

#### Correspondence



# An analysis of online messages about probiotics Margaret A Brinich<sup>1</sup>, Mary Beth Mercer<sup>1</sup> and Richard R Sharp<sup>12\*</sup>

- \* Corresponding author: Richard R Sharp sharpr3@ccf.org
- <sup>1</sup> Department of Bloethics, Cleveland Clinic, 9500 Euclid, Ave., JJ60, Cleveland, OH, 44195, USA
- <sup>2</sup> Center for Genetic Research Ethics & Law, Case Western Reserve University, Cleveland, OH, 44106, USA

For all author emails, please log on.

BMC Gastroenterology 2013, 13:5

doi:10.1186/1471-230X-13-5

The electronic version of this article is the complete one and can be found online at: <a href="http://www.biomedcentral.com/1471-230X/13/5">http://www.biomedcentral.com/1471-230X/13/5</a>

Received: 15 June 2012 Accepted: 28 December 2012 Published: 11 January 2013

© 2013 Brinich et al.; Ilcensee BioMed Central Ltd.

This is an Open Access article distributed under the terms of the Creative Commons Attribution License (<a href="http://creativecommons.org/licenses/by/2.0">http://creativecommons.org/licenses/by/2.0</a>), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

#### **Abstract**

Formula display: Vilatin Jax

Internet websites are a resource for patients seeking information about probiotics. We examined a sample of 71 websites presenting probiotic information. We found that descriptions of benefits far outnumbered descriptions of risks and commercial websites presented significantly fewer risks than noncommercial websites. The bias towards the presentation of therapeutic benefits in online content suggests that patients are likely interested in using probiotics and may have unrealistic expectations for therapeutic benefit. Gastroenterologists may find it useful to initiate conversations about probiotics within the context of a comprehensive health management plan and should seek to establish realistic therapeutic expectations with their patients.

Keywords: Probiotics; Doctor-patient communication; Complementary and alternative medicine; Ethics

#### **Findings**

A recent analysis by the Pew Internet & American Life Project found that 80% of Internet users in the U.S. search the Internet to obtain health-related information [1]. Those who are living with chronic disease report going online to search for health related information at a slightly higher rate (83%) [1]. These trends reflect the increasing popularity of the Internet as a key source of information about health and highlight the need for physicians to be

familiar with the content of health-related websites their patients may be accessing [2-5].

Like many patients with chronic illness, individuals with gastrointestinal (GI) diseases often turn to the Internet for information on treatment options [ $\underline{6}$ ]. Of particular interest to many patients living with inflammatory bowel disease (IBD), irritable bowel syndrome (IBS), and other chronic GI diseases is the potential utility of probiotics and other so-called complementary and alternative medicines [ $\underline{7}$ ]. Patients can purchase probiotics directly and the marketing of these products is not regulated by the Food and Drug Administration (FDA) in a manner that is comparable to the regulation of pharmaceutical drugs. These considerations highlight the need to examine Internet messages about probiotics. Knowing what information is communicated to patients, and by whom, is critical for gastroenterologists and other clinicians who care for patients with chronic GI diseases, many of whom are using probiotics or may be considering their use as a supplement to ongoing care regimens [ $\underline{7}$ - $\underline{10}$ ].

We present a systematic description of Internet websites discussing probiotics. Our analysis describes: 1- Internet depictions of the potential benefits and risks of probiotics; 2- common misrepresentations of probiotics; and, 3- other information that may influence patient decisions about the use of probiotics. To the extent that these Internet messages may shape patient attitudes and beliefs, gastroenterologists should be aware of how the potential benefits and risks of probiotics are being presented to the patients they serve. By understanding the full spectrum of Internet messages concerning probiotics, gastroenterologists can better engage their patients in discussions about probiotics and their potential use as a tool for managing chronic GI diseases.

