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Kombucha: Formulation, chemical composition, and therapeutic potentialities.

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1 **Kombucha: Formulation, chemical composition, and therapeutic potentialities.**

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26 **Abstract**

27 Kombucha is a millennial beverage with great potential due to its functional claims. The
28 infusion of black or green tea leaves (*Camellia sinensis*) and sugar is fermented by a
29 symbiotic culture of bacteria and yeasts (SCOBY) resulting in an acidic and lightly
30 carbonated beverage, kombucha. It offers in its composition phytoconstituents with
31 relevant nutritional valor, among these, flavonoids that stand out for their antioxidant,
32 anti-inflammatory characteristics and their association with decreasing the risks of
33 various diseases. Previous studies *in vivo* and *in vitro* have shown promising results using
34 kombucha as a functional beverage. Those studies promote the search for alternative raw
35 materials for the production of kombucha, in addition, new ingredients interfere in the
36 production, constitution, and nutritional potentialities of the beverage, as well as its
37 functionality in the face of diseases. Thus, this graphical review compiles relevant
38 scientific data on kombucha involving its origin, production, nutritional potential, and
39 possible health benefits associated with its consumption.

40 **Keywords:** Fermented tea; Bioactive compounds; Fermented beverage; Analog
41 kombucha.

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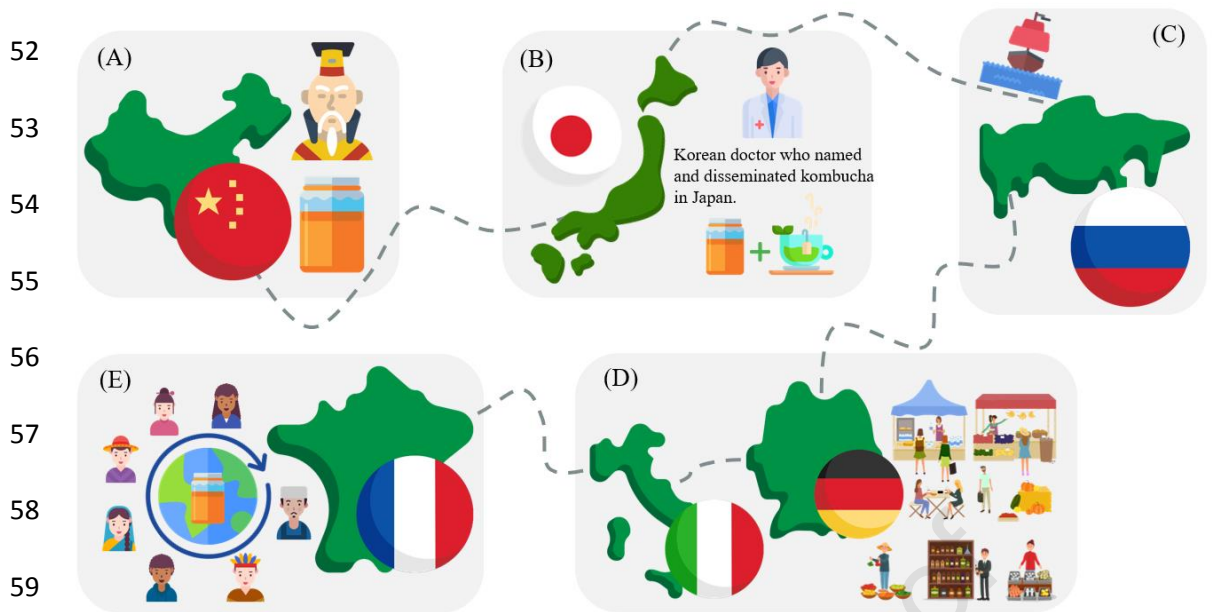
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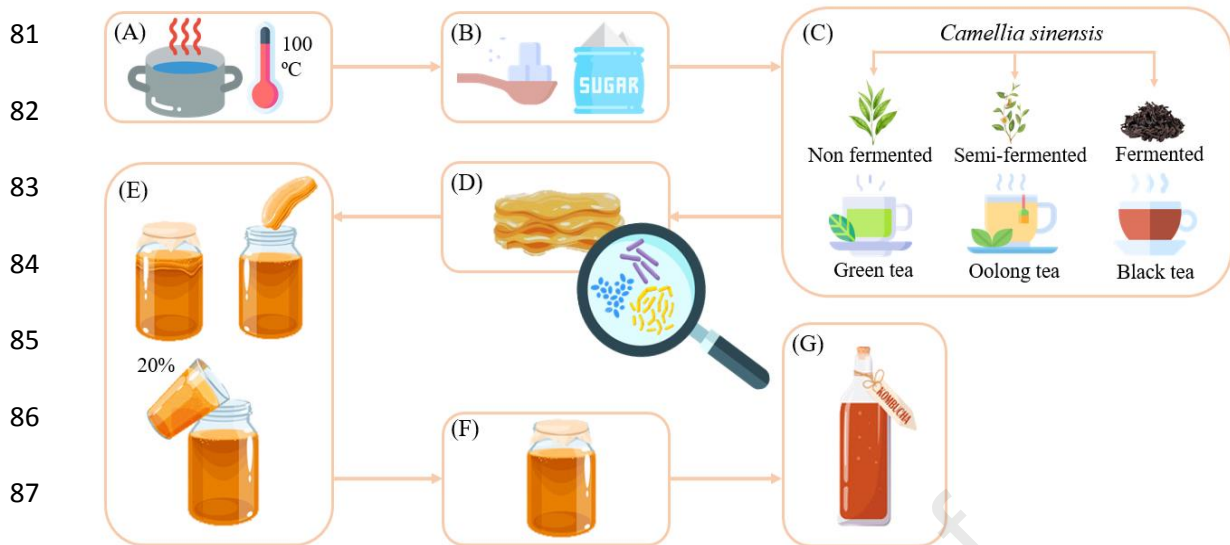
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62 **Fig 1. The historical context of kombucha, its origin, and expansion.** Records show
 63 that kombucha has been a product appreciated for thousands of years, (A) and its
 64 consumption began around 220 BC., in Manchuria, located in northeastern China. During
 65 the Chinese dynasty "*Tsin*", the beverage became popular in the country, as it was already
