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Potential pharmacological benefits of Ginger (*Zingiber officinale*) – A review

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Abstract

Ginger (*Zingiber officinale* Roscoe, Zingiberacae) is widely used as medicine since ancient times. This review summarizes the most relevant reports implying diverse pharmacological efficacies of ginger. Several scientific investigations focus on isolation, purification and characterization of active phytoconstituents from *Zingiber officinale* with pharmacological benefits. These include flavonoids, essential oils, carbohydrates, phenolic compounds, alkaloids, saponins, terpenoides, tannin, glycosides, steroids, minerals, proteins, lipids, fibers, fatty acids, lecithins, protease, calcium, phosphorous, potassium along with many vitamins like riboflavin, thiamine, vitamin C and niacin. It has the ability to act as antioxidative, anti-diabetic, anti-inflammatory, antimicrobial, neuro-protective and anti-carcinogenic agent to manage and cure a variety of ailments with minimal or no side effects. Further clinical and preclinical trials are warranted to elucidate the consumption of ginger as an alternative pharmacological adjunct.

Key words: Zingiber officinale, anti-oxidants, anti-diabetic and anti-microbial

 Full length article
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1. Introduction

Since the earliest times, traditional medicine systems are based upon plants. Even the modern era is dependent upon the same centuries old system to manage and treat disease. World Health Organization investigated that 80% population of the globe depend on folk medicines [1]. The demand of herbal medicines increase in both developed and under developed countries, therefore, the pharmaceutical companies are taking interest in herbal medicines [2].

Zingiber officinale (Zingiberacae) has been used in Indian and Chinese medicines for more than 25 era. It has been used in the world as a herbal remedy, condiment and cooking spice. Chinese people used Zingiber officinale as an anti-nausea remedy, as digestive aid and to cure bleeding ailments. Ginger is also used to cure toothache, baldness, respiratory and snakebite conditions. It is considered a dry, pungent and yang herb which is used for disorders caused by cold conditions. Ginger has been used widely in Ayurveda medicine to block excessive clotting, for arthritis and hypercholesterolemia treatment [3].

Z. officinale can grow in both tropical and in subtropical regions and can adopt both humid and warm *Hussain and Manzoor*, 2017 conditions up to 1500 meter sea level. *Z. officinale* is produced in Bangladesh, Pakistan, India, US, Nepal, Taiwan, Nigeria, China and in other regions of world. Full grown *Z. officinale* plant is about 2 meter long. The shoots grow from bulb on the bottom of the ginger plant. The rhizome of *Zingiber officinale* grows underground. Its rhizome is thick and tuberous. Mostly rhizome is broad lobed, aromatic, enclosed like scars, fleshy and knobbly. While the leaves are long, green, blade slowly reduce to a point, sheathing bases and 2 to 3 centimeter broad [4].

2. Phytochemistry

Z. officinale rhizome constituents vary depending upon the area of origin as well as on rhizome conditions. Z. officinale rhizome is widely used in food products due to its flavor, aroma and nutritional composition. Rhizome extracts have elevated level of compounds such as gingerol, shogaols, zingerone and paradol. It was confirmed that gingerol and shogaol are the major components that show pharmacological activities [5]. From the volatile oil of Z. officinale rhizome, main active ingredients isolated are the zingiberene, sesquiterpenes, zingiberol and bisapolene [3].

Other rhizome constituents include fat, minerals, proteins, lipids, fibers, fatty acids, carbohydrates, lecithins, protease, calcium, phosphorous, potassium along with many vitamins like riboflavin, thiamine, vitamin C and niacin [6]. It was reported by [7] that rhizomes possess essential flavonoids, oils, carbohydrates, phenolic compounds, alkaloids, saponins, terpenoides, tannin, glycosides, steroids, and proteins as the main phytochemical groups.

