Gut Health – Looking Beyond the Hype

Graham MacKenzie Ph.C Stone's Pharmasave, Baddeck

Author: Healthy Logic, Gut Logic



1

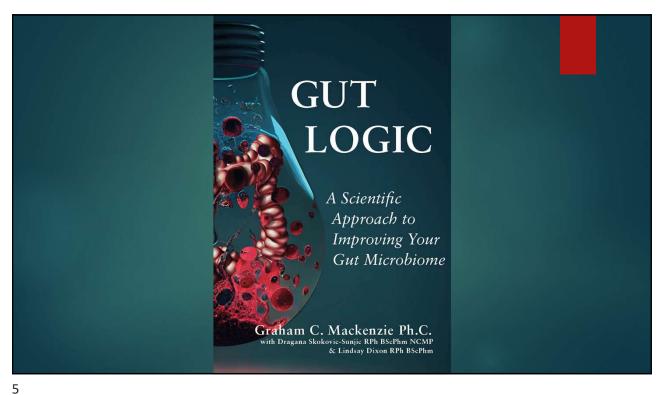
Presenter Disclosure

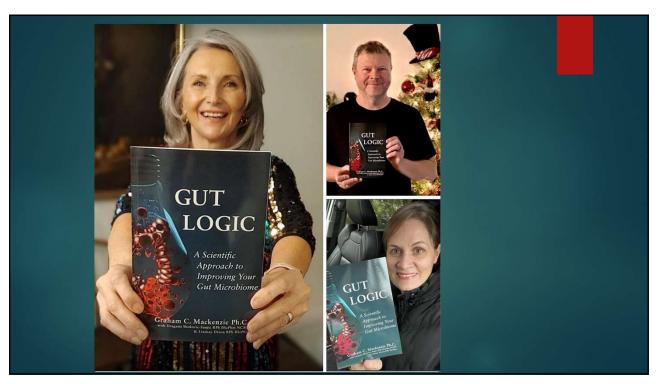
- ▶ No potential or perceived conflict of interest
- ▶ No financial or in-kind support from any commercial or other organization other than an honorarium from Dalhousie to present this webinar
- ▶ This program has no industry involvement but will involve using commercial brand names for clarity in giving recommendations.



Learning Objectives

- ▶ Learn the basics of probiotic nomenclature and terms
- ▶ Become familiar with using AEProbio.com in making probiotic recommendations
- ▶ Challenges with probiotic recommendations
- ▶ Understand brand specific recommendations for various conditions and the evidence behind them
- ▶ Learn the best way to take probiotics and when not to
- ▶ Understand basic information on what prebiotics are and what they do for gut health







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Pick a side

- ▶ A) For the most part, I recognize the value probiotics and recommend them quite often
- ▶ B) For the most part I am too skeptical of probiotics and rarely recommend them
- C) I rarely recommend probiotics mainly because I don't know enough about them to feel comfortable recommending them

Naming a Probiotic

- ▶ genus species strain
- ▶ Bifidobacterium longum CECT 7347
- ▶ Genus Bifidobacterium
- ▶ Species longum
- ▶ Strain CECT 7347
- ► CFU's

| GENUS & SPECIES | STRAIN | BRAND | |
|---|-------------------------|-----------|--|
| Bifidobacterium lactis | CNCM I-2494 | Activia | |
| Bifidobacterium infantis | 35624 | Align | |
| Lactobacillus rhamnosus | HA111 | Bacid | |
| Lactobacillus reuteri | DSM 17938 | Biogaia | |
| Lactobacillus acidophilus | CL1285 | | |
| Lactobacillus casei | LBC80R | Bio-K | |
| Lactobacillus rhamnosus | CLR2 | | |
| Lactobacillus casei | CNCM I-1518 | DanActive | |
| Saccharomyces boulardii | lyo | Florastor | |
| Several genus and species | Several strains | Probaclac | |
| Lactobacillus plantarum | 299v | TuZen | |
| Several genus and species | Several defined strains | VSL#3 | |
| Bifidobacterium lactis Lactobacillus acidophilus | BB-12 LA-5 | Yoptimal | |

Essential Terms

- ▶ Probiotic live microorganisms that, when administered in adequate amounts, confer a health benefit on the host
- Prebiotic Source of food and nutrition for the healthy bacteria of the gut
- ▶ Postbiotic non-living substances/remnants produced by microorganisms. Cell wall fragments, vitamins, metabolites, proteins, peptides.
- Synbiotic a mixture of probiotic and prebiotic
- ► Microbiota the entire mix of microbiome in the body including bacteria, virus, parasites and fungi.

