Natural Honey Beneficial to Health, Its Chemical Composition, and Biochemical Activities: A Review

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Authors’ contributions

This work was carried out in collaboration between both authors. Both authors read and approved the final manuscript.

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ABSTRACT

Since early antiquity, honey has been utilized worldwide as a balanced meal and in supplementary treatment. It contains over 200 different substances, the majority of which are water, sugars, vitamins, enzymes, amino acids, minerals, and phytochemicals. It has health advantages, including microbial inhibition, healing, and its influence on other disorders. Additionally, it stimulates the immune system and helps immune cells mature. The nutritional value of honey is substantially influenced by the type of plants in the region, the climate, and the production process. Different types of honey were examined by various workers, they found a strong relationship between honey type and HMF, saccharose, and phenolic content, as well as acidity and antioxidant activity. For the level of vitamin C, glucose, and fructose, the combination of variety and manufacturing technique has a substantial influence. The focus of this study is on the biochemical processes and its potential health benefits. However, the precise mechanisms of honey’s influence on many illnesses and activities have not yet been fully defined, and more study is necessary to understand their precise contributions.
**INTRODUCTION**

A sweet liquid produced by the honey bees is called honey. Honey is well-known on a global scale since it contains several nutrients that are good for people's health. Egyptians, Greeks, Romans, and Chinese people have all utilized it in the past to treat gastrointestinal injuries and illnesses, such as stomach ulcers. Additionally, it has been employed as a treatment for earaches, sore throats, and coughs [1]. Honey is used internally [2] as a functional meal to offer energy and nutrients to improve the body's organs [3] in addition to being utilized externally. This has been done since the start of history. The quality of honey is significantly influenced by its active ingredients, which include organic acids, flavonoids, polyphenols, glucose, and fructose [4]. Due to its useful qualities and nutritional benefits, honey is produced in many nations across the world and is acknowledged as a significant food that provides energy as well as a treatment.

Furthermore, the biochemical, physiological, and pharmacological properties of honey are well established. The biochemical functions, prospective health advantages, and effects of chemical contaminants in honey are the main areas of this review.

Apiculture is the study and practice of employing substances from honeybee colonies, such as honey, bee bread, bee venom, bee pollen, propolis, and royal jelly, to maintain, lengthen, and prolong life. Bee products have been rapidly incorporated into both conventional and contemporary medicine in recent years. Due to their effectiveness, bee products are currently the subject of several research aimed at determining their specific health benefits and pharmacological qualities, which has accelerated the creation of nutraceuticals and functional foods derived from these products. Functional food is defined as food that, when compared to conventionally remediated and nutritive food, can promote improved physiological or psychological health. These outcomes favourably impact great health preservation, wellbeing, and decrease in chronic disease [5].

Due to its useful qualities and nutritional benefits, honey is produced in many nations across the world and is acknowledged as a significant food that provides energy as well as a treatment. Furthermore, the biochemical, physiological, and pharmacological properties of honey are well established. The biochemical functions, prospective health advantages, and effects of chemical contaminants in honey are the main areas of this review.

**CHEMICAL COMPOSITION OF HONEY**

Each floral source has a unique honey composition, but then again seasonal, environmental factors and processing conditions are similarly significant. More than 200 bioactive compounds are present. Some other term for honey is "supersaturated sugar solution." Natural honey has 82.4% of its mass in carbohydrates, 38.5% in fructose, 31% in glucose, 12.9% in other sugars, 17.1% in water, 0.5% in protein, organic acids, multimineral, amino acids, vitamins, phenols, and a plethora of other minor constituents. The ratio of one form of sugar to another is influenced by the source, such as floral pastures, and to a lesser measure by the enzyme invertase, which dissolves normal sugar in grapes and other fruits. This enzyme can be found in the flower where the bees get their nectar, but it is also in the bee itself [6].

