

Historical Significance and Modern Applications of Turmeric (*Curcuma longa*) in Traditional Medicine and Beyond

Ali Zeeshan¹, Humaira Akram¹, Awon Shahzeb Nasir², Saba Manzoor³

¹Department of Biological Sciences, University of Veterinary and Animal Sciences Lahore, Punjab, Pakistan

²Department of Plant Breeding and Genetics, College of Agriculture, University of Sargodha, Punjab, Pakistan

³Department of Agriculture Economics, College of Agriculture, University of Sargodha, Punjab, Pakistan

ABSTRACT:

Ayurveda and ancient Chinese medicine have used turmeric rhizome of ginger family Zingiberaceae specie *Curcuma longa*, in the past to treat chronic illnesses like cardiovascular and metabolic disorders. Although turmeric's chemical makeup is complicated, two main groups that exhibit bioactive qualities are known as curcuminoids and lavender oil. Turmeric used in anti-inflammatory, anti-cancer, wound healing and in cosmetics etc. In cure of different biological activities turmeric used in foodstuff. Clinical trial evidence is also provided to highlight potential treatment and protective effectiveness. To fully evaluate the long-term protective effects of turmeric, clinical studies with longer intervention durations and defined endpoints for assessing health outcomes are required.

Key words: *Curcuma longa*; Antidiabetics; Antidepressant; Anti fibrinogen; Turmeric;

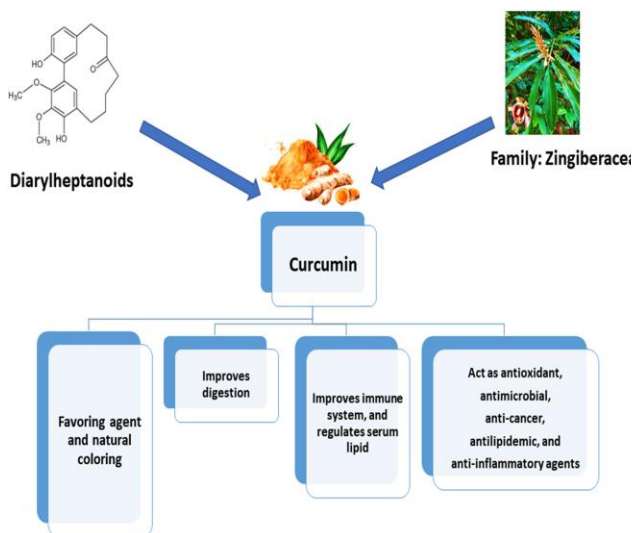
INTRODUCTION

The never ending plant *Curcuma longa* Linn. also called as *Curcuma domestica* Valetton, is a specie of the ginger family Zingiberaceae and comes from the region of India (Kocaadam et al., 2017). The epithet "longa" alludes to the long shape of plant's rhizome, from which turmeric is made. It also possesses a distinctive orange-yellow tint. The name varies depending on the language, turmeric typically refers to bright colour or the "yellow colour" Prasad S, Aggarwal BB (2011). *Curcuma* is a Latin word derived from Arabia root called *kurkum* meaning "saffron" also called Indian saffron. Mostly grows in tropics and

subtropics regions including Pakistan. The turmeric primary roots are egg and pear like while lateral roots are tubers. Ebrahim et al 2020. The tubers contain curcumin pigment main component of turmeric which acts as bioactive and polyphenolic compounds which has used to cure of different ailments (Tapal and Tiku 2012, Gilani et al). curcumin in turmeric has antioxidant property and used in food materials (Mohammadian et al 2019, Rafiee et al 2019). The chemically curcumin consists of 2 hydroxyl, methoxy and two aromatic rings with diarylheptanoid. The unsaturated aliphatic carbon chain with carbonyl groups

attached with phenolic rings (Rafiee et al 2019, Pan et al 2014)

like china, Africa and Southern sides. Nair 2013



SCIENTIFIC CLASSIFICATION OF TURMERIC (*Curcuma longa* L.)