We constructed a sample of problotic-related websites using criteria that would mirror the search patterns of patients with GI diseases. Online searches were completed initially using the five most popular search engines at the time of sampling—Google, Yahoo, LiveSearch, AOL, and Ask [11]. Due to the similar search algorithms employed by these five search engines, keyword searches using the term "probiotics" returned similar results. Therefore, we chose to rely on two popular search engines, Google and Yahoo, in identifying relevant Internet websites. We conducted Internet queries using these search engines from May 6 to 26, 2009. Based on the work of Eysenbach and colleagues, which suggests that consumers' online health information searches are typically limited to initial search results, we limited our sample to the first 50 results of each search engine query (5 pages of results per query) [12]. Websites were excluded from the sample if the website: 1- existed solely as a retail site intended for direct purchase; 2- was not able to be accessed at the time of sample selection; 3- was determined to be an erroneous result upon closer examination (e.g. a clearly unrelated website with no content related to probiotics); 4- focused on non-human applications of probiotics; or 5- was a peer reviewed scientific, medical, or research article or book (Figure 1).

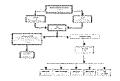


Figure 1. Description of Sampling Methods.

Due to the highly dynamic nature of online content, discrete images of each website were saved as PDF files along with the date on which the content was captured. Two analysts independently reviewed each of these files using a code sheet to document specific content areas and qualities of the websites. A code sheet was developed and used to evaluate major content areas (Table 1). The Sandvik scale was used to evaluate overall website quality based on seven characteristics measured on a scale from 0-worst to 2-best [13]. These results were then summed for a best possible score of 14 [13].

#### Table 1. Coding categories used to analyze the content of 71 Internet websites on probiotics

After two analysts independently coded websites for content, their respective code sheets were compared and used to establish a single consensus code sheet. Cohen's kappa statistic was used to evaluate agreement between the two analysts and showed very high levels of agreement across the 10153 items (143 items per website) evaluated [14]. Kappa values for each of the 71 websites ranged from 0.41 to 0.94, with a mean value of 0.72 and standard deviation of 0.10. Results were entered into an Access database and imported into SPSS version 19.0 for further

analysis [15]. Descriptive statistics were examined to check for outliers and data-entry errors. Independent samples t-tests (for ordinal data) and Fisher's exact tests (for nominal data) were used to assess differences between commercial and noncommercial websites. P values of  $\leq 0.05$  were considered statistically significant.

A total of 71 websites were identified and analyzed for content. As shown in Table 2, these websites were diverse with respect to content and intended audience. Approximately half of the websites (51%) were categorized as commercial. Websites that were not commercial in nature included those run by non-profit organizations (13%), news media (13%), Internet content farms<sup>a</sup> (4%), and personal websites (4%). Two websites included in the sample were maintained by medical institutions, one from the Mayo Clinic and another from the University of Alabama Health System. The only government site included in the analysis was supported by the National Center for Complementary and Alternative Medicine.

#### Table 2. Characteristics of 71 Internet websites on probiotics

A majority of these websites (76%) identified at least one problotic product using a commercial brand name. It was also common for websites to have some connection or "linkage" to more information about a specific problotic product (72%) (Table 2). Many of those links were to product advertisements (75%). Other links directed readers to outside websites where problotic products could be purchased (47%).

By utilizing a standard metric to examine website quality, the Sandvik score set a mean overall quality score at 9.3 (Table 3). When commercial and noncommercial websites were compared, their Sandvik scores differed significantly with mean scores of 7.7 and 10.7 respectively (p=.001). Commercial websites received consistently lower mean ratings than their noncommercial counterparts in six of the seven content areas examined using the Sandvik scale, including "balance," "authorship," and "currency" of content, with significantly lower mean scores across each of these three areas (p=.001).

## <u>Table 3.</u> Comparison of general quality criteria scores (Sandvik Scores) between 71 commercial and noncommercial Internet websites

Websites were generally accurate in their depictions of what probiotics are and how they function in the host (Table 4). Probiotics typically were described as "complementary" to other approaches to promoting good health. Although a widely accepted definition of probiotics recommended by the Food and Agricultural Organization of the United Nations and World Health Organization (FAO-WHO) was not used consistently across websites, over half of the websites dld contain all four components of this definition (56%). Two primary elements of this definition, that probiotics are "live microorganisms" (89%) ingested to provide "health benefit" (86%), were present in most websites. Websites often suggested that probiotics may provide benefits related to general health and wellness (70%) as well as improvements in GI health (96%).

