - Supreme-PB30+ DF (without FOS)
 - ♦ Limonen-E
 - Hepasylin

Thank You!

• Special thank you to Dr. Ibby Omole

References

- ♦ Furnari, Manuele, et al. "Clinical trial: the combination of rifaximin with partially hydrolysed guar gum is more effective than rifaximin alone in eradicating small intestinal bacterial overgrowth." *Alimentary pharmacology & therapeutics* 32.8 (2010): 1000-1006.
- Furnari, M., V. Savarino, and E. Savarino. "Letter: treatment for small intestinal bacterial overgrowth–where are we now?." *Alimentary pharmacology & therapeutics* 39.4 (2014): 442-442.
- Shah, Shailja C., et al. "Meta-analysis: antibiotic therapy for small intestinal bacterial overgrowth." *Alimentary pharmacology & therapeutics* 38.8 (2013): 925-934.
- Grover, Madhusudan, et al. "Small intestinal bacterial overgrowth in irritable bowel syndrome: association with colon motility, bowel symptoms, and psychological distress." *Neurogastroenterology & Motility* 20.9 (2008): 998-1008.
- Lauritano et al. "Rifaximin dose-finding study for the treatment of small intestinal bacterial overgrowth." *Alimentary pharmacology & therapeutics* 22.1 (2005): 31-35.
- Bijkerk, C. J., et al. "Systematic review: the role of different types of fibre in the treatment of irritable bowel syndrome." *Alimentary pharmacology & therapeutics* 19.3 (2004): 245-251.
- Hofmann, Alan F., and Lars Eckmann. "How bile acids confer gut mucosal protection against bacteria." *Proceedings* of the National Academy of Sciences of the United States of America 103.12 (2006): 4333-4334.
- Chedid, Victor, et al. "Herbal therapy is equivalent to rifaximin for the treatment of small intestinal bacterial overgrowth." *Global Advances in Health and Medicine* 3.3 (2014): 16-24

- Thompson, John Richard. "Is irritable bowel syndrome an infectious disease?." *World journal of gastroenterology* 22.4 (2016): 1331.
- Rezaie, Ali, Mark Pimentel, and Satish S. Rao. "How to Test and Treat Small Intestinal Bacterial Overgrowth: an Evidence-Based Approach." *Current gastroenterology reports* 18.2 (2016): 1-11.

۵

۵

۵

۵

۵

۵

- Ghoshal, Uday C., et al. "A proof-of-concept study showing antibiotics to be more effective in irritable bowel syndrome with than without small-intestinal bacterial overgrowth: a randomized, double-blind, placebocontrolled trial."*European journal of gastroenterology & hepatology* (2016).
- Steven Sandberg-Lewis, ND, DHANP and Allison Siebecker, ND, MSOM. *SIBO: Dysbiosis Has A New Name.* Townsend Letter March 2015.
- Yarandi, Shadi Sadeghi, et al. "Overlapping gastroesophageal reflux disease and irritable bowel syndrome: increased dysfunctional symptoms." *World J Gastroenterol* 16.10 (2010): 1232-8.
- Dukowicz, A.C. and E. Lacy. "Small intestinal bacterial overgrowth: A comprehensive review." Gastroenterology & Hepatology Vol. 3, No. 2 (2007): 112–122.
- Pimentel M, Park S, Mirocha J, Kane SV, Kong Y. The effect of a nonabsorbed oral antibiotic (rifaximin) on the symptoms of the irritable bowel syndrome: a randomized trial. Ann Intern Med. 2006;145(8):557-563.
- Sachdev, Amit H., and Mark Pimentel. "Gastrointestinal bacterial overgrowth: pathogenesis and clinical significance." *Therapeutic advances in chronic disease* 4.5 (2013): 223-231.

Patil, Anant D. "Link between hypothyroidism and small intestinal bacterial overgrowth." *Indian journal of endocrinology and metabolism* 18.3 (2014): 307.

- Saad, Richard J., and William D. Chey. "Breath testing for small intestinal bacterial overgrowth: maximizing test accuracy." *Clinical Gastroenterology and Hepatology* 12.12 (2014): 1964-1972.
- Quigley, Eamonn MM. "Small intestinal bacterial overgrowth: what it is and what it is not." *Current opinion in gastroenterology* 30.2 (2014): 141-146.
- Grace, E., et al. "Review article: small intestinal bacterial overgrowth-prevalence, clinical features, current and developing diagnostic tests, and treatment." *Alimentary pharmacology & therapeutics* 38.7 (2013): 674-688.
- Pimentel, Mark, et al. "Development and validation of a biomarker for diarrhea-predominant irritable bowel syndrome in human subjects." *PloS one*10.5 (2015): e0126438.

۵

۵

٨

Miyano, Yuki, et al. "The role of the vagus nerve in the migrating motor complex and ghrelin-and motilin-induced gastric contraction in suncus." *PloS one* 8.5 (2013): e64777.