3. Antioxidant activity

Oxidative or radical stress can destroy molecules and antioxidants especially those of natural origin play an important role in against oxidative damage. Several studies suggested antioxidant efficacies of Z. officinale rhizomes [4, 8-11]. [12, 13] stated that Z. officinale extract possess free radical scavenging activity and gingerol inhibits lipid peroxidation. The oleoresin of ginger and oil extract also revealed significant antimicrobial and antioxidant activities [14]. Ginger components such as 1-dehydro-6-gingerdione and 6 dehydroshogaol are effective inhibitors of the NO compound in stimulated macrophages [15]. In another study, [16] observed that 6-shogaol has powerful antioxidant activity. Furthermore, [17] indicated that phenolic components possess strong antioxidative, anti-inflammatory and anti-carcinogenic properties. The active pungent compound 6-gingerol lessened gene expression decreased of cysts in ovaries and restored biological parameters to normal in rat models [18].

4. Anti-diabetic activity

Diabetes mellitus, a disabling metabolic disorder has affected about 150 million people worldwide [19]. Several studies indicated at anti-diabetic potentials of *Z. officinale*.

Diverse diabetes suppressive mechanisms of Z. *officinale* are proposed [20, 21]. It was observed by [22] that flavonoids components of Z. *officinale* act against diabetes mellitus either by avoiding glucose absorption or by enhancing glucose tolerance. The flavonoids either act as insulin mimetics or as insulin secretagogues. When [23] studied the hypoglycaemic potential of Z. *officinale* in streptozotocin induced diabetic mice, it decreased the cholesterol, triacylglycerol and serum glucose levels in diabetic rats as compared to control rats. The inhibition of carbohydrate digestive enzymes such as alpha glucosidase and alpha amylase through plants can be attributed to their phenolic components [24]. Z. *officinale* inhibits the enzymes that regulate carbohydrate metabolism and ultimately hyperglycemia in diabetic conditions.

Similarly, [25] conducted *in vitro* study to assess enzyme inhibitory potentials of *Z. Officinale* extracts. Ginger activities against two enzymes were found to connect with total phenolic constituents of shogaols and gingerol in extracts. However, [26] reported alpha glucosidase inhibitory effects of *Z. officinale* extracts with no effect on alpha amylase activity. Recently, it was stated by [27] that 6-paradol, a pungent ginger component can cause hypocholesterolemia and hypoglycemia. Similarly, [28] indicated reduced glucose level, insulin, leptin and enhanced lipid profile in high fat diet rats by the use of hydro ethanolic extract of *Z. officinale*.

5. Antimicrobial activity

Plant derived antimicrobial compounds have massive remedial potential. These are useful against infectious ailments with minimal side reactions that are associated with synthetic antimicrobial agents. The antimicrobial properties of plant extracts usually arise from secondary plant metabolites such as steroids, tannins, alkaloids, flavonoids, resins, gums and phenolic compounds [29]. [30] stated that antimicrobial properties of medicinal plant extracts depends on parameters such as technique employed, microorganism tested, plant material used and growth medium. For better study valuable quality of plant extracts should be preferred. Extraction and the solvent system possibly both modify final results. Different medicinal plant extracts may show different results.

Rhizomes of *Zingiber officinale* possess antimicrobial abilities [31, 32]. It has been investigated by [33] that ethanolic extract of *Z. officinale* rhizome inhibited bacterial growth. When methanolic extracts of *Z. officinale* were tested against nineteen strains of *Helicobacter pylori*, crude extracts that contain gingerols, inhibited the growth of all strains [34].

[35] reported antibacterial activities of crude flavonoids, polysaccharides, ethanolic and aqueous extracts of Z. officinale against microbes S. pyogenes, H. influenza, S. aureus and S. pneumonia. Ethanolic and flavanoids extracts revealed antibacterial properties against bacteria while aqueous and polysaccharide extracts showed no activity. Earlier, [36] demonstrated antimicrobial potencies of various extracts (ethyl acetate, water, n-hexane and ethanol) of Z. officinale rhizome against Staphylococcus epidermidis, Streptococcus viridians and Coliform bacillus. It was observed that all the extract except H₂O extract inhibited the bacterial growth. Ethanolic extract showed highest antibacterial activity. Strong antifungal activity of ethanolic extracts of Z. officinale against C. albicans was demonstrated by [37]. [38] conducted a study to examine antiviral action of gingerol, shogoal and ingenol isolated from Z. officinale rhizome. Gingerol was reported as effective inhibitor of *M. tuberculosis* and *M. avium*.

