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Diet and Intestin	nal Bacterial Overgrowt	th: is there evidence	?	
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Abstract

The intestinal microbiota and its role in health and disease processes have been the subject of several studies. It is known that changes in the intestinal microbiota occur due to several factors, such as the use of medication, age, lifestyle and diseases, which can modify intestinal homeostasis and lead to excessive growth of bacteria in the small intestine, triggering a clinical condition called small bowel bacterial overgrowth (SIBO). Individuals with SIBO may present gastrointestinal symptoms ranging from nausea, diarrhea and/or constipation, and flatulence to distension and abdominal pain, resulting from poor absorption of nutrients or changes in intestinal permeability. The gold-standard treatment is based on the use of antibiotics to eradicate bacterial overgrowth. Some studies have evaluated diets in the treatment of SIBO; however, the studies are of low methodological quality, making the extrapolation of the results to clinical practice unfeasible. Thus, there is still not enough scientific evidence to support a specific type of diet for the treatment of SIBO.

Key Words: Small intestinal bacterial overgrowth; treatment; diet; nutrition

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Core Tip: Some dietary interventions have been used in clinical practice for the treatment of SIBO, however the available evidence to support such approaches is of low quality and scarce. Thus, the treatment of SIBO must be focused on controlling the underlying disease and observing the improvement in the symptoms presented by the patient, since so far there is no specific diet for the treatment of SIBO

INTRODUCTION

Small intestinal bacterial overgrowth (SIBO) is defined as a clinical condition caused by excessive numbers of small intestinal bacteria (≥10³ CFU/mL) that include

predominantly gram-negative aerobic and anaerobic species^[1]. In the physiological state, there are mechanisms to prevent excessive colonization of bacteria in the small intestine, such as an acidic stomach pH, pancreatic enzymes, the intestinal immune system, small intestine peristalsis, the ileocecal valve and the intestinal barrier itself. However, when changes in any of these mechanisms occur, SIBO can develop^[2].

Given the growing knowledge of the intestinal microbiota and its role in health and disease processes, a series of studies have linked SIBO with diseases such as irritable bowel syndrome (IBS)^[3], inflammatory bowel disease (IBD)^[4], nonalcoholic fatty liver disease^[5], postgastrectomy syndrome^[6] and several other conditions that are considered risk factors for the development of SIBO.

The main gastrointestinal symptoms are nausea, diarrhea and/or constipation, flatulence, distension and abdominal pain. Such symptoms may be due to poor absorption of nutrients or changes in intestinal permeability, as well as the inflammation and/or immune activation that result from pathological bacterial fermentation in the small intestine^[7].

The noninvasive diagnostic method most commonly used in clinical practice is the measurement of hydrogen gas and/or methane exhaled in the breath after the ingestion of a fixed amount of a carbohydrate substrate. The indication of colonization by methanogens, which are not bacteria but belong to the Archaea domain, led to the proposal of a new term called intestinal overgrowth of methanogens (IOM). Given the diagnostic limitations, data on the prevalence of SIBO in the general population are unreliable, and the association of SIBO with the main risk factors is supported^[1].

In terms of treatment, the use of antibiotics has currently been the gold standard for eradicating bacterial overgrowth^[1]. Rifaximin has been shown to be effective in the treatment of SIBO, despite the heterogeneity found in the studies as well as the lack of a recommendation regarding the dose and duration of treatment^[8]. The antibiotics used are based on whether SIBO is associated with methane production (M-SIBO) or hydrogen production (H-SIBO). M-SIBO is associated with delayed small bowel and colon motility and is produced under anaerobic conditions. The treatment of M-SIBO

tends to be easier than that of H-SIBO and is often associated with constipation. H-SIBO is associated with an overgrowth of gram-positive bacteria in the small intestine and usually manifests with diarrheal symptoms^[9]. However, the recurrence rate after treatment with antibiotics is high, showing that there are no universally accepted treatment approaches for the treatment of SIBO^[1].

Regarding diet, studies have focused on diets with a reduction in fermentable products, involving an approach with few fibers and no gluten. The use of probiotics and phytotherapeutics has also been studied, but the results emphasize the need for well-conducted studies so that the results can be extrapolated to clinical practice^[1,9].