11

Checking Health Canada's Information on Specific probiotic (Florastor) falling under Drugs

https://www.canada.ca/en/health-canada/services/drugs-health-products/natural-non-prescription.html

Recommended use or purpose:

Helps prevent antibiotic associated diarrhea. Helps reduce the symptoms related to acute diarrhea (infectious). Source of Probiotics. Helps support intestinal/gastrointestinal health. Could promote a favourable gut flora. Helps to reduce the risk of antibiotic-associated diarrhea. Helps reduce the symptoms related to acute infectious diarrhea. Helps to reduce the risk of traveller's diarrhea. Helps to treat acute infectious diarrhea in children. An adjunct to physician-supervised antibiotic therapy in patients with Helicobacter pylori infections. Helps reduce recurrent Clostridium Difficile Associated Diarrhea (CDAD) when used as an adjunct to antibiotics.

Risk Information:

Cautions and Warnings

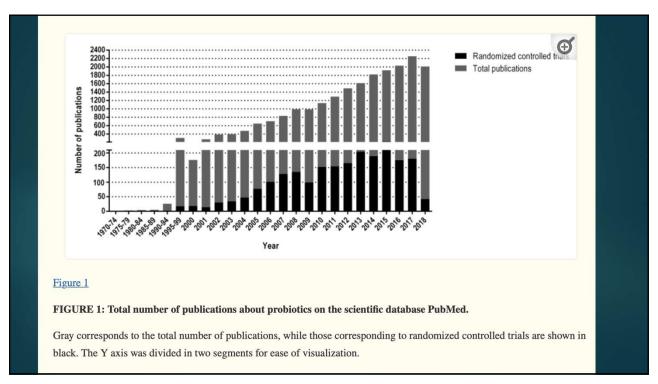
Discontinue use and consult a health care practitioner if symptoms of digestive upset (e.g. diarrhea) occur, worsen or persist beyond 3 days. Consult a health care practitioner prior to use if you have an abnormal condition of the gastrointestinal tract lining (i.e. ulcer or other GI tract disorders). The concurrent administration of Saccharomyces boulardii lyo and intestinally acting antimycotics may decrease the efficacy of Saccharomyces boulardii lyo. Treatment does not replace rehydration when this is necessary.

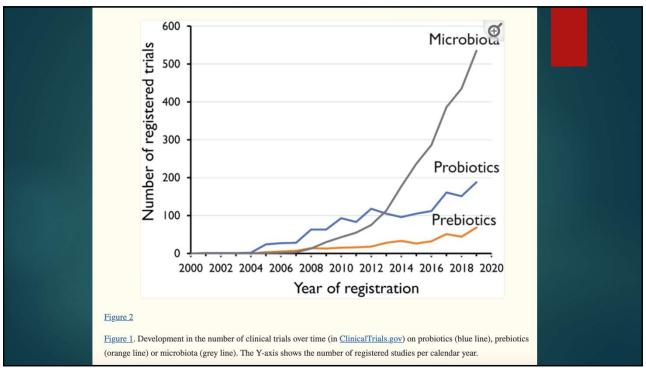
Contra-Indications

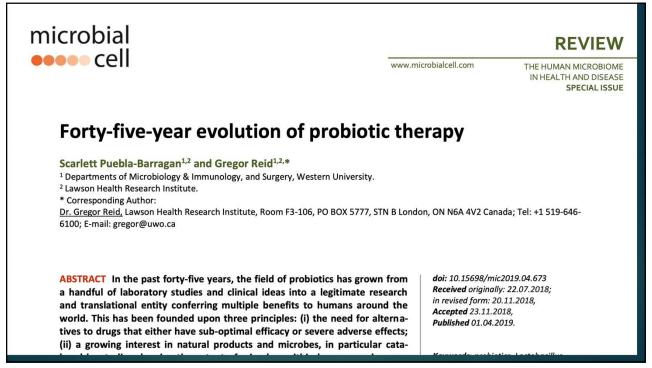
Do not use in the acute-care setting (ICU, CCU, NICU, etc.), with critically ill patients, especially in those individuals with open arterial and venous access required (such as a Vascath, Permoath, or AV fistula). Do not use for any individual with a central line or port or in the surroundings of any patient with a central line or port. Central lines include short- and long-term central venous catheters (CVCs) and peripherally inserted central catheters (PICCs). Do not use if you are severely immunocompromised, have an immune-compromised condition (e.g. AIDS, lymphoma, long-term corticosteroid treatment, pre- or post- organ/bone marrow transplant). Do not use and consult a health care practitioner if you are experiencing nausea, fever, vomiting, bloody diarrhea or severe abdominal pain. Do not use if you have hypersensitivity to any component of the product (especially allergy to yeast).