Honey contains 76% sugars (34% glucose, 40.5% fructose, 1.9% sucrose) and 5.5% other carbs. While rapeseed honey stands out for having a larger amount of glucose, acacia and chestnut honey are both quite high in fructose. Numerous studies on the fructose/glucose ratio of different honey samples have been undertaken, discovered that samples of honey dew honey from Croatia had slightly more fructose content (32.4%) than glucose (31%), whereas samples from Macedonia had more...
glucose (36.8%) than fructose (33.6%). According to Ahmed et al. (2014), four samples of honey from various locations in western Algeria range were taken, the glucose and fructose concentration are found from 21.45 to 28.26 g/100 g and 25.20 to 37.64 g/100 g, respectively [7-10]. With a range of 15 to 23%, water is the second-most significant component of honey. The amount of water in honey affects its viscosity, specific weight, maturity, flavour, and crystallization, and is influenced by the weather, the type of bees, the strength of the bee colonies, the humidity and air temperature in the hive, the processing and storage conditions, and the honey's botanical origin.

Due to its hygroscopicity, honey contains a variable amount of water that varies throughout storage according on the air's humidity. Because it affects the stability and resistance of honey to microbiological degradation during storage, it may be claimed that the amount of water in honey plays a significant role in defining both its quality and durability. The probability of fermentation increases with water content [11,12]. Because nectar and pollen are essential components of plants, proteins can be found in honey. Proteins in honey can take the form of simple substances like amino acids or very complicated structures [13]. Protein and amino acid content together make up no more than 0.7% of the total. Nearly all amino acids that are crucial for health are present in honey. Proline, the primary amino acid, is used to gauge how ripe honey is. Normal honeys should include more than 200 mg/kg of proline. Values below 180 mg/kg indicate that the honey has likely been tampered with by the addition of sugar [14].

The components that give honey its scent is known as honey volatiles. Early in the 1960s, studies on honey volatiles were initiated. The majority of volatile volatiles are most likely derived from plants, although some of them are also likely to be bee-added, according to recent research [15] of volatiles recovered from honey. A large number of chemicals have been identified in various honeys up to this point. Secondary metabolites generated from plants include polyphenols and phenolic acids. In plant systematics, these substances have been employed as chemotaxonomic markers. They have been proposed as potential indicators for identifying the honey's botanical origin [16]. According to reports, dark-coloured honeys contain less flavonoids and more phenolic acid derivatives than light-coloured ones [17].

Mineral compounds are present in honey in various concentrations. In addition to several other components, potassium is the major element in honey (as shown in Table 1). The main mineral element is potassium, which makes up an average of around one third of the total, however there are many different trace elements. Numerous studies have revealed that the trace element composition of honey is mostly influenced by its botanical source. 3.68% or so are in minerals [18]. Minerals in honey increase the value of honey for human consumption even if this portion of the honey is not produced in great quantities. The majority of minerals, including potassium, chlorine, sulphur, calcium, sodium, phosphorus, magnesium, silicon, iron, manganese, and copper, are found in honey [19]. Darker honey varieties contain more minerals than lighter ones when compared to the observed mean value. Of course, a darker species can be found that is less wealthy than some lighter species [20-23].

![Fig. 1. (A-B)- Structure of sucrose, glucose and fructose](image-url)
Table 1. Chemical composition of honey per 100 g