DIET AND SIBO: WHAT IS THE EVIDENCE

The principle for treating SIBO involves treating the underlying disease or condition to eradicate the bacterial overgrowth and possible associated nutritional deficiencies^[10].

A recent systematic review included studies that evaluated the use of probiotics, an elemental diet, and herbal therapy for the treatment of SIBO^[11]. It was observed that in the randomized clinical trials that evaluated the use of probiotics, there was heterogeneity between the strains used and the duration of treatment, and regarding the methodological quality, most studies were of moderate quality. These results corroborate the position of the American College of Gastroenterology that treating SIBO with probiotics seems to be counterintuitive and that the studies published thus far are of low quality^[1].

Although it may seem effective, an elemental diet is difficult to follow, and the literature lacks further studies for the clinical applicability of these diets. Regarding the use of medicinal herbs, although a few studies showed promising results in relation to the improvement of symptoms, these studies were of low methodological quality, and further studies are needed to standardize and prove the safety for the clinical applicability of these herbs^[11].

Dietary strategies for the treatment of SIBO are based on a reduction in the consumption of fermentable products, which involves a diet low in fiber, sugar alcohols

and other fermentable sweeteners such as sucralose. However, the dietary data for SIBO are an extension of nutritional therapy for irritable bowel syndrome (IBS)^[3], highlighting diets low in FODMAPs (fermentable oligosaccharides, disaccharides, monosaccharides and polyols), which are short-chain carbohydrates that are osmotically active and fermentable by small intestinal bacteria^[1].

However, it is not clear whether the clinical improvement resulting from dietary changes is the result of a modification of the intestinal microbiota or is simply due to a reduction in fermentation and gas production. Although dietary modification is used in clinical practice for patients with SIBO, its role and effectiveness have not yet been determined^[10].

It is important to highlight that the type and amount of bacteria found in the gastrointestinal tract depend on an individual's diet^[12]. Thus, changes in the intestine lead to a significant impairment of nutrient absorption, including that of macronutrients and some vitamins^[10]. In SIBO, some patients may show some signs of lactose intolerance, but it is believed to be bacterial lactose intolerance and not a deficiency of the lactase enzyme^[13,14].

Other types of diets, such as a carbohydrate elimination diet and the "specific SIBO" diet, which are commonly disseminated on websites, do not have scientific evidence to support such a prescription.

Obese individuals seem to have an increased risk for SIBO^[15], with an estimated prevalence of 41%^[16] and with a predominance of bacteria from the phylum Firmicutes to the detriment of Bacteroidetes^[17].

A recent meta-analysis in which most articles included were from Western countries, with only one Asian study, assessed the association between SIBO and obesity. It was observed that the risk for SIBO was three times higher among individuals with obesity than among individuals without obesity when evaluated only in the Western population [18]. This result highlights the importance that diet plays in the intestinal microbiota, since a Western diet rich in sugars and fats has been shown to reduce beneficial intestinal microorganisms in obese individuals [19].

However, the causal relationship between SIBO and obesity is not clear, and more comprehensive studies are needed that can consider ethnicity, eating habits and potential confounding factors [18].

Another meta-analysis with cohort studies evaluated the relationship of a usual diet, intestinal microbiota and intestinal inflammation, comparing patients with intestinal diseases with the general population. It was observed that a diet rich in plant foods, whole foods and sources of polyunsaturated fatty acids and polyphenols has the potential to prevent intestinal inflammatory processes, and these recommendations are plausible to increase the abundance of the intestinal microbiota^[20].

Thus, despite studies showing apparently promising results, the quality of evidence of these studies is low, and these types of restrictive diets should not be maintained for long, as they are not nutritionally complete and are poor in important substrates for colonic bacteria, which can impact the goal of healthy colonization^[1].

The recommended treatment for SIBO remains empirical, and consideration should be given to identifying and correcting underlying causes, correcting nutritional deficiencies and administering antibiotics^[21].

CONCLUSION

Although diets for the treatment of SIBO seem promising, studies are still scarce and of low quality, limiting their clinical applicability. Therefore, it is important to be careful when prescribing restrictive diets, since the most severe cases of SIBO can cause malabsorption syndrome, which can lead to nutritional risk, and a healthy diet rich in vegetables and whole foods seems the best option to obtain a balanced microbiota. Thus, further studies with well-designed methodological designs are needed to support the clinical applicability of diets as treatments for SIBO.

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