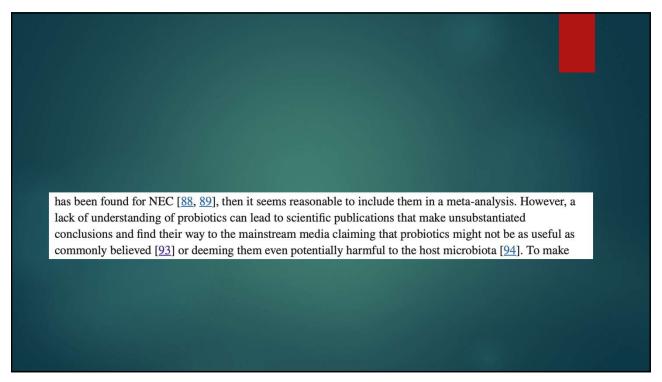
Probiotics may also fall under Food with Health Canada

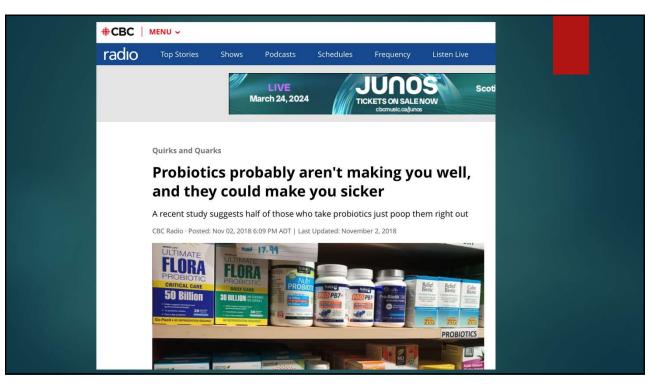
https://www.canada.ca/en/health-canada/services/foodnutrition/food-labelling/health-claims/accepted-claims-aboutnature-probiotic-microorganisms-food.html











Are probiotics today even worth taking?

According to Brett Finlay from the University of British Columbia, there are a few health ailments that probiotics available on the market today that have been scientifically proven to work. Examples might be antibiotic associated diarrhoea, eczema, and some cases of colic in babies.

"But the problem is, there's a bewildering number of probiotics."

He suggested if people want to be smart consumers when it comes to buying probiotics, they can check the Clinical Guide to Probiotic Products Available in Canada. The site charts clinical evidence for the efficacy of specific probiotics for specific illnesses, and has been reviewed by a team of health professionals.

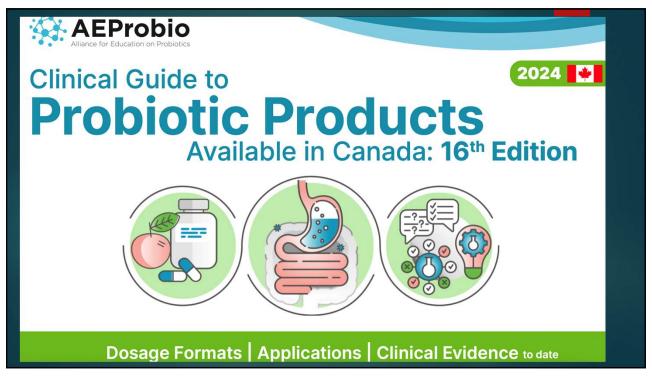
(f) The idea that] taking one probiotic that is going to cure all and make you feel better and [is] just wonderful wonder drug, that doesn't exist.

- Dr. Brett Finlay, UBC

"I think if people are serious about probiotics, they really should take the time to look at this site and say, "What is the clinical evidence for a particular probiotic and a particular condition," said Finlay.

19

Many healthcare professionals have had the same questions you have about probiotics over the last few decades. Most of us were amazed by the published research and evidence growing exponentially year after year, looking at the potential for probiotics as an intervention. However, most of us were confused with the question: How do I know what probiotics to recommend? This particular study is telling me I should be recommending this complicated-sounding strain of Lactobacillus rhamnosus with a series of letters and numbers behind it. If I gave this name to my patient, would they know what product to buy?





| AAD | Antibiotic associated diarrhea - Prevention | ID | Infectious diarrhea |
|--------|---|------------------|--|
| BV | Bacterial vaginosis | LDL-C | Reduces LDL and total cholesterol |
| C | Constipation | LH | Liver Health (NASH/NAFLD/MHE; as adjunct to standard therapy; see studies for specific population) |
| CDAD | Clostridium difficile associated diarrhea - Prevention | M/A | Mood and Affect (symptoms related to stress/anxiety; not a substitute for standard treatment) |
| CE/AD | Childhood eczema/ Atopic dermatitis | Mastitis | Mastitis in breastfeeding mothers (treatment and prevention) |
| CID | Common infectious disease - community acquired | NEC* | Necrotizing Enterocolitis (newborn) *as per hospital protocol, not for self-administration |
| СМРА | Cow Milk Protein Allergy (including Colic due to CMPA) | NI | Nosocomial infections prevention - hospital acquired |
| Colic | Colic | ОН | Oral health (reductions of tonsillitis, laryngitis, and dental caries) |
| FAP | Functional abdominal pain | Regurg/ GIMot | Reduces regurgitation/ Improves gastrointestinal motility |
| HP | Helicobacter pylori - Adjunct to standard eradication therapy | TD | Traveler's diarrhea prevention |
| IBD-P | Inflammatory bowel disease - Pouchitis | vc | Vulvovaginal candidiasis |
| IBD-UC | IBD - Ulcerative colitis - Adjunct to standard therapy | WM | Weight Management (aids in reduction of body weight, body fat mass, and waist circumference) |
| IBS | Irritable bowel syndrome | | |