<table>
<thead>
<tr>
<th>A- Minerals (Average amount present in mg)</th>
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<tbody>
<tr>
<td>Calcium</td>
<td>4-30</td>
</tr>
<tr>
<td>Chlorine</td>
<td>2-20</td>
</tr>
<tr>
<td>Zinc</td>
<td>0.05–2.00</td>
</tr>
<tr>
<td>Sodium</td>
<td>1.6–17.0</td>
</tr>
<tr>
<td>Copper</td>
<td>0.02–0.60</td>
</tr>
<tr>
<td>Iron</td>
<td>0.03–4.00</td>
</tr>
<tr>
<td>Magnesium</td>
<td>0.7–13</td>
</tr>
<tr>
<td>Phosphorous</td>
<td>2–15.0</td>
</tr>
<tr>
<td>Selenium</td>
<td>0.001–0.003</td>
</tr>
<tr>
<td>Potassium</td>
<td>40.0–350.00</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>B- Proximate (g)</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Fructose</td>
<td>38.2</td>
</tr>
<tr>
<td>Glucose</td>
<td>31.3</td>
</tr>
<tr>
<td>Sucrose</td>
<td>0.7</td>
</tr>
<tr>
<td>Other disaccharides</td>
<td>5.0</td>
</tr>
<tr>
<td>Organic acids</td>
<td>0.5</td>
</tr>
<tr>
<td>Proteins, amino acids</td>
<td>0.3</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>C- Vitamins (mg)</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Ascorbic acid</td>
<td>2.2–2.5</td>
</tr>
<tr>
<td>Thiamin</td>
<td>0.0–0.01</td>
</tr>
<tr>
<td>Riboflavin</td>
<td>0.01–0.02</td>
</tr>
<tr>
<td>Niacin</td>
<td>0.1–0.2</td>
</tr>
<tr>
<td>Pantothenic acid</td>
<td>0.02–0.11</td>
</tr>
<tr>
<td>Pyridoxine (B6)</td>
<td>0.01–0.32</td>
</tr>
</tbody>
</table>

3. BIOACTIVE COMPOUNDS IN HONEY

Honey has a lot of beneficial chemicals (Table 1). Bioactive substances include both essential and non-essential elements found in food chains, such as polyphenols and vitamins. These substances are naturally occurring in food and offer advantageous health advantages. Bioactive substances include phenolic compounds. When there are appropriate functional derivatives present, phenols are described as organic compounds having an aromatic ring that is chemically linked to one or more hydrogenated substituents [24]. Honey also contains trace levels of beneficial substances as phenolic acid, flavonoids, and tocopherol [25]. The antioxidant, antibacterial, antiviral, anti-inflammatory, antifungal, wound-healing, and cardioprotective effects of phenolic compounds, among others, contribute to the functional qualities of bee products [26]. Phenolic acids, flavonoids, ascorbic acid, proteins, carotenoids, and specific enzymes like glucose oxidase and catalase are among the components of honey that have positive health effects [27].

According to the quantity of vitamins, minerals, antibiotic-rich inhibin, carotenoids, free amino acids, enzymes, proteins, Maillard reaction products, and phenolic compounds present in honey composition, the non-sugar components, though present in minor quantities, define a specific type of honey and are bioactive [28]. Flavanoids (apigenin, chrysir, galantine, hesperidin, kaempferol, luteolin, myricetin, and quercurin), which are typically heterocyclic ring compounds, are made up of two aromatic rings connected by a carbon bond. Different flavonoid classes, such as flavanols, flavones, flavonones, flavanols, isoflavones, flavononols, and anthocyanidins, result from variations in the heterocyclic ring compounds. Ring substitutions result in a variety of molecules for each flavonoid family [29]. Phenolic chemicals exert antioxidant capacity (AOC) in a variety of ways, such as metal chelators, free-radical scavengers, or gene modulators of enzymatic and non-enzymatic systems that control cellular redox balance, depending on the molecular structures [30]. A particular phytochemical or mixture of them may be able to identify the location and type of plant from which honey was made [31,32]. Manuka honey, citrus honey, sunflower honey, and lavender honey, among others, all contain methylglyoxal, as do hesperidin, quercetin, and luteolin [33–36].
4. HEALTH BENEFITS OF HONEY

4.1 Wound Management

Honey has long been used as a remedy for boils, ulcers, burns, insect bites, and skin conditions. The effectiveness of honey as an antibacterial agent and a stimulator of wound regeneration has been supported by scientific research [37]. In the wound matrix, honey encourages the activation of dormant plasminogen, which causes the dynamic production of the proteolytic enzyme. Blood clots contract and fibrin is destroyed by plasmin. It is an enzyme that disintegrates fibrin clots in the wound bed that have adhered dead tissues [38]. Clinical research demonstrating the efficacy, specificity, and sensitivity of honey in wound care suggests that it performs better than traditional and contemporary wound care dressings [39].