Table 1: Diverse Pharmacological benefits

Effective Components	Pharmacological activity	Mode of action	References
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6 Shogaol	Anti-cancerous effect	Stimulation of apoptosis, inactivation of VEGF pathways lead to antineoplastic effects	[39]
Ginger extract	Osteoarthritis therapy	By decreasing inflammation mediators	[40,41]
6 Shogaol	Anti-inflammation effect	Suppression of pro-inflammatory chemokines and cytokines (RANTES, MCP-1)	[42]
Gingerol and its related compound	Antioxidant activities	By scavenging free radicals	[43]
Sesquiterpenes, shogaols, gingerols, monoterpenes and shogaols	Cure of vomiting and nausea	By anti-serotonin and anticholinergic activity	[44]
6-Gingerol	Anti-tumor activity	Via induction of apoptosis, modulation of genetics.	[45]
6-Shogaol (Alkanone from Ginger)	Antioxidant activity	Induces apoptotic cell death of human hepatoma p53 mutant mahlavu subline via an oxidative stress-mediated caspase	[46]
10-gingerol	Antimicrobial activity	Antibacterial activity against periodontal bacteria	[47]
Ginger extract	Anti-diabetic activity	By lowering glucose level, insulin, body weight etc. in rats	[48]
6-shogaol	anticancer activities against breast cancer	Inhibition of cell invasion reduction of matrix metalloproteinase-9 expression	[49]
Ginger extract in ethanol	Anti- hyperlipidemic effect	By decreasing cholesterol, triglycerides to elevate HDL	[50]
Ginger extract in ethanol Zingiberone	Hypoglycemic effect	Decreasing of blood sugar level by suppression of oxidative stress and anti-inflammation suppressors, to elevate insulin sensitivity	[51, 52]
Phenolic, 6 shogaol, flavonoids	Neuro-protective Effect	Acceleration of brain anti-oxidant resistance mechanisms	[52]
Terpenoides from Zingiber officinale	Induction of apoptosis in endometrial cancer cells	By the activation of p53.	[53]
Ginger and its constituent	Treatment of Colorectal cancer	By increasing the lymphocyte counts in colorectal cancer patients	[54]

Essential oil and oleoresin of Zingiber officinale Roscoe.	Antimicrobial and antioxidant activity	Scavenge ABTS+ free radical. Inhibit NO synthesis.	[14]
Active compound zerumbone	Treatment of cancer cells	Enhanced radiation-induced DNA damage and inhibited nuclear expression of DNA repair proteins ataxia-telangiectasia mutated (ATM) and DNA-PKcs	[55]
Zingiber officinale Essential oils	Anticancer Activity	Cytotoxic potential against Hela cells	[56]
Zingiber officinale	Induction of apoptosis	Programmed death in <i>T. vaginalis</i>	[57]
6-gingerol and 6-shogaol	Relief of oral ulcerative mucositis-induced pain	Blockage of sodium channel	[58]
Zingerone and its novel derivative	Anticancer activity	Inhibition of beta factor (TGF-β1) induced epithelial-mesenchymal transition	[59]
6-gingerol	Effect on adenosine monophosphate (AMP) activated protein kinase Nuclear factor kappa B pathway (AMPK- NF-Kb)	Up regulation of Sirt-6, enhanced phosphorylation of AMPK, down regulation of resistin, decrease of inflammatory molecules P65.	[60]

4. Conclusions

Ginger is an essential medicinal herb and widely used in Siddha, Chinese and in Ayurveda, medicine etc. Consumption of the *Z. officinale* plays a vital role in controlling of human ailments.

Biochemical analysis of ginger revealed its remedial potential and development of new drugs from ginger could be emphasized for prevention of numerous diseases.