Levels of Evidence **Level of Recommendation:** Evidence obtained from at least one appropriately designed trial (e.g. randomization, blinding, appropriate population Levell comparisons) with a power calculation for the outcome(s) of interest. The expert review board reserves the right to make the final decision for the level of recommendation. (HIGHEST LEVEL) Evidence obtained from well-designed controlled trials without randomization. Evidence obtained from randomized trials not **Level II** satisfying all criteria listed in Level I. Evidence obtained from well-designed cohort or case-control analytic studies, preferably from more than one center or research group. Evidence obtained from multiple time series with or without the intervention. Dramatic results in uncontrolled trials might also be regarded as this type of evidence. Opinions of respected authorities, based on clinical experience, descriptive studies, or reports of expert committees. Level III Readers are strongly encouraged to review the evidence listed for each product for themselves. Please refer to individual publications for specifics of patient populations studied (e.g. age, gender, comorbidities) and details of clinical effect (statistical significance, size of treatment effect, precision of the confidence intervals).



| Brand Name | Probiotic Strain(s) | Dosage Form | CFU/dose | No. of doses/day | O | ės ės | CDAD | BS | AAD | IBD-P | = |
|-------------------------------------|--|-------------------|--|-----------------------------|--------------------|-------------------|------------|-----------------------|---------|------------|-------------------|
| Purica Probiotic Intensive GI 🐇 | L. plantarum CECT7484/KABP022® L. plantarum CECT7485/KABP023® P. acidilactici CECT7483/KABP021® | Capsule | 3B/capsule | 1 capsule | | | | I ♠ (73,74) | | | |
| Ultra Probiotic Complex by GNC 🐇 | L. acidophilus CUL-60 L. acidophilus CUL-21 B. bifidum CUL-20 B. lactis CUL-34 | Packet Capsule | 25B/packet 25B/ 50B/ 75B per capsule | 1-2 packets 1-2 capsules | | | II (52) | II (53) | | | |
| UltraFlora® Balance | L. acidophilus NCFM® B. lactis Bi-07® | Capsule | 15B/capsule | 2 capsules | | | | II (78) | | | |
| UltraFlora® Intensive Care 🦸 | L. plantarum 299v | Capsule | 10B/capsule | 2 capsules | | | (32,36) | (33-35) | (31) | | |
| UltraFlora® Restore 🐧 | B. lactis Bi-07® L. acidophilus NCFM® B. lactis Bi-04 L. paracasei Lpc-37 | Capsule | 20B/capsule | 1 capsule | | | | | (75,76) | | |
| Visbiome® (| L. acidophilus DSM24735/SD5212 L. paracasei DSM24733/SD5218 L. delbrueckil subsp. bulgaricus DSM24734/SD5210 L. plantarum DSM24730/SD5209 B. longum DSM24736/SD5220 B. infantis DSM24737/SD5219 B. breve DSM24737/SD5219 S. thermophilus DSM24731/SD5207 | Sachet | 450B/sachet | 1-4 sachets | Ⅱ ♠ (81,92) | I⊕ (82-84) | | | | (83,85-87) | (88-91, 93,94) |

| Br | and Name | Probiotic Strain(s) | Dosage Form | CFU/dose | No. of doses/day | ė S S | £ | CDAD | BS | AAD | ę |
|-----|--|---|-------------------|-------------------------|-----------------------------|----------------|-----------------------|--------------------|--------------------|---------|---------|
| Flo | rastorMax® 🐇 | Saccharomyces boulardii Iyo CNCM | Sachet | 10B/sachet | 1 sachet | (43,47-50) | 1 % (44-46) | (8,9,38, 39,51) | | (37,38) | (40-42) |
| Flo | rastor® 🦸 | 1-745 | Capsule Sachet | 5B/capsule 5B/sachet | 1-2 capsules 1-2 sachets | (43,47-50) | (44-46) | (8,9,38, 39,51) | | (37,38) | (40-42) |
| | Forte 🦸 | L. acidophilus CUL-60 L. acidophilus CUL-21 L. acidophilus CUL-21 B. bilidum CUL-22 P Escherichia coli Nissle 1917 | Capsule | 10B/capsule | 2-3 capsules | | | (52) | II 👶 | | |
| | IBS Relief 🐇 | | Capsule | 25B/capsule | 1 capsule | | | (52) | H& | | |
| ш | Intensive 🐇 | | Capsule | 25B/capsule | 1 capsule | | | (52) | II 😂 (53) | | |
| HMF | Intensive 50 🐔 | | Capsule | 50B/capsule | 1 capsule | | | (52) | II 🚓 | | |
| _ | Intensive Powder 🕻 🏶 | | Powder | 25B/1 scoop (1 gram) | 1 scoop | | | (52) | II 🚓 (53) | | |
| | Super Powder 4 🕸 | | Powder | 10B/1 scoop (1 gram) | 2-3 scoops | | | (52) | Ⅱ ♣ (53) | | |
| Μι | itaflor® 🔆 | | Capsule | 2.5-25B/ capsule | 1-2 capsules | (57-60) | | | | | |
| Pre | egnancy Care Probiotic | | Capsule | 2B/capsule | 1-2 capsules | | | | III (69) | | |
| | obiotic 10 Billion Active Ils Daily Maintenance | Lactococcus lactis (UALI-08) L. gasseri (UALg-05) L. rhamnosus (UALI-06) B. animalis subsp. lactis (UABIa-12) B. breve (UABbr-11) L. paracasei (UALp-04) L. rhamnosus (UAL-18) L. acidophilus (DDS@-1) L. plantarum (UALp-05) B. longum subsp. longum (UABI-14) B. birlidum (UABB-10) L. casei (UALc-03) L. reuteri (UALre-16) B. longum subsp. lindits (UABI-13) | Capsule | 10B/capsule | 3 capsules | II (68) | | | | | |