According to certain situations, honey promotes wound healing even in infected wounds that do not react to antiseptics or antibiotics and wounds that have been contaminated with bacteria that are resistant to antibiotics, like methicillin-resistant Staphylococcus aureus (MRSA). Additionally, honey promotes the formation of a healthy granulated wound bed and helps with autolytic debridement [40]. Malodor is a characteristic of all serious wounds brought on by Peptostreptococcus spp. and Bacteroides spp. anaerobic bacterial species [41]. The metabolism of amino acids from putrefied serum and tissue proteins by bacteria results in the production of foul-smelling substances such as ammonia, amines, and sulphur. As honey releases a significant amount of glucose, a substrate that bacteria prefer to consume over...
amino acids, these molecules are replaced by lactic acids [42]. Fast wound healing, infection clearance, tissue regeneration, reduced inflammation, and improved dressing comfort are some of the therapeutic outcomes seen following honey administration, as well as reduces the tissue adhesion [43].

### 4.2 Paediatric Care

By promoting epithelialization of the affected skin surface, honey also manages skin damage around stomas, such as ileostomy and colostomy [44]. Paediatric dermatitis brought on by frequent use of napkins and diapers, eczema, and psoriasis respond favourably to honey. Patients with psoriasis or atopic dermatitis were studied to see how honey combined with beeswax and olive oil affected their condition. A clinical trial revealed that a honey-based mixture was very well tolerated and produced noticeable improvements. Numerous nitric oxide metabolites found in honey lower the risk of skin infection in psoriasis [45].

### 4.3 Oral Health

Numerous oral conditions, such as halitosis, stomatitis, and periodontal disease, can be helped by honey. Additionally, it has been used to prevent periodontitis, dental plaque, gingivitis, and mouth ulcers. Honey's antibacterial and anti-inflammatory qualities can promote the development of granulation tissue, which in turn helps damaged cells heal [46]. Periodontitis is brought on by the Gram-negative bacterium Porphryromonas gingivalis. In addition to preventing periodontal disease, honey has antibacterial effects on this anaerobic bacterium [47]. Stomatitis, an inflammation of the mouth's mucous membranes, can result in ulcers that are clearly visible and extremely painful. Honey is good against stomatitis and quickly penetrates the tissues [48,49]. Bad breath is a symptom of the oral health disorder halitosis. The majority of the oral cavity's odour is brought on by degrading microbial activity [50]. According to a recent study, honey consumption reduces halitosis because of its potent antibacterial properties brought on by its methylglyoxal component [51].

### 4.4 Gastrointestinal (GI) Disease

Enzymes found in natural honey aid in the absorption of compounds like sugars and starches. The sugar molecules in honey are in a shape that makes them simple for the body to absorb. Additionally, honey contains nutrients like minerals, phytochemicals, and flavonoids that support the body's digestive functions [52]. Salmonella spp., Escherichia coli, Shigella spp., and numerous other Gram-negative species are susceptible to the bactericidal effects of pure honey [53]. Numerous crucially important beneficial microorganisms are found in the gastrointestinal tract (GIT). Bifidobacteria, for instance, is one of the microorganisms needed to maintain a healthy GI system. Consuming foods high in probiotics may help the GIT contain more Bifidobacteria, according to some research. In the presence of prebiotics, this bacteria's biological processes and growth are significantly improved. Natural honey has a high prebiotic content, according to studies [54]. Honey has been identified as a notable dietary supplement that promotes the proliferation of Lactobacillus and Bifidobacteria and catalyses' their probiotic effectiveness in the GIT, according to certain in vitro and in vivo experimental studies [55,56]. Prebiotic components in honey such inulin, oligofructose, and oligosaccharides encouraged an increase in Lactobacillus acidophilus and L. plantarum populations by 10-100 folds under in vitro conditions, which was advantageous for the gut microbiota [57].