#### <u>Table 4.</u> Characterizations of probiotics on 71 Internet websites

The description of probiotics as consumable products almost uniformly included some mention of the species (94%) and manner in which it is to be ingested (99%). In addition to defining what probiotics are and suggesting appropriate indications for their use, all but one website noted at least one mechanism of action by which probiotic bacteria function in the host. The two most common mechanisms described were to regulate immune response (83%) and inhibit pathogenic bacteria colonization (82%).

#### Focus on therapeutic benefit

The representation of potential benefits and risks varied greatly across websites (Table  $\underline{5}$ ). Seventy percent of websites identified at least one specific disease or clinical indication for problotics. Specific benefits cited ranged from relieving general gastrointestinal distress such as gas and bloating (n=63), to managing behavioral symptoms related to autism (n=6) and preventing cardiovascular disease (n=4). Specific risks associated with problotics were noted less frequently than benefits and typically focused on minor GI issues such as the potential for diarrhea or gas

(n=28). A few websites also discussed less common risks related to bacterial Infection (n=7) and headaches (n=4). Other potential safety issues focused on heightened risks for specific populations, such as probiotic use by children (n=20) and pregnant women (n=13). Of the websites examined, 47% did not mention any risks associated with probiotic use.

#### Table 5. Specific benefits and risks of probiotics described on 71 Internet websites\*

On average, websites mentioned 10.2 benefits and 1.8 risks per website, with an upper range of 36 benefits and 11 risks on any individual site (Table  $\underline{6}$ ). The number of risks described on commercial websites was significantly lower than noncommercial websites (1.0 and 3.0 respectively; p=.002). Although no formal evaluation was performed, websites often reiterated one or more benefits of using problotics multiple times throughout the text. Conversely, the discussion of risks associated with problotics was less commonly reiterated in multiple locations.

#### Table 6. A comparison of how commercial and noncommercial websites portray probiotics

Statements regarding the general safety of probiotics appeared in approximately half (49%) of the websites reviewed (Table 6). Broad statements of general health benefit—such as claims that probiotics can help "Maintain optimal health and wellness"—were present in more than half of the websites (59%). Commercial websites were significantly more likely to include such a statement (62%) in comparison to noncommercial websites (32%) (p=.031).

Nearly half (47%) of the websites encouraged readers to discuss problotics with a heath care provider and 56% explicitly endorsed the use of problotics (Table  $\underline{6}$ ). These endorsements were often made by an identifiable individual, such as a named physician (23%), other medical professional (31%) or patient/consumer (21%). Commercial websites were more likely to recommend the use of problotics than noncommercial websites (p=.001).

When gastroenterologists and other healthcare providers (HCPs) encounter patients who are interested in using probiotics as therapy for their GI disease, they should anticipate that many of these patients will have been exposed to a variety of messages concerning the health benefits of probiotics. Our data indicate that many Internet websites contain unsubstantiated and misleading claims about the therapeutic benefits of probiotics. This benefit-focused messaging is particularly troubling given that probiotics have not been studied extensively in clinical trials and there is little to no evidence supporting many of the benefits claimed on some websites. For example, although there is evidence suggesting that probiotic bacteria may have some utility in the care of patients with IBS, ulcerative colitis, and pouchitis [16-21], claims that probiotics can prevent cancer or assist in managing autistic behaviors lack empirical support.

In addition to unsupported claims of benefit, our analysis also revealed a comparative lack of information about the risks of using probiotics. Studies suggest that the level of risk posed to healthy individuals by a typical probiotic is very low [22]. In the absence of high quality data documenting harms associated with typical uses of probiotics, the lack of online information about possible side effects is unsurprising. For some individuals, however, such as immunocompromised patients who may be seeking probiotics to manage a chronic GI disease, there may be additional risks that are not discussed on most Internet websites [23-27].