Important quotes from Gut Logic

- ▶ " As a consumer, you may be left out in the cold, not knowing what brand of probiotic has the exact strains you need for a given ailment."
- ► "As I outlined in Healthy Logic (2021), my first priority, either with an OTC selection or a prescription medication, is that the therapy is safe. My second concern is that it works."
- ▶ "Just to put into context the difficulty in interpreting the results of these experiments and applying them to real-life recommendations, the microbial species Bifidobacterium longum is a microbial species, but there are over 400 strains of this species."

29

Table Of Contents

Introduction

(Contribution by Dragana Skokovic-Sunjic)

Chapter 1: The Logic about our Gut

Chapter 2: Exploring Upper Respiratory Tract Infections (URTI's)

Chapter 3: Understanding Diarrhea

Chapter 4: Insight into Irritable Bowel Syndrome (IBS)

Chapter 5: Nurturing Mental Health

Chapter 6: Bacterial Vaginosis/Vulvovaginal Candidiasis (yeast infection)

Chapter 7: Unveiling Urinary Tract Infections (UTI's)

Chapter 8: Cervical Cancer Awareness

Chapter 9: Strategies for Effective Weight Loss

Chapter 10: Managing Cholesterol Levels

Chapter 11: Tackling Eczema

Chapter 12: Exploring the World of Fermented Foods

and Prebiotic



► The most effective strains of probiotics require refrigeration

31

True or False

► The expiry date of refrigerated probiotics is typically longer than unrefrigerated probiotics

True or False

▶ Probiotics that are unrefrigerated on the shelf before purchased should be refrigerated once opened.

33

Select a specific strain for your particular problem.

Select the amount that is proven to work; do not go with the "the more, the merrier" idea.

Most of us do not need to take probiotics forever unless we have a chronic condition and our healthcare provider recommends it.

A wrong belief exists that refrigerated probiotics are better than those at room temperature. It all depends on the strain. Some strains are resilient and resistant to the influence of oxygen, light, and temperature and stay viable after exposure; other strains are very fragile and need to be protected, kept in dark bottles, specific formulations, and refrigerated to prolong their viability.

You do not need to get your gut tested to see what is missing. Each of us has a different blend and composition of the gut microbiome. Each of us has a different

AEProbio.com. The reason why this site is so valued, in my opinion, is because it does something that most other references do not. Dragana and her colleagues give the pedestrian reader the exact commercial product that corresponds to the strain or strains that have evidence of working. As I have explained to many health professionals and in this book, simply telling someone probiotics work or don't work for a medical condition means nothing without the actual product referenced as well. Most people will hunt forever for

35

Knowing the specificity of the recommendation

- ▶ 2008 The guide recommended Bacid (ERFA)
- ▶ Lactobacillus rhamnosus GG (10B CFU / capsule)
- ► Common hospital formulary inclusion
- ▶ 2009 The Guide learned the company changed the strain to Lactobacillus rhamnosus HA111.
- ▶ Health Canada requirement was to report the strain designation to them but no requirement to be on the label
- ▶ It continued to be dispensed in hospitals for AAD and CDAD even though there was no evidence for efficacy for the new strain
- ▶ Removed from The Guide in 2011. The original strain in Bacid was since sold as Culturelle in Canada

Knowing the specificity of the recommendation

- ▶ Similar situation with Danactive.
- ▶ Dropped strength from 10B to 1B CFU's per dose
- Insufficient to provide the health benefits as before