### 4.5 Dyspepsia, Gastritis, and Peptic Ulcer

In dyspepsia, a persistent condition, the stomach and first portion of the small intestine mostly act improperly. Epigastric discomfort, heartburn, bloating, and nausea are symptoms of the condition. The early sign of a peptic ulcer, which has the potential to develop into cancer, is dyspepsia. The term “gastritis” describes the lining of the stomach wall's irritation and inflammation. Peptic ulcers are painful erosions or open sore ulcers that develop on the lining of the duodenum or stomach. Helicobacter pylori (H. pylori), the causative agent of peptic ulcers, and gastritis have both been linked to honey [58]. Clinical studies have demonstrated that honey boosted healing effects while reducing stomach acid output. As a result of its antibacterial qualities and preventive effects, honey is consumed as a dietary supplement [59]. Honey's high sugar content and acidic pH are caused by glucose oxidase's conversion of glucose into gluconic acid during the oxidative process. Hydrogen peroxide, which serves as an antibacterial agent, is released by this method. Additionally, fibroblasts and epithelial cell activators are impacted by glucose oxidase, which is necessary for the healing of ulcers brought on by H. pylori [43-44].
4.6 Gastroesophageal Reflux disorder

The condition known as gastroesophageal reflux disease (GERD) is a mucosal infection brought on by the abnormal reflux of stomach contents into the esophagus and even the lungs. Heartburn, inflammation, and acid regurgitation are GERD symptoms. By covering the lining of the esophagus and stomach and inhibiting the upward flow of food and gastric fluid, honey consumption aids this condition. Finally reducing the likelihood of acid reflux, honey can further stimulate the sphincter tissues to help with their renewal [60].

4.7 Constipation and Diarrhea

A widespread and diverse condition known as chronic constipation is characterized by uncomfortable feces (irregular stools and difficult stool passage). Straining, difficult stool expulsion, a feeling of incomplete evacuation, hard or lumpy stools, and a longer time to pass stool are all signs of difficult stool passage [61]. A high frequency of bowel motions accompanied by watery stools is referred to as diarrhea. In comparison to traditional antiviral medication, honey has reduced the aetiology and duration of viral diarrhea [62]. In another instance, raw Manuka honey taken on an empty stomach was beneficial in treating persons with inflammatory bowel syndrome (IBS) who were experiencing severe diarrhea or constipation, bloating, and stomach discomfort [63].

4.8. Pharyngitis and Coughs

Streptococcus spp. causes pharyngitis, also referred to as sore throat, an acute infection of the oropharynx and nasopharynx [64]. In addition to streptococci, sore throats can also be brought on by viruses, non-streptococcal bacteria, fungi, and irritants such chemical pollution. Manuka honey’s anti-inflammatory, antiviral, and antifungal qualities make it useful for treating sore throats. In addition to relaxing the throat, honey coats the inside lining of the throat and kills any potentially hazardous microorganisms [65,66]. According to a study, honey is more effective than dextromethorphan and diphenhydramine in treating cough brought on by upper respiratory infections [67]. Honey’s anti-inflammatory and antibacterial characteristics helped both kids and adults sleep better after consuming it, reducing the severity of recurrent coughs (2.5 ml). According to comparison research on children using various natural items, honey was discovered to be the most frequently utilized treatment for pneumonia 82.4% of the time [68].

4.9 Metabolic and Cardiovascular Fitness

Natural wild honey has cardioprotective and therapeutic effects against the heart diseases and vasomotor dysfunctions brought on by adrenaline. The total phenolic content of honey and radical scavenging activity have been found to be correlated [69]. Consuming honey significantly decreased the risk factors for cardiovascular and metabolic disorders. In addition to improving lipid profiles and maintaining vascular homeostasis, honey demonstrates cardioprotective benefits [70]. Honey contains flavonoids that enhance coronary vasodilation, reduce platelet clotting potential, limit low-density lipoprotein (LDL) oxidation, raise HDL levels, and enhance endothelial function [71]. Honey has been shown to have beneficial effects on metabolic syndromes (MetS) in a study comparing the metabolic response of honey [72]. Hyperglycaemia, hypertension, abdominal obesity, dyslipidaemia, and heightened susceptibility to diabetes, renal, and heart disorders are all signs of MetS. By suppressing inflammatory and angiogenic pathways, honey’s polyphenols lessen atherosclerotic plaques [73]. A clinical trial on hyperlipidaemic individuals revealed that honey reduced total cholesterol (TC) and significantly slowed the rise in plasma glucose levels. A molecule found in honey called nitric oxide (NO), also serves as a cardioprotective agent [74].