References

- WHO. (2002). Traditional Medicine Strategy Launched: WHO News, Geneva, Switzerland. 80: 610.
- P. C. Chikezie, O. A. Ojiako and K. C. Nwufo. (2015). Overview of anti-diabetic medicinal plants: The Nigerian research experience. Journal of Diabetes and Metabolism. 6: 6.
- [3] M. S. Moghaddasi, and H. H. Kashani. (2012). Ginger (Zingiber officinale): A review. Journal of medicinal plant research. 6: 4255-4258.
- [4] G. Kumar, L. Karthik and K. V. B. Rao. (2011). A review on pharmacological and phytochemical properties of Zingiber officinale Roscoe (Zingiberaceae). Journal of Pharmacy Research. 4: 2963-2966.

- [5] R. K. Mishra, K. Anil and K. Ashok. (2012). Pharmacological activity of Zingiber officinale. International Journal of Chemical Sciences. 1: 1423-1426.
- [6] T. A. Ibrahim, I. B. O. Dada and R. A. Adejare. (2010). Comparative phytochemical properties of crude ethanolic extracts and physicochemical characteristics of essential oils of Myristical fragrans (nutmeg) seeds and Zingiber officinale (ginger) roots. Electronic Journal of Environmental, Agricultural and Food Chemistry. 9: 1110-1116.
- [7] P. R. S. Adel, and J. Prakash. (2010). Chemical composition and antioxidant properties of ginger root (Zingiber officinale). Journal of Medicinal Plants Research. 4: 2674-2679.
- [8] I. Hinneburg, D. Dorman and H. J. Hiltunen. (2006). Antioxidant activities of extracts from selected culinary herbs and spices. Food Chemistry. 97: 122-129.
- [9] G. S. Kumar, H. Nayaka, S. M. Dharmesh and P. V. Salimath. (2006). Free and bound phenolic antioxidants in amla (Emblica officinalis) and turmeric (Curcuma longa). Journal of Food Composition. 19: 446-452.

- [10] M. Cousins, J. F. Adelberg, Chen, F and J. Rieck. (2007). Antioxidant capacity of fresh and dried rhizomes from four clones of turmeric (Curcuma longa L.) grown in vitro. Industrial Crops and Products. 25: 129-135.
- [11] P. Jegajeevanram, N. M. I. Alhaji and S. Velavan. (2015). In vitro antioxidant activity of mango ginger rhizome. World Journal of Pharmaceutical Research. 4: 1739-1745.
- [12] A. R. H. M. Tahtawy, E. A. M. Bastawesy, A. M. G. Monem, Z. K. Zekry, A. H. A. Mehdar and E. M. M. Merzabani. (2011). Antioxidant activity of the vola¬tile oils of Zingiber officinale (ginger). Spatula DD Journal on Complementary Medicine and Drug Discovery. 1: 1-8.
- [13] A. H. Rahmani, M. F. M. Shabrmi and S. M. Alyl. (2014). Active ingredients of ginger as potential candidates in the prevention and treatment of diseases via modulation of biological activities. International Journal of Physiology, Pathophysiology and Pharmacology. 6:125-136.
- [14] Y. Bellik. (2014). Total antioxidant activity and antimi¬crobial potency of the essential oil and oleo¬resin of Zingiber officinale Roscoe. Asian Pacific Journal of Tropical Disease. 4: 40-44.
- [15] F. Li, Y. Wang, K. L. Parkin, V. Nitteranon, J. Liang, W. Yang, Y. Li, G. Zhang and Q. Hu. (2011). Isolation of qui¬none reductase (QR) inducing agents from gin¬ger rhizome and in vitro anti-inflammatory activity. Food Research International. 44: 1597-1603.
- [16] S. Dugasani, M. R. Pichika, V. D. Nadarajah, M. K. Balije-palli, S. Tandra and J. N. Korlakunta. (2010). Comparative antioxidant and anti-inflammatory effects of [6]-gingerol, [8]-gingerol, [10]-gingerol and [6]-shogaol. Journal of Ethnopharmacology. 127: 515-520.
- [17] M. Khader, N. Bresgen and P. M. Eckl. (2010). Antimutagenic effects of ethanolic extracts from three Palestinian medicinal plants. Journal of Ethnopharmacology. 127: 319-324.
- [18] P. S. Pournaderi, P. Yaghmaei, H. Khodaei, Z. Noormohammadi and S. H. Hejazi. (2017). The effects of 6-Gingerol on reproductive improvement, liver functioning and cyclooxygenase-2 gene expression in estradiol valerate-Induced polycystic ovary syndrome in wistar rats. Biochemical and Biophysical Research Communications. 17: 1-8.
- [19] M. Iroganachi, C. O. Eleazu, P. N. Okafor and N. Nwaohu. (2015). Effect of unripe plantain (Musa paradisiacal) and ginger (Zingiber officinale) on blood glucose, body weight and feed intake of streptozotocin induced diabetic rats. Open Biochemistry Journal. 9: 1-6.