37

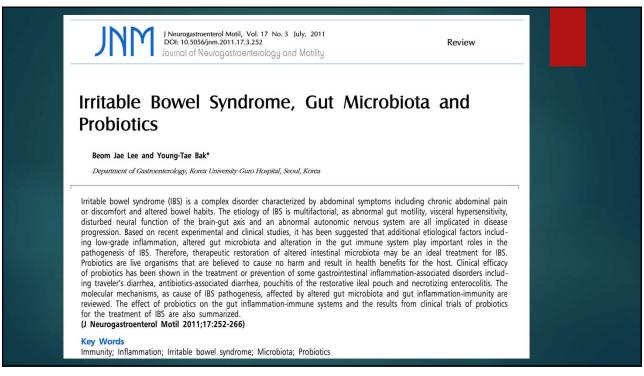
True or False

Contrary to popular belief, there are no indications for taking a probiotic in an otherwise healthy person



You should know who this recommendation of probiotics is for. It may surprise you to know that supplementing with probiotics isn't for everyone. Most people who are relatively healthy and symptom-free of bothersome issues really aren't going to benefit much from probiotics. There really is very little evidence to support taking random probiotics or even targeted probiotics, for that matter, for long-term health outcomes. This

39



Irritable Bowel Syndrome

- ▶ Symptoms well known, pathophysiology not as much
- Correspondingly, how probiotics work is hypothesized
- ▶ Microbiota between healthy and IBS patients shows a difference (some with SIBO and some not) – coliforms, lactobacilli and Bifidobacterium species lower in IBS. Varies geographically.
- ► Concentrations of short chain fatty acids, acetic acid and propionic acids are higher in IBS symptoms
- ▶ Non absorbed antibiotics (neomycin and rifaximin) reduce the concentrations of these products and can improve IBS flare ups.
- ▶ Align (1/day), BioK+IBS Pro (2/d), Ultraflora Intensive care(2/d)

41

RESEARCH ARTICLE Lactobacillus acidophilus CL1285, Lactobacillus casei LBC80R and Lactobacillus rhamnosus CLR2 improve quality-of-life and IBS symptoms: a double-blind, randomised, placebo-controlled study K. Preston ①, R. Krumian ①, J. Hattner ①, D. de Montigny ①, M. Stewart ①, S. Gaddam ① *Corresponding author: morgan.stewart@sprim.com Beneficial Microbes: 9 (5)- Pages: 697 - 706 https://doi.org/10.3920/BM2017.0105 References Cited by Full-text Abstract **Abstract** A combination of Lactobacillus acidophilus CL1285, Lactobacillus casei LBC80R and Lactobacillus rhamnosus CLR2 was compared to placebo for relief of symptoms of irritable bowel syndrome (IBS). A total of 113 subjects at 3 clinical sites were randomised in a 2:1 ratio and followed for 12 weeks. Subjects ingested either 2 capsules of active study product, containing 50×10⁹ cfu of live organisms, or 2 placebo capsules daily. Endpoints included improvement in abdominal pain, days of pain, distention, stool consistency and frequency, quality of life (QOL), and adequate relief (AR) of IBS symptoms. IBS subtypes constipation (IBS-C), diarrhoea (IBS-D), and mixed (IBS-M) were evaluated separately; the effect of gender was also examined. For all efficacy endpoints improvement of 30% or more vs placebo was considered clinically significant. With the exception of pain intensity and AR, the endpoints demonstrated a therapeutic advantage of active over placebo for IBS symptoms in at least some subject subgroups. The IBS-D and female subgroups showed the largest and most consistent effects. Stool frequency and consistency were evaluated in the IBS-C and IBS-D subgroups, and improvement of active vs placebo was noted in both. QOL improvement was seen overall and in specific domains. Adverse events (AEs) were limited to 7 subjects; all were of mild or moderate intensity except one, severe cramping. Four AEs in the same subject in the placebo group were judged to be related to study product; these resolved by the end of study. There were no serious AEs.

Traveller's Diarrhea

- ▶ Florastor (1-2/day) and
- ► Florastor Max (1/day)

Meta-analysis of probiotics for the prevention of traveler's diarrhea

Lynne V McFarland 1

Affiliations + expand

PMID: 17298915 DOI: 10.1016/j.tmaid.2005.10.003

Abstract

Background: Traveler's diarrhea (TD) is a common health complaint among travelers. Rates of TD can range from 5% to 50%, depending on the destination. The use of probiotics for this disease remains controversial. The objective of this study was to compare the efficacy of probiotics for the prevention of TD based on published randomized, controlled clinical trials.

Methods: PubMed, Google Scholar, metaRegister, NIH registry of clinical trials and Cochrane Central Register of Controlled Trials were searched from 1977 to 2005, unrestricted by language. Secondary searches of reference lists, authors, reviews, commentaries, associated diseases, books and meeting abstracts. Inclusion criteria included: randomization, controlled, blinded, efficacy trials, in humans, peer-reviewed journals. Exclusion criteria were: pre-clinical, safety, phase 1 studies in volunteers, reviews, duplicate reports, trials of unspecified probiotics, trials of prebiotics, and inconsistent outcome measures.

Results: Twelve of 940 screened studies met the inclusion and exclusion criteria. The pooled relative risk indicates that probiotics significantly prevent TD (RR=0.85, 95% CI 0.79,0.91, p<0.001).