- **Kingdom:** Plantae
- **Subkingdom:** Tracheobionts
- **Super division :** Spermatophyta
- **Division:** Mangoliophyta
- **Order** Zingiberales
- **Family** Zingiberaceae
- **Genus** Curcuma
- **Species** longa
- **Scientific name** *Curcuma longa*

MORPHOLOGY OF TURMERIC PLANT

Indonesia and Malaysia are home to the turmeric plant. it is perennial ,herbaceous and leafy ginger like plant with underground (undifferentiated root) rhizome found in America etc. it require humidity and temperature between 20 to 30 degree for proper growth or require high rain fall for flowering stage. Yellow flowers are gathered as remedies for many ailments. Since ancient times, turmeric has been cultivated in Southeast Asia with a height of 10-15 cm. The turmeric plant's thin, oval, long sheath-like leaf blades were introduced to both Africa and China

HISTORY OF TURMERIC

According to doctrine turmeric as a fundamental plant Rafiee et al 2019, Liu et al 2016Turmeric is called as ‘Indian saffron’ and has been used for 4000 years as Ayurveda in Indian medicines. Turmeric is used for flavoring with different digestive properties and highly used in India as Prasad (Nasir *et al.*, 2013). Europeans in 14th century took turmeric to western side. About 4000 years ago crushed rhizome of turmeric mostly used in different purposes like cooking, medicines and fabrics. Turmeric with 40 species have Indian origin (Nasir et al., 2013). Around about 70 to 110 species of turmeric have recorded in Asia, India, and Thailand. However, approximately 70–110 species have recorded in tropical Asia, and India other are found in Myanmar, and Thailand and some other tropical regions

CHEMICAL COMPOSITION OF TURMERIC

Analysis of turmeric shows that is a herb the herb that consists of carbohydrates, amino acids, lipids, minerals, vitamins and water 60 to 70% (RR et al., 2019) Its rhizome is a source of essential oils which

also contains terpenoids, monoterpenoids, demethoxycurcumin and curcuminoids and curcumin etc. its essential oils and curcumin classified into secondary metabolites (Nelson *et al.*, 2017).

STRUCTURE OF CURCUMIN.

Curcumin is a turmeric antioxidant and bioactive part. It has anti-inflammatory, anti-diabetics, anti-depressant, anti-fibrinogen properties. It is beneficial in treating all type of cancer (Luthra *et al.*, 2011) curcumin has potential to stop proliferations of mitogen induced of lymphocytes and blood cells that act as anti-carcinogenic activity (Huang., *et al* 1992). It has also some therapeutically values like wound healing, anti-potent, anti thrombic activites. Curcuminoids are separated from different mechanisms but solvent extraction is mostly used.



THERAPEUTICAL VALUES.....

INFLAMMATION

Organism's most important and striking defence responses to tissue damage, brought on by an ischemia injury brought on by an insufficient blood supply to a tissue physical injury, vulnerability to toxins, poison, or other types of trauma (Singh, 2019). When the inflammatory response is no longer necessary, it must be vigorously stopped to prevent tissue and cellular degeneration that could result in chronic inflammation. (Eming, Krieg, & Davidson, 2007). The inflammatory responses neutrophils, macrophages, leukocytes, cytokines and amines are included. These factors moderate the inflammatory process to cure healing and tissue damage. (Abdulkhaleq *et al.*, 2018). Different investigations have reported that inflammatory, osteoarthritis, dermatitis and bowel diseases are cured by turmeric containing curcumin (Aggarwal, Gupta, & Sung, 2013; Ahmad *et al.*, 2020; Shimizu *et al.*, 2019)

WOUND HEALING PROCESS

Wound healing are complex procedure that include reshaping of tissue and inflammation Sidhu *et al* (1998). In wounds treated with curcumin as opposed to untreated wounds, it was noticed that the transforming beta growth factor and fibronectin localization are crucial factors in wound healing, increased. Phan *et al* (2001) in an effort to understand the process of curcumin's wound healing researchers examined the effects of curcumin on damage caused by hydrogen peroxide on cultured human keratinocytes and fibroblasts. It was found that exposing human keratinocytes to curcumin (10 g/ml)

significantly reduced the damage caused by hydrogen peroxide. But there were no safeguards against harm to the oxidase of hypoxanthine- xanthine. The researchers came to the conclusion that curcumin is a powerful inhibitor of keratinocyte and fibroblast that cause damage in humans. Thangapazham et al (2007). Revealed that curcumin acts proangiogenic agent to aid in the healing of wounds by generating factor of beta growth which promotes angiogenesis during the remodelling stage of the wound healing. Panchatcharam et al (2006) found that wounds treated with curcumin healed more quickly than control wounds, as evidenced by improved rates of wound contraction and higher tensile strength.

ANTIFIBRINOGEN ACTIVITY

It was reported that Curcumin show to be a useful antifibrinogenic agent that contain collagen mRNA that prevent in Vivo and Vitro DNA synthesis. Kang et al (2002).

ANTIDIABETIC ACTIVITY

When curcumin effect were tested in rats it showed that blood sugar level decreases ad standard test revealed that lessen the oxidative stress. The result showed that curcumin reduce the level of glucose in polyol that enhanced NADPH ratio and antioxidant enzyme called glutathione peroxidase. (Arun and Nalini 2002, Hussain 2002, Murugan and Pari 2007).