- [20] J. S. Ahmad and S. Abbas. (2011). Hypoglycemic effect of Zingiber Officinale in alloxan induce diabetic rat. Pakistan Veterinary Journal. 5: 160-162.
- [21] R. Smit, K. Neeraj and K. Preeti. 2013. Traditional medicinal plants used for the treatment of diabetes. International Journal of Pharmaceutical and Phytopharmacological Research. 3: 171-175.
- [22] G. Brahmachari. (2011). Bio-flavonoids with promising antidiabetic potentials: A critical survey. Opportunity, Challenge and Scope of Natural Products in Medicinal Chemistry. 3: 187-212.
- [23] A. Z. M. Amin, M. Thomson, A. K. K. Qattan, P. R. Shalaby and M. Ali. (2006). Antidiabetic and hypolipidaemic properties of ginger (Zingiber officinale) in streptozotocin-induced diabetic rats. British Journal of Nutrition. 96: 660-666.
- [24] A. Stephen, Adefegha, O. Ganiyu, S. Olasunkanmi, Omojokun, M. Tech, O. Tajudeen, Jimoh, and I. Oyeleye. (2016). In vitro antioxidant activities of African birch (Anogeissus leiocarpus) leaf and its effect on the a-amylase and a-glucosidase inhibitory properties of acarbose. Journal of Taibah University Medical Sciences. 10: 1-7.
- [25] M. P. Rani, K. P. Padmakumari, B. Sankarikutty, O. L. Cherian, V. M. Nisha and K. G. Raghu. (2011). Inhibitory potential of ginger extracts against enzymes linked to type 2 diabetes inflammation and induced oxidative stress. International Journal of Food Sciences and Nutrition. 62: 106-110.
- [26] L. G. Ranilla, Y. I. Kwon, E. Apostolidis and K. Shetty. (2010). Phenolic compounds, antioxidant activity and in vitro inhibitory potential against key enzymes relevant for hyperglycemia and hypertension of commonly used medicinal plants, herbs and spices in Latin America. Bioresource Technology. 101: 4676–4689.
- [27] C. K. Wei, Y. H. Tsai, M. Korinek, P. H. Hung, M. El-Shazly, Y. B. Cheng, Y. C. Wu, T. J. Hsieh and F. R. Chang. (2017). 6-Paradol and 6-Shogaol, the pungent compounds of ginger, promote glucose utilization in adipocytes and myotubes, and 6paradol reduces blood glucose in high-fat diet-fed mice. International Journal of Molecular Sciences. 18: 168.
- [28] D. L. N. Heras, V. M. Munoz, M. B. Fernandez, S. Ballesteros, L. A. Farre, R. B. Roso and V. Lahera. (2017). Molecular factors involved in the hypolipidemic- and insulin-sensitizing effects of a ginger (Zingiber officinale Roscoe) extract in rats fed a high-fat diet. Applied Physiology, Nutrition and Metabolism. 42: 209-215.
- [29] A. Kapoor, G. Kaur and R. Kaur. (2015). Antimicrobial activity of different herbal plants