Conclusion: Several probiotics (Saccharomyces boulardii and a mixture of Lactobacillus acidophilus and Bifidobacterium bifidum) had significant efficacy. No serious adverse reactions were reported in the 12 trials. Probiotics may offer a safe and effective method to prevent TD.

43

C.Diff and Antibiotic Associated Diarrhea Prevention

- ▶ BioK+ Antibio Pro (1-2/d)
- ▶ Bio-K+ Drinkable Probiotic (1/day)

Dose-response efficacy of a proprietary probiotic formula of Lactobacillus acidophilus CL1285 and Lactobacillus casei LBC80R for antibiotic-associated diarrhea and Clostridium difficile-associated diarrhea prophylaxis in adult patients

Xing Wang Gao ¹, Mohamed Mubasher, Chong Yu Fang, Cheryl Reifer, Larry E Miller

Affiliations + expand

PMID: 20145608 DOI: 10.1038/ajg.2010.11

Abstract

Objectives: Standard therapies for antibiotic-associated diarrhea (AAD) and Clostridium difficile-associated diarrhea (CDAD) have limited efficacy. Probiotic prophylaxis is a promising alternative for reduction of AAD and CDAD incidence.

Methods: In this single-center, randomized, double-blind, placebo-controlled dose-ranging study, we randomized 255 adult inpatients to one of three groups: two probiotic capsules per day (Pro-2, n=86), one probiotic capsule and one placebo capsule per day (Pro-1, n=85), or two placebo capsules per day (n=84). Each probiotic capsule contained 50 billion c.f.u. of live organisms (Lactobacillus acidophilus CL1285 +Lactobacillus casei LBC80R Bio-K+ CL1285). Probiotic prophylaxis began within 36 h of initial antibiotic administration, continued for 5 days after the last antibiotic dose, and patients were followed for an additional 21 days.

45

Results: Pro-2 (15.5%) had a lower AAD incidence vs. Pro-1 (28.2%). Each probiotic group had a lower AAD incidence vs. placebo (44.1%). In patients who acquired AAD, Pro-2 (2.8 days) and Pro-1 (4.1 days) had shorter symptom duration vs. placebo (6.4 days). Similarly, Pro-2 (1.2%) had a lower CDAD incidence vs. Pro-1 (9.4%). Each treatment group had a lower CDAD incidence vs. placebo (23.8%). Gastrointestinal symptoms were less common in the treatment groups vs. placebo and in Pro-2 vs. Pro-1.

Conclusions: The proprietary probiotic blend used in this study was well tolerated and effective for reducing risk of AAD and, in particular, CDAD in hospitalized patients on antibiotics. A dose-ranging effect was shown with 100 billion c.f.u., yielding superior outcomes and fewer gastrointestinal events compared to 50 billion c.f.u. (ClinicalTrials.gov number NCT00958308).

Notable Level I evidence and Health Canada Recommendations

- ► Constipation BioGaia Protectis (baby and child products)
- Bacterial Vaginosis and Vulvovaginal Candidiasis Provacare vaginal ovule, Ultraflora Women's oral caps, RepHresh ProB probiotic oral caps
- ▶ Colic Purica Probiotic Baby Colic drops, HMF Baby drops, BioGaia Protectis drops and chewable tablets.

47

Efficacy of *Bifidobacterium animalis* subsp. *lactis* BL-99 in the treatment of functional dyspepsia: a randomized placebo-controlled clinical trial

Qi Zhang, Guang Li, Wen Zhao, Xifan Wang, Jingjing He, Limian Zhou, Xiaoxu Zhang, Peng An, Yinghua Liu, Chengying Zhang, Yong Zhang, Simin Liu, Liang Zhao, Rong Liu, Yixuan Li, Wenjian Jiang, Xiaoyu Wang, Qingyu Wang, Bing Fang, Yuyang Zhao, Yimei Ren, Xiaokang Niu, Dongjie Li, Shaoqi Shi, ... Fazheng Ren ✓ + Show authors

Nature Communications 15, Article number: 227 (2024) | Cite this article

432 Accesses | 3 Altmetric | Metrics

Abstract

Current treatment for functional dyspepsia (FD) has limited and unsustainable efficacy. Probiotics have the sustainable potential to alleviate FD. This randomized controlled clinical trial (Chinese Clinical Trial Registry, ChiCTR2000041430) assigned 200 FD patients to receive placebo, positive-drug (rabeprazole), or *Bifidobacterium animalis* subsp. *lactis* BL-99 (BL-99; low, high doses) for 8-week. The primary outcome was the clinical response rate (CRR) of FD score after 8-week treatment. The secondary outcomes were CRR of FD score at other periods, and PDS, EPS, serum indicators, fecal microbiota and metabolites. The CRR in FD score for the BL-99_high group [45 (90.0%)] was significantly higher than that for placebo [29 (58.0%), p = 0.001], BL-99_low [37 (74.0%), p = 0.044] and positive_control [35 (70.0%), p = 0.001].