RADIO PROTECTIVE ACTIVITY

A possible chemo preventive agent, curcumin reduces the effect of protein kinase that are reduced by radiations.

Therefore, it may be able to reduce the emergence of radio resistance after radiation by suppressing PKC. It inhibits the effect against toxic radiations and organo chlorine pesticides. it is beneficial for lowering cancer risk.. (Cheng et al 2001). Varadkar et al (2001) reported curcumin is mostly used in different radio resistant therapy techniques. Inano and Onada (2002) it also oticed the effect of curcuma in pregnant rats that play role in reducing the risk of mammary and different cancers Khafif et al (2005) explained that curcumin stopped the senitizing effect in different carcinoma cells. Kunwar et al (2007) it also play role as shielding cells in different enzymes like catalase, dismutase against dangerous radiations.

SAFETY DOSAGE

Different forms of turmeric like extracts, powders, with high quality has o tee associated with any toxicity. Asians who consumed turmeric per day on average did not experience any harmful side effects. (Eigner and Scholz 1999). Male and female pigs, monkeys etc were given turmeric at greater doses, however the structure and mass of the body did not alter. (Chattopadhyay et al 2004). It also reported that a experiment conducted in which 25 humans received 8000mg of turmeric on daily basis for 3 months but they show no any danger. Five further human studies utilizing daily doses of 1125 to 2500 mg of curcumin also revealed to be healthy. Curcumin's anti-inflammatory activity was based on human investigations.

ANTI TUMOR AND ANTI CANCER ACTIVITY

Huang et al., 1992) reported that ornithine decarboxylase protein that causes skin cancer in mouse is also cured by turmeric. According to Lin et al (2000) and Johnson and Mukhtar (2007) turmeric curcumin has capacity to enhance chemotherapeutic cytotoxicity that prevent from colon cancers. When curcumin was coupled with common chemotherapy drugs, synergistic benefits were seen. Due to its free radical action, it successfully inhibits UV radiation producing cells damage and decreased the prevalence cancer of skin. In vitro models, it was discovered that curcumin prevented the multiplication of breast cancer cells. (Zhang et al 2007). Pure turmeric reduced the growth of malignant and premalignant reducing in the glandular stomach and had anti - carcinogenic actions in rats. (Ozaki et al (2000) kappa B nuclear factor and activated protein I transcriptional factors were shown to be strongly reduced by curcumin. These elements were understood to perform crucial functional roles in osteoclast preservation. It has been demonstrated that curcumin induces osteoclast mortality in a dose- or manner of moment. As evidence that it increases osteoclast apoptosis, it displays osteoplastic and osteoclast activity. Wahl et al (2007) revealed that curcumin might overcome the chemo resistant phenotype seen in many cells of by apoptotic ways.

TURMERIC IN COSMETICS

Turmeric is used in different cosmetics used for skin as colorant agent. (Nair, 2013). As a skin beautifier, it is frequently combined

with milk and applied externally and internally by Indian. In different Asians countries turmeric extract is used on bride and groom skin before the wedding ceremony in the hopes that it will make the skin sparkle and ward off hazardous bacteria. (Sabale, Modi & Sabale, 2013). Numerous multinational enterprises and worldwide businesses already employ it in the manufacture of several sunscreens and turmeric-based face treatments. (Prasad & Aggarwaal, 2011).Hindu ladies utilise turmeric and turmeric-based products like kumkum and parani in their baths as an inexpensive and all-natural method of improving their appearance. It has been said that applying turmeric paste to the face and limbs before a bath will wash the skin and improve the appearance of the face. Its mending and antibacterial properties are regarded to be a preventive and therapeutic treatment for the terrible adolescent ailment known as pimples. (Sabale, Modi & Sabale, 2013). It is also source of removing unwated hair from female skin (Ratanshi, 2017; Sa & Das, 2008; Sabale, Modi & Sabale, 2013). After giving birth or after pregnancy woman consume turmeric paste including ginger and milk that help in recovery soon, (Nair, 2013; Krishnaswamy. 2008)

ANTI DEPRESSED

The American Cancer Society (2016) reported curcumin is effective in controlling depression symptoms in animals despite the fact that few clinical trials have been performed Prasad and Aggarwal (2011), turmeric can effective in depression. They postulated that swim stress resulted in a considerable reduction in the ethanolic

extract's serotonin, acid, noradrenaline, and dopamine concentrations as well as serotonin turnover. In a study published Chen (2002) reported that mice administrating turmeric paste about 140 to 150 mg for 14 days show reduced stiffness in tail dispersion and forced swim tests. 560 mg of turmeric have antidepressant activity while doses reduced brain oxidase activity while large amount have an impact on brain activities while in rats fluoxetine reduced the effect of MAO-A in brain cells. Curcumin is not soluble in water, hence it is uncertain what molecule in turmeric aqueous preparations is responsible for its activity.