 66 believed in its energy and detoxifying properties. (B) The Korean doctor named Kombu,
 67 who used the beverage to treat intestinal problems of the emperor at the time, around the
 68 year 414 AD. introduced the beverage in Japan. After this, the beverage gained
 69 prominence and came to be called "Kombucha" in honor of the doctor Kombu
 70 (Chakravorty et al., 2019). (C) Given its nutritional and functional potentialities,
 71 kombucha spread throughout the world and became also known in Europe. Studies report
 72 that it first arrived in Russia via commercial sea routes and (D) expanded to Germany and
 73 Italy in the 20th century, shortly after World War II. In the 1950s, (E) Kombucha also
 74 became popular in France and North Africa (Jayabalan, Malbaša, & Sathishkumar,
 75 2016). Currently, kombucha has gained prominence again and has been widely
 76 disseminated in the world market for beverages and products with functional claims
 77 (Dutta & Paul, 2019).

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91 **Fig 2. Formulation of traditional kombucha.** The process of obtaining kombucha is
 92 considered simple since it does not require large equipment and hard-to-reach ingredients.
 93 According to the region and the studies developed, there are variations in the specificities
 94 and proportions of the materials used, but the methodology described by Jayabalan et al.,
 95 (2014) is referred to as the standard process (Dutta & Paul, 2019). The elaboration process
 96 of kombucha starts from the preparation of the tea, by infusion of *Camellia sinensis* leaves
 97 (green tea, black tea, or oolong tea), then the sugar is added and, at room temperature, a
 98 cellulosic film called "Symbiotic Culture of Bacteria and Yeasts" (SCOBY) is inoculated.
 99 In this film there is a predominance of acetic acid bacteria and yeasts, the SCOBY is
 100 responsible for fermentation and gives the characteristics of the beverage. Some authors
 101 also report the use of previously fermented tea as starters, which can be used to start the
 102 fermentation process (Chakravorty et al., 2019; Jayabalan et al., 2014, 2016; Kapp &
 103 Summer, 2019).