#### Effective patient communication amid mixed health messages

Alongside the specific health benefits noted above, our analysis identified many broad statements promoting problotics as a general wellness tool. The promotion of probiotics as a holistic "complementary and alternative medicine" (CAM) situates these products in a market space occupied primarily by healthy consumers seeking to prevent disease and maintain overall health and wellness. Unfortunately, a number of studies have demonstrated that patients often are reluctant to discuss their interest in CAM with physicians [28-32]. In the case of probiotics, wellness messaging on Internet websites may make patients less likely to consult with a HCP prior to using probiotics. By employing a combination of marketing messages, focusing on both specific symptom management and promotion of overall health, probiotic products are depicted as both benign CAM and as powerful disease management tools, even within the very same website. This dual messaging may create confusion among patients with chronic GI diseases, who may wonder where these products fit within the context of their current therapies. Of

particular concern is that these patients may discontinue medication use without consulting a gastroenterologist, in favor of over-the-counter probiotic products.

The difference in language used to market probiotics to healthy consumers contrasts with marketing strategies designed to elicit interest among patients with chronic GI diseases. In a study by Mercer and colleagues, when GI patients were asked to comment on their personal perceptions of the risks and benefits of probiotics, the authors found that many GI patients were reluctant to initiate new treatments, even over-the-counter treatments, without consulting their physicians [2]. Mercer and colleagues also found that patients expressed concerns that while probiotics may be effective in promoting good digestive health for many individuals, as patients with chronic GI diseases, they felt they needed to exercise greater caution in using products that affect digestive balance. For many of these patients, claims of improved digestive health were not sufficient to motivate probiotic use.

Another source of inconsistent online messaging about probiotics is evident in the frequent suggestion that individuals consult a physician or other medical professional prior to using probiotics. Although the importance of seeking advice from a physician was explicit on many websites this message frequently was accompanied by tips on where to purchase over-the-counter problotics or invitations to purchase probiotics online. While suggestions to bring physicians into decisions about the use of probiotics may temper patient's enthusiasm to purchase probiotics independently, appeals to physician involvement may also suggest greater levels of medical benefit than are supported by available data.

We suggest that the responsibility to clarify the potential benefits and risks of probiotic therapies falls squarely in the hands of health care providers. By discussing probiotics with patients in a nonjudgmental manner that is mindful of the many mixed messages that patients may have received concerning these products, HCPs can help patients assess what role, if any, probiotics should play in their ongoing treatment approach.

#### Commercial interests

The complex and sometimes inconsistent mix of messages about probiotics presented online are further complicated by a strong presence of commercial interests in many of the websites patients may view. Structurally these sites interface closely with probiotic marketing and frequently identify specific brands or products by name. Even when patients are not offered the opportunity to purchase a particular product online, our data shows that online searches will often include multiple websites promoting the use of probiotic products. The Influence of commercial interests is of particular concern since the overall quality of commercial websites, as measured by the Sandvik score, was significantly lower in comparison to websites that lacked a commercial focus. Many commercial websites failed to identify an author, failed to indicate how current the information contained on the site was, and lacked balance. These findings suggest that HCPs should be cautious in promoting any particular product and direct interested patients to more balanced, higher quality websites maintained by noncommercial entities.

In an era when direct-to-consumer advertising plays a major role in the marketing of pharmaceutical drugs [33,34], the strategies employed on probiotic websites are not surprising. In addition, manufacturers' claims regarding the efficacy of probiotic products are not subjected to the rigorous oversight typical of pharmaceutical drugs [35]. Nonetheless, references to peer-reviewed medical literature appeared in the majority of the websites we reviewed (63%). The presence of such citations implies a certain degree of scientific validity behind claims of safety or efficacy, regardless of the accuracy of the specific statements made or relevance of the studies cited. In a best case scenario, potential inconsistencies between available research data and claims of health benefit would be discussed in a clinical setting, with the involvement of a knowledgeable gastroenterology specialist. However, it is unclear whether patients will seek this additional input from a HCP or pursue probiotics outside clinical settings, as a self-management approach.