4.10 Gastroenteritis

Inflammation of the digestive tract is brought on by gastroenteritis, often known as stomach flu or gastric flu. This syndrome could be brought on by the transfer of infectious organisms through food, water, and person-to-person contact. Dehydration, watery diarrhea, bloating, abdominal cramps, and nausea are all signs of gastroenteritis. This syndrome can be brought on by a variety of infectious organisms, including Salmonella, Shigella, and Clostridium [75]. According to a clinical trial by Abdulrahman from 2010, infantile gastroenteritis can be treated with honey. Due to honey’s high sugar content, which accelerates electrolyte and water absorption in the stomach, the study found that patients with gastroenteritis recovered more quickly when honey was used to replace the glucose in
standard electrolyte oral rehydration solution (ORS) [76].

4.11 Diabetic Foot Ulcer (DFU)

Consuming honey is a cheap and successful kind of therapy for DFU. Microbial infections can make DFU more challenging and impede the healing process. Due to their lowered immune response, patients with diabetic peripheral neuropathy may not exhibit symptoms such as pain, oedema, or redness in addition to the infection, further complicating the diagnosis [77]. According to a review, using honey to treat venous ulcers produced effective results with high patient compliance rates [78]. In patients with locally infected wounds, DFU, Charcot foot ulcerations, and complex concomitant disorders that have not responded to hospital therapy, honey is useful in wound management [79]. Additionally, the presence of honey results in high tolerability and minimal harm to the wound bed.

4.12 Liver and Pancreatic sicknesses

Honey aids in detoxification, liver system balance, and pain relief. Oxidative damage is a cause of complications in the hepatic system. Antioxidant properties of honey may have a preventative effect on a damaged liver. According to a study on rats with paracetamol-induced liver damage, honey's antioxidant and hepatoprotective properties significantly reduced liver damage. As it provides sufficient glycogen storage in liver cells, honey, which has a fructose to glucose ratio of 1:1, may assist to produce improved blood sugar levels, which is beneficial for patients with fatty liver disease. Stress hormones are released when the liver doesn't have enough glycogen store, which over time damages glucose metabolism. Insulin resistance is the main cause of fatty liver disease and is brought on by impaired glucose metabolism. Following treatment with Tualang honey, a different study found that blood glucose levels significantly decreased [80-82].

5. CANCERS AND ONCOGENESIS

5.1 Colorectal Cancer

A polyp, which often originates on the inner lining of the colon or rectum and develops toward the centre, is where the majority of colorectal malignancies start. While some polyps are not harmful, some will develop into adenomas and may eventually cause cancer. The multiplication of colon cancer cells was suppressed by the honey, according to a study, that examined the chemo preventive effects of Gelam and Nenas monofloral honeys against colon cancer cell lines. The impact of honey was investigated using colon cancer cells that had been exposed to hydrogen peroxide-induced inflammation. The findings demonstrated that honey reduced inflammatory activity in malignant cells. Another study looked into how crude honey affected the apoptosis of colon cancer cell types. The study verified honey's ability to inhibit cell proliferation in these tissues. Additionally, considerable antiproliferative effect against colon cancer cells was seen at high phenolic concentrations (like those of quercetin and flavonoids). Cell cycle arrest, activation of the mitochondrial pathway, induction of mitochondrial outer membrane permeabilization, induction of apoptosis, modulation of oxidative stress, reduction of inflammation, modulation of insulin signaling, and inhibition of angiogenesis in cancer cells are some of the molecular mechanisms underlying honey's antiproliferative and anticancer effects. Additionally, honey has the capacity to influence cancer-promoting proteins, genes, and cytokines. A number of honey constituents, including chrysin, quercetin, and kaempferol, have been demonstrated to stop the cell cycle in human melanoma, renal, cervical, hepatoma, colon, and oesophageal cancer cell lines at various phases, including G0/G1, G1, and G2/M. The mitochondrial pathway is affected by a number of stimuli, including nutrition, physical stress, oxidative stress, and damage from chemotherapy and radiation, two of the most common cancer therapies. These triggers lead to the release of many proteins from the mitochondria's intermembrane space (IMS), including cytochrome c, which ultimately results in cell death. Honey contains flavonoids that are excellent at triggering the mitochondrial pathway and releasing proteins with cytotoxic potential. The most common anticancer strategy, induction of mitochondrial outer membrane permeabilization (MOMP), resulting in protein leakage from the IMS, which inexorably leads to cell death. Honey reduces the propensity of the mitochondrial membrane to cause MOMP in cancer cell lines. It has also been shown that honey increases the depolarization of the mitochondrial membrane, enhancing the apoptotic impact of tamoxifen. It has been demonstrated that honey's flavonoid components, namely quercetin, cause MOMP and cancer cell death [83-85].
5.2 Liver Cancers