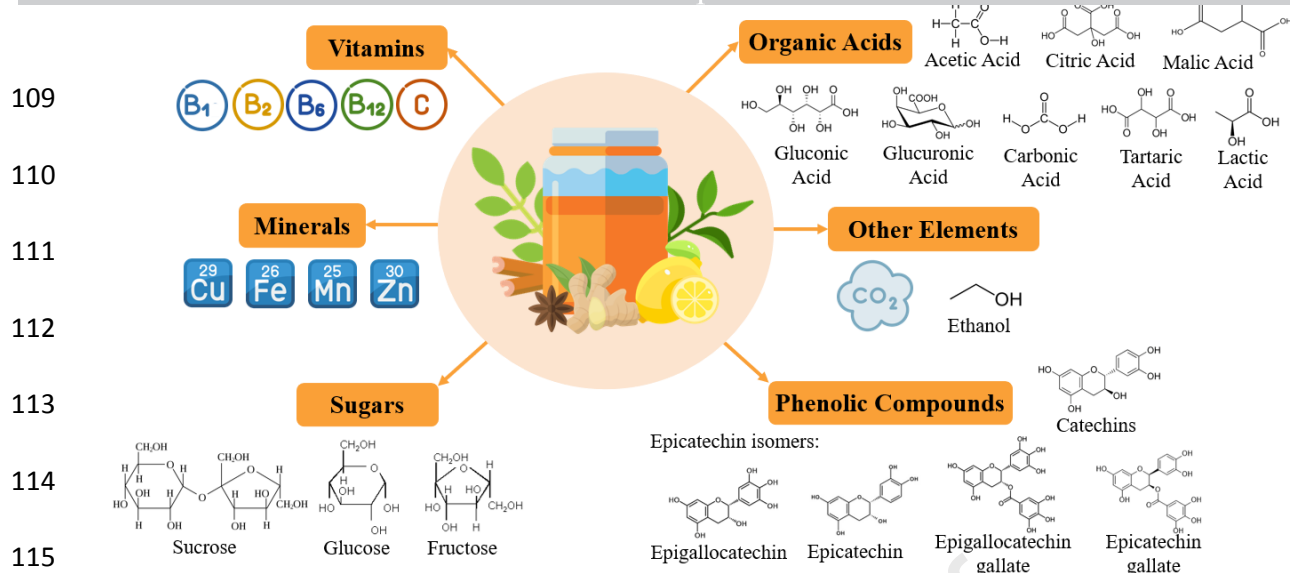
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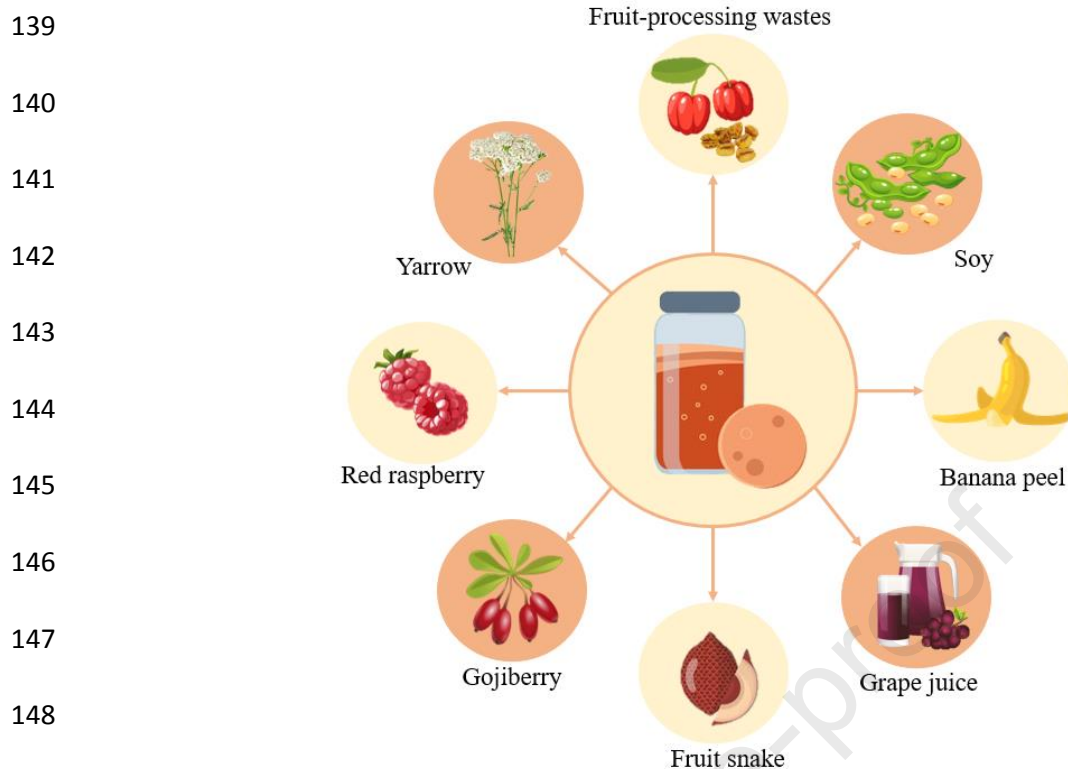
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Fig 3. Chemical composition of traditional kombucha. Kombucha has interesting nutritional valor, mainly due to the benefits of *Camellia sinensis* that are already well described in many literatures. As for the characteristic acidity of the beverage, it may vary according to the time and speed of fermentation and occurs due to the production of organic acids, especially acetic acid (Chakravorty et al., 2016; Vitas et al., 2018). Acetic bacteria, the major part of the SCOBY, synthesize acetic acid from the ethanol produced by yeasts. However, there are other acids in kombucha, such as citric, malic, gluconic, glucuronic, carbonic, tartaric, and lactic, among others (Chakravorty et al., 2019; Vitas et al., 2018; Zubaidah et al., 2019). Others compounds are found in kombucha; ethanol, sugars, mainly glucose and fructose, but also sucrose fractions that are not degraded by yeasts. As well as amino acids, vitamins of the B and C complex, minerals such as iron, zinc, and manganese, and polyphenols that may vary according to the ingredients used and the conditions of the fermentation (Abuduaibifu & Tamer, 2019; Jayabalan et al., 2014, 2016; Rahmani et al., 2019; Villarreal-Soto et al., 2018). Therefore, studies show that the chemical composition of the beverage is directly linked to the ingredients and their proportions, as well as the variation of the fermentation parameters. Thus, these variations can potentiate the production of specific nutritional compounds. Examples include phenolic compounds, known to have several health benefits and are associated with disease prevention (Aspiyanto et al., 2017). Among the compounds found in kombucha are flavonoids, especially catechins and their derivatives, considered functional substances (Chakravorty et al., 2019; Jayabalan et al., 2016; Kapp & Sumner, 2019; Leal et al., 2018).