extracts: A review. World Journal of Pharmaceutical Sciences. 4: 422-459

- [30] Z. M. Ross. (2001). Antimicrobial properties of garlic oil against human enteric bacteria: evaluation of methodologies and comparisons with garlic oil sulfides and garlic powder. Applied and Environmental Microbiology. 67:475–480.
- [31] M. Habsah. (2000). Screening of Zingiberaceae extracts for antimicrobial and antioxidant activities. Journal of Ethnopharmacology. 72: 403-410.
- [32] D. Srinivasan. (2001). Antimicrobial activity of certain Indian medicinal plants used in folkloric medicine. Journal of Ethnopharmacology. 74:217-220.
- [33] A. P. Martins. (2001). Essential oil composition and antimicrobial activity of three Zingiberaceae from Sao Tome and Principe. Planta Medica. 67: 580-584.
- [34] G. B. Mahady. (2003). Ginger (Zingiber officinale Roscoe) and the gingerols inhibit the growth of Cag A+ strains of Helicobacter pylori. Anticancer Research. 23:3699-3702.
- [35] D. Gao and Y. Zhang. (2010). Comparative antibacterial activities of crude polysaccharides and flavonoids from Zingiber officinale and their extraction. Asian journal of traditional medicines. 5: 235-238.
- [36] S. P. Malu, G. O. Obochi, E.N. Tawo and B. E. Nyong. (2009). Antibacterial activity and medicinal properties of ginger (Zingiber officinale). Global Journal of Pure and Applied Sciences. 15: 365-368.
- [37] Z. Atai, M. Atapour and M. Mohseni. (2009). Inhibitory effect of ginger extract on Candida albicans. American Journal of Applied Sciences. 6: 1067-1069.
- [38] P. Miri, J. Bae and D. S. Lee. (2008). Antibacterial activity of [10]-gingerol and [12]-gingerol isolated from ginger rhizome against periodontal bacteria. Phytotherapy Research. 22: 1446-1449.
- [39] Y. J. Surh, K. K. Park, K. S. Chun, L. Lee, E. Lee and S. S. Lee. (1998). Anti-tumor-promoting activities of selected pungent phenolic substances present in ginger. Journal of Environmental Pathology, Toxicology and Oncology. 18: 131-139.
- [40] H. Bliddal, A. Rosetzsky, P. Schlichting, M. S. Weidner, L. A. Andersen, H. H. Ibfelt, K. Christensen, O.N. Jensen and J. Barslev. (2000). A randomized, placebo-controlled, cross over study of ginger extracts and ibuprofen in osteoarthritis. Osteoarthritis and Cartilage. 8: 9-12.
- [41] R. D. Altman, and K. C Marcussen. (2001). Effects of a ginger extract on knee pain in patients with osteoarthritis. Arthritis & Rheumatology. 44: 2531-2538.

- [42] S. C. Penna, M. V. Medeiros, F. S. Aimbire, N. H. C. Faria-Neto, J. A. Sertie and L. R. A. Martins. (2003).Anti-inflammatory effect of the hydralcoholic extract of Zingiber officinale rhizomes on rat paw and skin edema. Phytomedicine. 10: 381-385.
- [43] Y. Masuda, H. Kikuzaki, M. Hisamoto and N. Nakatani. (2004). Antioxidant properties of gingerol related compounds from ginger. Biofactors. 21: 293-296.
- [44] E. Bryer. 2005. A literature review of the effectiveness of ginger in alleviating mild-to-moderate nausea and vomiting of pregnancy. Journal of Midwifery & Women's Health. 50: 1-3.
- [45] Y. J. Park, J. Wen, S. Bang, S. W. Park and S. Y. Song. (2006). [6]-Gingerol induces cell cycle arrest and cell death of mutant p53-expressing pancreatic cancer cells. Yonsei Medical Journal. 47: 688-697.
- [46] C. Y. Chen, T. Z. Liu and Y. W. Liu. (2007). 6-Shogaol (Alkanone from Ginger) induces apoptotic cell death of human hepatoma p53 mutant mahlavu subline via an oxidative stress-mediated caspasedependent mechanism. Journal of Agricultural and Food Chemistry. 55: 948–954.
- [47] M. Park, J. Bae and D. S. Lee. (2008). Antibacterial activity of [10]-gingerol and [12]gingerol isoated from ginger rhizome against periodontal bacteria. Phytotherapy Research. 22: 1446-1449.
- [48] S. Nammi, S. Sreemantula and B. D. Roufogalis. (2009). Protective effects of ethanolic extract of Zingiber officinale rhizome on the development of metabolic syndrome in high-fat diet-fed rats. Basic & Clinical Pharmacology & Toxicology. 104: 366– 373.
- [49] H. Ling, H. Yang, S. H. Tan, W. K. Chui and E. H. Chew. (2010). 6-Shogaol, an active constituent of ginger, inhibits breast cancer cell invasion by reducing matrix metalloproteinase-9 expression via blockade of nuclear factor-κB activation. British Journal of Pharmacology. 161: 63-77.
- [50] A. O. Bolanle. (2011). Effect of ginger powder (Zingiber officinale) on plasma lipid profile and liver enzyme activities of hypercholesterolemic rats. Journal of Life Science. 5: 12-716.
- [51] V. Vats, J. K. Grover and S. S. Rathi. (2002). Evaluation of anti-hyperglycemic and hypoglycemic effect of Trigonella foenumgraecum linn, Ocimum sanctum linn and Pterocarpus marsupium Linn. in normal and diabetic alloxanized rats. Journal of Ethnopharmacology. 79: 95-100.
- [52] K. R. Shanmugam, K. Mallikarjuna, N. Kesireddy and K. S. Reddy. (2011). Neuroprotective effect of