#### Study limitations

This study of online content related to probiotics has several limitations that reflect our approach to defining an appropriate sample. First and foremost is the challenge of characterizing the ever-changing information available on the Internet. By taking snapshots of relevant websites within a defined time period we were able to minimize the amount of change that may have occurred during data collection and analysis. Subsequent searches of Internet content returned a similar set of websites, suggesting that this information may be reasonably stable. We did not

assess the stability of this information formally, however, which is a ilmitation of our approach. In addition, our analysis of website content, which employed well established qualitative methods and duplicate analysis by two independent coders, has the potential to introduce human error or evaluator bias. Finally, our approach focused narrowly on website content and did not consider how individuals who are exposed to this information may understand or use the information presented. Although not within the scope of our study, our findings highlight the need for additional research examining how patients interpret online information about probiotics and make healthcare choices based on that information.

As ever larger volumes of information are available directly to patients via the Internet, one of the greatest challenges facing HCPs is to stay current with the many health-related messages that patients may encounter outside traditional clinical settings. Although this challenge is not unique to gastroenterology, the marketing and direct availability of probiotic products combine to create additional challenges in maintaining good communication between patients and their HCPs.

Our results suggest that the combination of directive product branding, unsubstantiated claims of health benefit on Internet websites, and direct patient access to problotics, should be cause for concern among gastroenterologists. The messages patients receive from Internet websites may require that gastroenterologists and other HCPs revisit their patients' expectations about probiotics and provide a more scientifically grounded and balanced overview of their therapeutic value. These discussions should aim to clarify the potential benefits and risks of probiotics for individuals with chronic GI diseases, highlighting potential differences in the benefit-to-risk profiles of probiotic usage for healthy consumers in comparison to patients with chronic GI diseases. Gastroenterologists may also find it helpful to direct interested patients to well established sources of health information that are free of commercial influence. By initiating these discussions, gastroenterologists and other patient educators can help to establish realistic expectations about probiotics.

#### Endnotes

<sup>a</sup> According to MacMillan Dictionary a content farm is, "A website which publishes huge volumes of low-quality content."

#### Competing interests

The authors declare that they have no competing interests.

#### **Authors' contributions**

MB contributed to study design, data collection, data analysis and the writing of the manuscript. MBM contributed to study design, data collection, data analysis, and the writing of the manuscript. RS conceived of the study, participated in its design and coordination, and provided critical review and revision of the manuscript.

#### Acknowledgements

This work was funded by grant number R01HG004877 from the National Human Genome Research Institute of the National Institutes of Health. We thank Sabahat Hizlan for her contributions to data analysis.

#### References

Pew Internet and American Life Project:
 Health topics February 1, 2011.
 http://www.pewInternet.org/~/media//Files/Reports/2011/PIP\_HealthTopics.pdf | webcite |.

#### FULL Lane Library TEXT @ STANFORD

2. Boucher J: Technology and patient-provider interactions: improving quality of care, but is it improving communication and collaboration?

Diabetes Spectrum 2010, 23:142-144. Publisher Full Text TEXT



3. Cowan M: Millennial transformation for primary care.

Military Med 2010, 175:379-381.



4. Murray E, Lo B, Pollack L: The impact of health information on the Internet on the physician-patient relationship.

Arch Intern Med 2003, 163:1727-1734. PubMed Abstract | Publisher Full Text



5. Hesse B, Nelson D, Kreps, Croyle RT, Arora NK, Rimer BK, Viswanath K: **Trust and sources of health** information: the impact of the Internet and its implications for health care providers: findings from the first health information national trends survey.

Arch Intern Med 2005, 165:2618-2624. PubMed Abstract | Publisher Full Text



6. O'Connor JB, Johanson JF: Use of the web for medical information by a gastroenterology clinic population.

JAMA 2000, 284:1962-1964. PubMed Abstract | Publisher Full Text



7. Mercer M, Brinich MA, Geller G, Harrison K, Highland J, James K, Marshall P, McCormick JB, Tilburt J, Achkar JP, Farrell RM, Sharp RR: How patients view probiotics: findings from a multicenter study of patients with inflammatory bowel disease and irritable bowel syndrome.

J Clin Gastroenterol 2012, 46:138-144. PubMed Abstract | Publisher Full Text



8. Langhorst J, Anthonisen IB, Steder-Neukamm U, Luedtke R, Spahn G: Patterns of complementary and alternative medicine (CAM) use in patients with inflammatory bowel disease: perceived stress is a potential indicator for CAM use.

