

DRINKING CURE BEVERAGE COMPRISING THREE COMPONENTS FOR BUILDING UP, PROMOTING AND MAINTAINING INTESTINAL HEALTH

[0001] The invention relates to a special beverage consisting of three components, to be drunk in a drinking cure which should preferably be used daily. To understand the motivation for this special composition of this beverage and the biological and medical background for the selection of the components, and to explain this drinking cure and its effectiveness, some important definitions are given, some of which are referenced in square brackets to scientific literature:

Microbiome: In the narrower sense, this refers to the totality of microbial DNA in the human body. In a broader and colloquial sense, the term “microbiome” refers to the totality of all *microorganisms* in the human body. In humans, microbiomes are differentiated according to body niche (intestine, skin, oral cavity, vagina, etc.). The intestinal microbiome is located in the lumen in the outer mucus layer of the intestine.

Commensals: Describes the organism that lives together with another organism on the same food without harming it.

Eubiosis: Describes the balance of bacterial cultures in the microbiome of a healthy person.

Dysbiosis: Describes a disturbed balance of bacterial cultures in the microbiome, the opposite of eubiosis.

Vitality: Describes the body's own mental, physical, emotional and social performance.

Probiotics: Describes living, non-pathogenic microorganisms that provide health benefits to the host (here, specifically the intestine) [1].

Prebiotics: Describes substrates that are selectively used by host microorganisms (here, specifically the microbiome) and provide health benefits [2].

Synbiotics: Describes a mixture of probiotics and prebiotics with a health benefit for the host (here, specifically for the intestine) [3].

[0002] In the microbiome of the human gut, around 100 trillion (10^{14}) bacteria are present. No other habitat on Earth is more densely populated. In the large intestine, there are $10^{10} - 10^{12}$ bacteria per gram of biomass. The bacteria consist of >90% of the two anaerobic phyla Bacillota (>75%) and Bacteriodota
5 (>15%) and form about 10,000 different commensal species. In the small intestine, the main species are *Lactobacillus* and *Enterococcus*, and in the large intestine the main species are *Bifidobacterium* alongside *E. coli*, *Enterobacterium* and *Clostridium*. This complex bacterial diversity makes humans a superorganism, possessing around 300 times more genes than a
10 human cell and producing a metabolic potential around 100 times higher than the human liver [4].

[0003] The intestinal microbiome is considered an (endocrine) “superorgan” [5] or “second brain” [6]. It is a complex microbial ecosystem in which bacteria live
15 together, communicate, exchange metabolic products and recombine. And this complex bacteria-gut interaction is strongly influenced by nutrition. Depending on the composition of the food and the bacterial composition, a variety of different physiologically active metabolites are formed in the microbiome.

[0004] The prerequisite for healthy vitality is balance (eubiosis) of the bacterial
20 cultures in the microbiome. This requires the microbiome to be supplied with a balanced mix of the macronutrients carbohydrates, proteins and fats [7]. In the microbiome, indigestible nutrients such as complex glucans and glucoconjugates of plant cell walls (colloquially called dietary fiber or fiber),
25 proteins along with glycine, taurine, bile acid and other conjugates, i.e., products resulting from the covalent coupling of at least two molecules, are metabolized. The anaerobic saccharolytic fermentation of the fiber results in “short-chain fatty acids” (SCFA) via pyruvic acid, i.e., acetic acid, propionic acid and butyric acid.

[0005] The change in the bacterial composition of the microbiome due to
30 disturbances significantly affects vitality. Over 90% of all diseases and complaints are caused by dysbiosis. Prominent examples include chronic

intestinal inflammation, obesity, diabetes, acne, atherosclerosis, asthma, neurodegenerative diseases and certain types of cancer [8]. Dysbiosis is caused by inadequate nutrient intake, poor nutrition, the intake of medication (especially antibiotics) and chronic stress.

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[0006] Known metabolites of the intestinal microbiome include serotonin, short-chain fatty acids, various B vitamins (B1, B2, B5, B7, B9, B12), vitamin K, secondary bile acids and branched-chain amino acids. Functionally, these metabolites improve nutrient metabolism and regulate the cardiovascular, immune, and nervous systems via the gut-brain [9], gut-lung [10], gut-heart [11], and gut-skin [12] axes.

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[0007] Carbohydrates, proteins and fats make up about 90% of the dry weight of food and 100% of its energy.

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[0008] Various lactic acid bacteria strains of the genus *Bifidobacterium* also have a detoxifying effect. Lactic acid bacteria from the strain *Enterococcus faecalis* can also have a detoxifying and health-promoting effect. They cause oxalate breakdown and thus prevent the formation of kidney stones [13].

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[0009] Bacteria of the genus *Lactobacillus* are heterofermentative lactic acid bacteria and ferment fiber via the phosphoketolase pathway to acetic acid and lactic acid. They are the most common bacterial genus in the microbiome [14].