ginger on anti-oxidant enzymes in streptozotocininduced diabetic rats. Food and Chemical Toxicology. 49: 893-897.

- [53] Y. Liu, R. J. Whelan, B. R. Pattnaik, K. Ludwig, E. Subudhi, H. Rowland, N. Claussen, N. Zucker, S. Uppal, D. M. Kushner, M. Felder, M. S. Patankar and A. Kapur. (2012). Terpenoids from Zingiber officinale (Ginger) induce apoptosis in endometrial cancer cells through the activation of p53. PLoS One. 7: 53-78.
- [54] S. Khiewkhern, S. Promthet, A. Sukprasert, W. Eunhpinitpong and P. Bradshaw. (2013). Effectiveness of aromatherapy with light thai massage for cellular immunity improvement in colorectal cancer patients receiving chemotherapy. Asian Pac J C P. 14: 3903-3907.
- [55] A. Deorukhkar, N. Ahuja and A. L. Mercado. (2015). Zerumbone increases oxidative stress in a thiol-dependent ROS-independent manner to increase DNA damage and sensitize colorectal cancer cells to radiation. Cancer Medicine. 4: 278– 292.
- [56] P. A. Santos, G. B. Avanço, S. B. Nerilo, R. I. Marcelino, V. Janeiro, M. C. Valadares and M. Machinski. (2016). Assessment of cytotoxic activity of Rosemary (Rosmarinus officinalis L.), Turmeric (Curcuma longa L.), and Ginger (Zingiber officinale R.) essential oils in cervical

cancer cells (HeLa). Scientific World Journal. 10: 1155.

- [57] M. Arbabi, M. Delavari, K. Z. Fakhrieh, M. Taghizadeh and H. Hooshyar. (2016). Ginger (Zingiber officinale) induces apoptosis in Trichomonas vaginalis in vitro. International Journal of Reproductive BioMedicine. 14: 691-698.
- [58] S. Hitomi, K. Ono, K. Terawaki, C. Matsumoto, K. Mizuno, K. Yamaguchi, R. Imai, Y.Omiya, T. Hattori, Y. Kase and K. Inenaga. (2016). [6]gingerol and [6]-shogaol, active ingredients of the traditional Japanese medicine hangeshashinto, relief oral ulcerative mucositis-induced pain via action on Na+ channels. Pharmacological Research. 30: 288-302.
- [59] Y. J. Kim, Y. Jeon, T. Kim, W. C. Lim, J. Ham, Y. N. Park, T. J. Kim and H. Ko. (2016). Combined treatment with zingerone and its novel derivative synergistically inhibits TGF-β1 induced epithelial-mesenchymal transition, migration and invasion of human hepatocellular carcinoma cells. Bioorganic & Medicinal Chemistry Letters. 16: 11-17.
- [60] R. M. Hashem, L. A. Rashed, K. M. Hassanin, M. H. Hetta and A. O. Ahmed. (2017). Effect of 6gingerol on AMPK- NF-κB axis in high fat diet fed rats. Biomedicine & Pharmacotherapy. 88: 293-301.