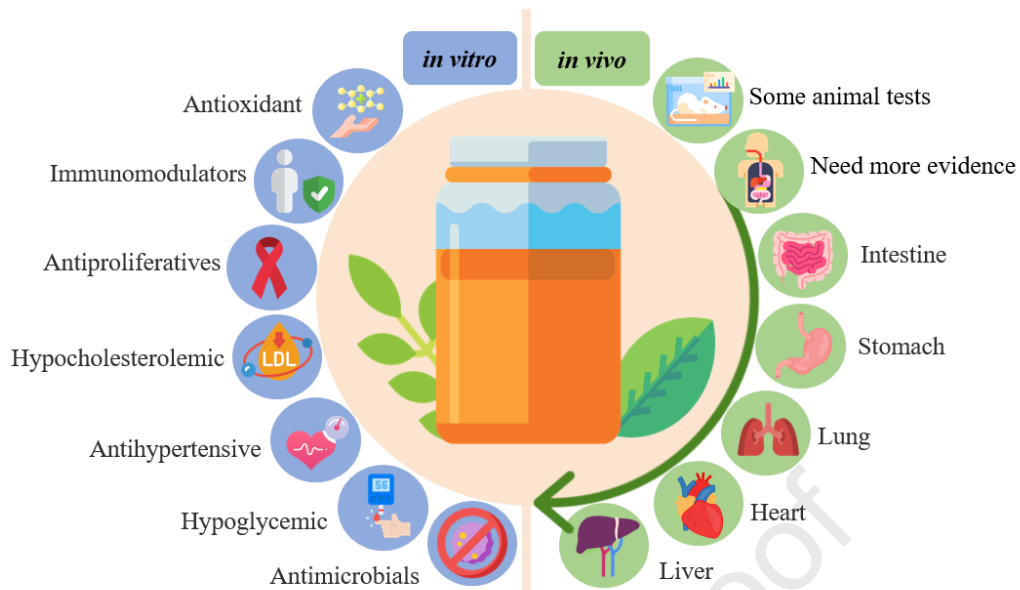
149 Images: Flaticon and Freepik (www.flaticon.com and www.freepik.com)