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[0010] It is known [15] that the bacteria of the genus *Bifidobacterium* present in the microbiome ferment the greatest variety of fiber. From the fermentation of the fiber, the hexoses are synthesized into acetic acid and lactic acid in a ratio of 3:2. Acetic acid is one of the “short-chain fatty acids” and is involved in the regulation of metabolism along with the immune and nervous systems.

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[0011] Lactic acid is used for energy production (gluconeogenesis) and has numerous vascular effects. The short-chain fatty acids serve as an energy

source for the epithelial cells of the intestine and are important signaling molecules. They bind to the receptors GPR41 and GPR43 on epithelial and immune cells. The receptors regulate the release of pro-inflammatory cytokines, which act directly on the central nervous system via GLP1 (glucagon-like peptide) and PYY (peptide-tyrosine-tyrosine). Furthermore, short-chain fatty acids inhibit histone deacetylase (HDAC) in immune cells and adipocytes and thus affect the transcription of these cells. In this way, the metabolism along with the immune and nervous systems are regulated.

10 [0012] The short-chain fatty acids are distributed throughout human tissue, including the brain, with the help of transporter proteins [16]. There, they regulate the metabolism along with the immune and nervous systems. In addition, the pH value is lowered to 5.5, thus preventing the proliferation of pathogenic intestinal bacteria.

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[0013] Enzymes of intestinal bacteria metabolize the secondary bile acids from the bile acid conjugates. These emulsify the fats from food and thus allow for enzymatic fat breakdown by lipases.

20 [0014] Enzymes of the intestinal bacteria hydrolyze short-chain peptides and individual amino acids from the proteins. These are used to build up and regenerate the intestinal mucosa and are transported via the bloodstream to the cells, where they are used for protein biosynthesis.

25 [0015] The incidence of diseases based on impairment of the cardiovascular, immune and nervous systems has increased in Western civilization. The increase correlates with the declining consumption of fiber. There is a causal relationship between nutrition, the intake of fiber and the metabolic products formed from it through fermentation, such as short-chain fatty acids.

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[0016] Nutrition and the resulting change in the composition of the microbiome is therefore directly linked to Western civilization diseases such as digestive

problems, obesity, attention deficits, concentration and sleep disorders. Prevention or correction through specifically tailored probiotic food supplements is possible.

- 5 [0017] The intake of prebiotics influences the relative growth of bacterial strains and thus changes the bacterial composition of the microbiome and thus the composition of the physiologically active metabolic products.

[0018] Bearing in mind all these scientific findings, the object of the present
10 invention is to provide a beverage for building up, promoting and maintaining intestinal health, which can be consumed daily as part of a drinking cure and is particularly effective and easy to use. The ingredients selected for the beverage are of crucial importance.

15 [0019] To achieve this object, a beverage is provided for building up, promoting and maintaining the intestinal microbiome (eubiosis), consisting of at least the following components:

- a probiotic, with around 30 billion live lactic acid bacteria per 10 mL, and at least 10 natural herbs and spices that have been preserved by fermentation
 - 20 by mixing the herbs and bacteria with water and sugar cane molasses, as a result of which the bacteria consume the sugar in the molasses and convert it into lactic and acetic acid,
 - a prebiotic combined with essential vitamins, minerals and trace elements,
 - a natural nutritional oil which contains more than 50 wt.% of omega 3 fatty
 - 25 acids and tocopherol,
- by processing these components into an emulsion as a beverage and administering them preferably daily as part of a drinking cure.

[0020] Daily intake of this beverage as part of a drinking cure supports build
30 up, promotion and maintenance of the intestinal health. The ingredients of the first component contain around 30 billion live lactic acid bacteria per 10 mL and over 10 different natural herbs and spices. Depending on the need, the

probiotic can also contain bacteria from the strain *Enterococcus faecium* and one or more strains of the genera *Lactobacillus* and *Bifidobacterium*, for example these, which all originate from Denmark:

• *Lactobacillus acidophilus* 43 | CIP 103598, ATCC 314, JCM 2121 |
5 BacDiveID:135307 - <https://bacdive.dsmz.de/strain/135307>

• *Enterococcus faecium* | DSM 2146, ATCC 6057, WDCM 00177, NBRC 113009, CIP 106742, CCM 2308, CECT 8108 | BacDiveID:5295 -
<https://bacdive.dsmz.de/strain/5295>

• *Bifidobacterium bifidum* Ti, Tissier | Type strain | DSM 20456, ATCC 29521,
10 JCM 1255, BCRC 11845, BCRC 14615, CCUG 18364, CCUG 45217, CECT 870, CGMCC 1.2212, CIP 56.7, HAMBI 1380, IFO 14252, KCTC 3202, KCTC 3281, LMG 11041, LMG 8810, NBRC 100015, NBRC 14252, NCAIM B.02021, NCFB 2715, NCIMB 702715, NCTC 13001, VTT E-97795, NCDO 2715 |
BacDiveID:1693 <https://bacdive.dsmz.de/strain/1693>

15 The product is naturally preserved through fermentation, in which the herbs and bacteria are mixed with water and sugar cane molasses. The bacteria consume the sugar in the molasses and convert it into lactic and short-chain fatty acids.