Complement Ther Med 2007, 15:30-37. PubMed Abstract | Publisher Full Text



9. Li FX, Verhoef MJ, Best A, Otley A, Hilsden RJ: Why patients with inflammatory bowel disease use or do not use complementary and alternative medicine: a Canadian national survey.

Can J Gastroenterol 2005, 19:567-573. PubMed Abstract | Publisher Full Text



10. Hilsden RJ, Verhoef MJ, Best A, Pocobelli G: Complementary and alternative medicine use by Canadian patients with inflammatory bowel disease: results from a national survey.

Am J Gastroenterol 2003, 98:1563-1568. PubMed Abstract | Publisher Full Text



Nielsen Wire :

Top U.S. Online search providers: February 11, 2009.

http://blog.nielsen.com/nielsenwire/online\_mobile/top-us-online-search-providers-january-2009/\_webcite\_l.

FULL Lane library TEXT & SYANFORD 12. Eysenbach G, Kohler C: How do consumers search for and appraise health information on the worldwide-web? qualitative study using focus groups, usability tests and in-depth interviews.

BMJ 2002, **324:**573-577. <u>PubMed Abstract | Publisher Full Text | PubMed Central Full Text</u>



13. Sandvik H: Health information and interaction on the Internet,

BMJ 1999, 319:29-32. PubMed Abstract | Publisher Full Text | PubMed Central Full Text



Landis JR, Koch GG: The measurement of observer agreement for categorical data.

Biometrics 1977, 33(1):159-174. PubMed Abstract | Publisher Full Text



15. SPSS for windows [computer program]. version 19.0. SPSS Inc; 2010.



16. Fedorak , Richard N, Dieleman LA, Madsen KL: Prebiotics, probiotics, antibiotics, and nutritional therapies in IBD. In Inflammatory bowel disease, clinical gastroenterology. 2nd edition. Edited by Cohen RD, Wu GU.

Totowa, NJ: Humana Press; 2011::123-150. FULL Land Librar



17. Floch MH, Walker WA, Guandalini S, Hibberd P, Gorbach S, Surawicz C, Sanders ME, Garcia-Tsao G, Quigley E, Isolauri E, Fedorak RN, Dieleman LA: Recommendations for probiotic use-2008.

J Clin Gastroenterol 2008, 42(Suppl 2):104-108.



18. Holubar SD, Cima RR, Sandborn WJ, Pardi DS: Treatment and prevention of pouchitis after ileal pouch-anal an astomosis for chronic ulcerative colitis.

Cochrane Database Syst Rev 2010, 6:CD001176. PubMed Abstract | Publisher Full Text



19. Sood A, Midha V, Makharia GK, Ahuja V, Singal D, Goswami P, Tandon RK: The probiotic preparation, VSL#3 induces remission in patients with mild to-moderately active ulcerative colitis.

Clin Gastroenterol Hepatol 2009, 7:1202-1209, PubMed Abstract | Publisher Full Text



20. Miele E, Pascarella F, Giannetti E, Quaglietta L, Baldassano RN, Staiano A: Effect of a probiotic preparation (VSL#3) on induction and maintenance of remission in children with ulcerative colitis.

Am J Gastroenterol 2009, 104:437-443. PubMed Abstract | Publisher Full Text



21. Mallon P, McKay D, Kirk S, Gardiner K: Probiotics for induction of remission in ulcerative colitis.

Cochrane Database Syst Rev 2007, 4:CD005573. PubMed Abstract | Publisher Full Text



22. Hempel S, Newberry S, Ruelaz A, Wang Z, Miles JNV, Suttorp MJ, Johnsen B, Shanman R, Slusser W, Fu N, Smith A, Roth E, Polak J, Motala A, Perry T, Shekelle PG: Safety of probiotics to reduce risk and prevent or treat disease. Evidence report/technology assessment No. 200. (Prepared by the southern California evidence-based practice center under contract No. 290-2007-10062-I.) AHRQ publication No. 11-E007. Rockville, MD: Agency for Healthcare Research and Quality; 2011.

http://www.ahrq.gov/clinic/tp/probiotictp.htm webcite .

rull lane Histori 9 STANFORD

23. Boyle RJ, Robins-Browne RM, Tang ML: Probiotic use in clinical practice: what are the risks.

Am J Clin Nutr 2006, 83:1256-1264. PubMed Abstract | Publisher Full Text



24. Zuccotti GV, Meneghin F, Raimondi C, Dilillo D, Agostoni C, Riva E, Giovannini M: **Probiotics in clinical** practice: an overview.