Hepatocellular carcinoma is the most typical kind of liver cancer (HCC). In a number of experimental experiments, the anticancer effects of honey on liver cancer cells have been examined. Nitric oxide (NO) levels were significantly reduced and the number of HepG2 cells was significantly reduced in HepG2 cells treated with honey. As a result, the cells' total antioxidant profile improved. Reactive oxygen species (ROS) encourage the survival of HepG2 cells, and sufficient amounts of ROS stimulate cell division and differentiation. This investigation was backed by the fact that the honey treatment reduced the level of NO. As a result, HepG2 cell numbers are decreased and malignant cell growth is inhibited by decreased ROS and increased antioxidant effectiveness. Abdel Aziz et al. also looked at how honey affected HepG2 cell lines in another study. According to the study, honey had distinct concentration-dependent cytotoxic, antimetastatic, and antiangiogenic effects on HepG2 cells [81,83,86].

5.3 Breast Cancer

Growth and spread of breast cancer are significantly influenced by imbalances in estrogen signalling pathways and estrogen propagating levels. The estrogen receptor (ER) signalling pathway is frequently targeted by breast cancer treatments. Because of their similar structural makeup to mammalian estrogen, phytoestrogens are a subclass of phytochemicals that can bind to estrogen receptors. The effectiveness of honey in modifying the ER signalling system has been examined in several experimental experiments. In MCF-7 cells, honey has biphasic action, according to another study. When phytoestrogens bind to estrogen receptors, this biphasic action of honey is characterized by an antiestrogenic effect at lower doses and an estrogenic effect at higher concentrations. Additionally, it has been noted that quercetin induces apoptotic effects via ER- and ER-dependent pathways. On the other hand, Tualang honey's cytotoxic effects against human breast cancer cells were shown by increased lactate dehydrogenase (LDH) release, which further showed the cytotoxic characteristics of honey. The study also demonstrated that non-malignant breast cells are not affected by honey's cytotoxic actions; rather, only breast cancer lines are affected. This proves that Tualang honey has a fair chance of success as a chemotherapeutic drug and exhibits highly specific and selective cytotoxic actions toward breast cancer cell lines [86-94].

6. CONCLUSION

The current review focuses on the potential health benefits of honey. It is very rich in bioactive components such as flavonoids, phenolic acids, phenolic compounds, terpenes, and enzymes, and it prevents several diseases and has health-promoting biological functions. Honey has a wide range of benefits and important nutritional properties and functional value. In this way, honey can be developed into a potent non-therapeutic drug. In conclusion, the complexity and variability of honey composition rely on its botanical and geographic origin. Each component has a specific role in nutrition, biology, and technology. They work together to increase honey's overall usefulness, making it special and superior to other natural sweeteners in terms of supplying energy and health advantages.

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COMPETING INTERESTS

Authors have declared that no competing interests exist.

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