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151 **Fig 4. Use of alternative substrates in the preparation of analogous kombucha.**

152 Because of the popularization and visibility of kombucha, researchers have been
 153 investigating the variation not only of the concentrations of the ingredients of its original
 154 formulation but of new raw materials and processes. These innovations increase the
 155 possibility of flavors and functionalities of the beverage, which further contributes to the
 156 acceptance of these products, called kombucha analogs. Given this reality, current
 157 research has been replacing *Camellia sinensis* or associating it with other herbs, fruits,
 158 and vegetables for the production of the beverage. Those tests have occurred, either using
 159 these raw materials to prepare the direct infusion and/or its addition to induce a second
 160 fermentation, which favors the flavoring and acceptance of the final product, besides
 161 potentiating the profile of bioactive compounds of the beverage (Emiljanowicz &
 162 Malinowska-Pańczyk, 2019). The use of soy (Xia et al., 2019), yarrow (Vitas et al., 2018),
 163 processed fruit residues (Leonarski et al., 2021), red raspberry (Ulusoy & Tamer, 2019),
 164 banana's peel (Pure & Pure, 2016), grape juice (Ayed, Ben Abid, & Hamdi, 2017), goji
 165 berry (Abuduaibifu & Tamer, 2019), snake fruits (Zubaidah, Dewantari, Novitasari,
 166 Srianta, & Blanc, 2018) and, more recently, the use of umbu-cajá and pitanga pulp (da
 167 Silva Júnior et al., 2021) have been reported in the literature.

EFFECTS ASSOCIATED WITH KOMBUCHA CONSUMPTION



Images: Flaticon and Freepik (www.flaticon.com and www.freepik.com)

Fig 5. Effects associated with kombucha consumption. The biological activities of kombucha and their respective functional and therapeutic potential have been associated with the beverage's chemical constituents and are commonly reported in studies involving *in vitro* and/or *in vivo* tests. Numerous *in vitro* tests have already been reported and found that the kombucha beverage has mainly antioxidant activity, which is well documented in the literature and is mainly associated with the plant used, being endorsed by various methods. Other biological activities linked to the consumption of kombucha have been reported and documented, such as immunomodulatory, antihypertensive, hypocholesterolemic, hypoglycemic, antiproliferative, and antimicrobial. Some tests were developed *in vivo* using animals. However, currently, there are no large studies involving direct effects on the human body associated with the ingestion of kombucha, both in the general shape of the organs and in a specific organ, making it a major obstacle in scientific evidence. However, there are records in the literature about the effect of kombucha on human cells (*in vitro*) (Kapp & Sumner, 2019; Morales, 2020; Sinir et al., 2019).

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202 Conflict of interest

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DECLARATION OF COMPETING INTEREST

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