J Int Med Res 2008, 36(Suppl 1):1A-53A. PubMed Abstract



- 25. Walker R, Buckley M: *Probiotic microbes: the scientific basis*. Washington, DC: American Academy of Microbiology; 2006. FULL Lane Library C STANEORU
- 26. Besselink MG, van Santvoort HC, Buskens E, Boermeester MA, van Goor H, Timmerman HM, Nieuwenhuijs VB, Bollen TL, van Ramshorst B, Witteman B, Rosman C, Ploeg RJ, Brink MA, Schaapherder A, Dejong C, Wahab PJ, van Laarhoven CJ, van der Harst E, van Eijck C, Cuesta MA, Akkermans L, Gooszen HG: Probiotic prophylaxis in predicted severe acute pancreatitis: a randomized, double-blind, placebo-controlled trial.

  Lancet 2008, 371:651-659. PubMed Abstract | Publisher Full Text | Full Lane Hibrary
- 27. Vogel G: Clinical trials: deaths prompt a review of experimental probiotic therapy.

  Science 2008, 319:557. PubMed Abstract | Publisher Full Text | Full Lang Library | C STANFORD
- 28. Mehta DH, Gardiner PM, Phillips RS, McCarthy EP: Herbal and dietary supplement disclosure to health care providers by individuals with chronic conditions.

  J Altern Complement Med 2008, 14:1263-1269. PubMed Abstract | Publisher Full Text | PubMed Central Full Text | Full Lang Library O STANFORD
- 29. Blendon RJ, DesRoches CM, Benson JM, Brodie M, Altman DE: Americans' views on the use and regulation of dietary supplements.
  Arch Intern Med 2001, 161:805-810. PubMed Abstract | Publisher Full Text
- 30. Eisenberg DM, Kessler RC, Van Rampay MI, Kaptchuk TJ, Wilkey SA, Appel S, Davis RB: Perceptions about complementary therapies relative to conventional therapies among adults who use both: results from a national survey.

Ann Intern Med 2001, 135:344-351. PubMed Abstract | Publisher Full Text Full



31. Adler SR, Fosket JR: Disclosing complementary and alternative medicine use in the medical encounter: a qualitative study in women with breast cancer.

J Fam Pract 1999, 48:453-458. PubMed Abstract



32. Pappas S, Perlman A: Complementary and alternative medicine. The importance of doctor-patient communication.

Med Clin North Am 2002, 86:1-10. PubMed Abstract | Publisher Full Text



33. Llang BA, Mackey T: Direct-to-consumer advertising with interactive internet media: global regulation and public health issues.

JAMA 2011, 305:824-825. PubMed Abstract | Publisher Full Text



34. Frosch DL, Grande D, Tarn DM, Kravitz RL: A decade of controversy: balancing policy with evidence in the regulation of prescription drug advertising.

Am J Public Health 2010, 100:24-32. PubMed Abstract | Publisher Full Text | PubMed Central Full Text

FULL Lane Library TEXT © STANFORD

35. Food and Agriculture Organization of the United Nations and World Health Organization:

Report on joint FAO/WHO expert consultation on evaluation of health and nutritional properties of probiotics in

food including powder milk with live lactic acid bacteria. 2001.

ftp://ftp.fao.org/es/esn/food/probio\_report\_en.pdf webcite.

FULL Lane Library TEXT © STANFORD

### Pre-publication history

The particular of the particul	
The pre-publication history for this paper	er can be accessed here:
http://www.biomedcentral.com/1471-230X/13/5/prepub	
	2 Springer