▶ Daily Bifidobacterium animalis lactis BL-99 supplementation for 8 weeks at a dose of 50 billion CFU per day improved the functional dyspepsia score clinical response rate in 200 trial participants. Randomized Controlled trial.

49

Probiotic for pathogen-specific Staphylococcus aureus decolonisation in Thailand: a phase 2, double-blind, randomised, placebo-controlled trial





Pipat Piewngam*, Sunisa Khongthong*, Natthrit Roekngam, Yongyuth Theapparat, Somkiat Sunpaweravong, Damrongsak Faroongsarng, Michael Otto

Summary

Background Decolonisation is considered a valuable means to reduce Staphylococcus aureus infection rates. However, Lancet Microbe 2023; 4: e75-83 previous topical strategies targeting the nose or skin had little success, and oral antibiotic-based decolonisation is ill advised because of eradication of the microbiota and development of antibiotic resistance. We previously showed that the probiotic Bacillus subtilis significantly diminished S aureus at the main intestinal colonisation site via specific bacterial interaction in mice; in this study, we tested this probiotic approach to control S aureus colonisation in

Methods We did a single-centre, phase 2, double-blind, randomised, placebo-controlled trial in adults from the Songkhla region of Thailand who were colonised by S aureus. Eligible participants were adults (aged ≥18 years) without history of intestinal disease, antibiotic treatment, or hospital admission within the previous 90 days. Participants were excluded if they were pregnant, breastfeeding, taking probiotics, or had diarrhoea. Participants were allocated (1:1) to groups by computer randomisation in blocks of four, and research coordinators were masked to group allocation. Participants received 250 mg of probiotic B subtilis MB40 or placebo once per day for 30 days and S aureus colonisation was determined after the last dose was received. The primary outcome was colonisation by S aureus (continuous, mean decrease in colony-forming-unit count) in the intestine (by faecal counts) and nares (by

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51

Findings The trial was done between Jan 29 and June 30, 2021, with enrolment taking place from Jan 29 to April 6, 2021. 115 participants were colonised by *S aureus*, either in the intestine (n=84), nose (n=50), or both (n=19), and were randomly assigned to treatment (n=55) and placebo groups (n=60). Oral probiotic *B subtilis* resulted in significant reduction of *S aureus* in stool (96 · 8%; p<0 · 0001) and nose (65 · 4%; p=0 · 0002). There were no differences in adverse effects or significant microbiome changes between the intervention and placebo groups.

Interpretation *B subtilis* probiotic eliminated more than 95% of the total *S aureus* colonising the human body without altering the microbiota. This probiotic strategy offers several key advantages over presently used decolonisation strategies for potential use in people with chronic or long-term risk of *S aureus* infection. Furthermore, by establishing a defining role of the intestinal colonisation site, our findings call for revisiting fundamental notions about *S aureus* colonisation.

When is the best time to take a probiotic?

- ▶ Information online is very contradictory
- ▶ It depends on the probiotic
- ▶ Stomach acid kills probiotics
- ▶ Enteric coated or encapsulated products take anytime
- ▶ Rapid passage through the stomach helps (i.e. not with food)
- ▶ Lactobacillus more viable in gastric acid
- ▶ Most Bifidobacteria are not acid stable (except lactis and animalis)
- ▶ Saccharomyces boulardii may be stable with or without a meal.
- ▶ Overall take as directed on label or ½ hour before a meal otherwise.
- ▶ Separate dose from antibiotic dose

53

Prebiotics

- ▶ Sometimes used instead of probiotics but with less evidence base
- ▶ Inulin, fructo-oligosaccharides (FOS) and galacto-oligosacharrides (gos)
- ▶ In return for this food, bacteria in the gut make acetate, propionate and butyrate
- ▶ This leads to a lower pH and increased absorption of Ca and Mg.
- ▶ SCFA like butyrate have proposed favorable effects on lipid metabolism and restrained lipogenesis and appetite suppression.
- Not all fiber is prebiotic. Bananas, pomegranate, apples, nectarines, asparagus, oinions, legumes, garlic, snow peas and green beans are prebiotic

Fermented food

- ▶ Created by microbial growth in a controlled fashion
- ▶ Fermented food with probiotics: some yogurt and some kifir.
- ▶ Fermented food with live microbes: uncooked sauerkraut, kimchi
- ▶ Fermented foods without any microbes: wine, beer, sourdough bread, tempeh and chocolate.

55

Recap of Learning Objectives

- ▶ Learn the basics of probiotic nomenclature and terms
- ▶ Become familiar with using AEProbio.com in making probiotic recommendations
- ▶ Challenges with Probiotic recommendations
- ▶ Understand brand specific recommendations for various conditions and the evidence behind them
- ▶ Learn the best way to take probiotics and when not to
- Understand basic information on what prebiotics are and what they do for gut health