[0021] The ingredients of the second component include >10 plant fibers as
20 prebiotic substrates and >30 different minerals, vitamins and trace elements. The main component with >25 wt.% contains plant fiber consisting of complex glycans, such as glucomannans, arabinogalactans, xyloglucans, galacturonans, fructooligosaccharides or galactomannans. Galactomannans consist of β -1,4 linked mannose as the main chain, which is partially linked to
25 α -1,6 galactose residues. For example, guar gum contains this glycan.

[0022] Clinical studies have shown that dietary supplements enriched with partially hydrolyzed guar gum achieve benefits and clinical effects in the reduction of diarrhea, in the treatment of irritable bowel syndrome and in
30 constipation [17]. In addition, the availability of fat and cholesterol for resorption is significantly reduced and the risk of vascular diseases is reduced [18].

[0023] The prebiotic substrates contained in the second component are metabolized by the saccharolytically fermenting bacteria in the microbiome into lactic acid and the short-chain fatty acids acetic acid, propionic acid and butyric acid.

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[0024] The purpose of the second component is to supply the microbiome with substrates that serve as precursors for the bacterial synthesis of short-chain fatty acids. Substrates that lead to toxic or harmful metabolites through bacterial synthesis are deliberately omitted in the second component.

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[0025] The third component is a natural nutritional oil highly enriched in omega 3, such as linseed oil or Sacha Inchi nut oil. It contains >50% omega 3 fatty acids and up to 50 mg/100 mL of tocopherol. Omega 3 fatty acids promote the growth of butyric acid, one of the short-chain fatty acid producing bacteria in the microbiome. In this way, they prevent neurodegenerative diseases and protect against inflammation, arteriosclerosis and heart attacks [19].

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[0026] Studies show that changes in microbiome composition are causally correlated with the occurrence of so-called lifestyle diseases. Lifestyle diseases are diseases that are related to the way of life. These include previously unknown incidences of metabolic diseases such as obesity and diabetes, inflammatory bowel diseases, asthma, allergies and immune deficiencies and neurodegenerative diseases [20].

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[0027] In these diseases, a significant decrease in bacterial diversity and the loss of specific metabolic functions are observed. The absence of specific bacteria or dysbiosis is the cause of the occurrence of these complex diseases [21].

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[0028] The daily, simultaneous, ritualized intake of the drinking cure based on the presented beverage with its three cardinal components has the purpose of ensuring that the supplied substances match the daily circadian fluctuations in the composition and function of the microbiome [22].

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[0029] The simultaneous intake of the three components in a daily, ritualized drinking cure combines pro- and prebiotic food and nutritional supplements and thus results in a unique synbiotic preparation.

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Claims

1. A beverage for building up, promoting and maintaining the intestinal microbiome (eubiosis), consisting of at least the following components:
 - 5 • a probiotic which contains approximately 30 billion live lactic acid bacteria per 10 mL, and at least 10 natural herbs and spices that have been preserved by fermentation by mixing the herbs and bacteria with water and sugar cane molasses, as a result of which the bacteria consume the sugar in the molasses and convert it into lactic and acetic acid,
 - 10 • a prebiotic combined with essential vitamins, minerals and trace elements,
 - a natural nutritional oil which contains more than 50 wt.% of omega 3 fatty acids and tocopherol,
 - by processing these components into an emulsion as a beverage and
 - 15 administering them preferably daily as part of a drinking cure.
2. The beverage according to claim 1, characterized in that the ingredients of the first component, the probiotic, additionally contain living bacteria from the strain *Enterococcus faecium* and one or more strains of the genera
- 20 *Lactobacillus* and *Bifidobacterium*.
3. The beverage according to any of claims 1 to 2, characterized in that the ingredients of the second component, the prebiotic, comprise more than 10 plant fibers as prebiotic substrates and more than 30 different minerals,
- 25 vitamins and trace elements, the main component containing more than 25 wt.% of vegetable fiber from complex glycans, such as glucomannans, arabinogalactans, xyloglucans, galacturonans, fructooligosaccharides or galactomannans, which consists of β -1,4 linked mannose as the main chain with partially linked α -1,6 galactose residues.
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4. The beverage according to any of claims 1 to 2, characterized in that the ingredients of the second component, the prebiotic, comprise more than 10 plant

fibers as prebiotic substrates and more than 30 different minerals, vitamins and trace elements, the main component containing more than 25 wt.% of fiber from the glycan family, namely guar gum, which contains such fiber.

- 5 5. The beverage according to any of claims 1 to 4, characterized in that the third component is a natural nutritional oil highly enriched in omega 3 fatty acids, a linseed oil or the oil of the Sacha Inchi nut, which contains more than 50% omega 3 fatty acids and up to 50 mg/100 mL of tocopherol.