CONCLUSION

Since ancient times, the medical systems of Ayurveda, Unani, and Siddha have all utilized turmeric. A review of the literature reveals that turmeric, whether in the various form such as extracts, powder or isolated components, shows a wide range of therapeutically actions with few effects. The phenyl ring with methoxy group the phenolic system, and the 1, 3-diketone system all play significant roles in the pharmacological actions of curcumin. Fortified with curcumin or turmeric, a number of useful varieties have been introduced in national and international markets for a variety of ailments. Even though this plant has been the subject of extensive research, there is still room for improvement in the area of medication development. In comparison to other phyto-antioxidants, curcumin is a safe, non-poisonous and potent natural antioxidant. In the near future, curcuma longa could be used as a cutting-edge herbal medication to treat a

variety of illnesses, such as carcinogenesis, diseases caused by inflammation and oxidative stress pathogenesis. Additional assessments must be conducted on Curcuma longa in order to learn about its numerous other uses for medication.

REFERENCES:

1. 1 Ebrahim AA, Elnesr SS, Abdel-Mageed MA and Aly MM, Nutritional significance of Aloe vera (*Aloe barbadensis* Miller) and its beneficial impact on poultry. *Worlds Poult Sci J* 76:803–814 (2020). <https://doi.org/10.1080/00439339.2020.1830010>
2. 10 Liu W, Chen XD, Cheng Z and Selomulya C, On enhancing the solubility of curcumin by microencapsulation in whey protein isolate via spray drying. *J Food Eng* 169:189–195 (2016). <https://doi.org/10.1016/j.jfoodeng.2015.08.034>
3. 7 Mohammadian M, Salami M, Momen S, Alavi F, Emam-Djomeh Z and Moosavi-Movahedi AA, Enhancing the aqueous solubility of curcumin at acidic condition through the complexation with whey protein nanofibrils. *Food Hydrocoll* 87:902–914 (2019). <https://doi.org/10.1016/j.foodhyd.2018.09.001>
4. 8 Rafiee Z, Nejatian M, Daeihamed M and Jafari SM, Application of curcumin-loaded nanocarriers for food, drug and cosmetic purposes. *Trends Food Sci Technol* 88:445–

- 458 (2019). <https://doi.org/10.1016/j.tifs.2019.04.017>
5. Aggarwal BB, Kumar A, Bharti AC 2003. Anticancer potential of curcumin: preclinical and clinical studies. *Anticancer Res* 23:363-398
 6. Arun N, Nalini N 2002. Efficacy of turmeric on blood sugar and polyol pathway in diabetic albino rats. *Plant Food Hum Nutr* 57:41-52
 7. Awasthi S 1996. Curcumin protects against 4-hydroxy-2-transnonenal-induced cataract formation in rat lenses. *Am J Clin Nutr* 64:761-766
 8. Azuine MA, Bhide SV 1994. Adjuvant chemoprevention of experimental cancer: catechin and dietary turmeric in forestomach and oral cancer models. *J Ethnopharmacol* 44:211-217
 9. Barquero LC, Villegas I, Sanchez-Calvo JM, Talero E, Sanchez-Fidalgo S, Motilva V, Lastra CA 2007. Curcumin, a *Curcuma longa* constituent, acts on MAPK p38 pathway modulating COX-2 and iNOS expression in chronic experimental colitis. *Int Immunopharm* 7:333-342
 10. Chattopadhyay I, Biawas K, Bandyopadhyay I, Banerjee RK 2004. Turmeric and curcumin: Biological actions and medicinal applications. *Curr Sci* 87:44-53
 11. Cheng A-L, Hsu C-H, Lin JK 2001. Phase I clinical trial of curcumin, a chemopreventive agent in patients with high risk or premalignant lesions. *Anticancer Res* 27:2895-2900
 12. Commandeur JN, Vermeulen NP 1996. Cytotoxicity and cytoprotective activities of natural compounds. The case of curcumin. *Xenobiotics* 26:667-680
 13. Cousins M, Adelberg J, Chenb F, Rieck J 2007. Antioxidant capacity of fresh and dried rhizomes from four clones of turmeric (*Curcuma longa* L.) grown in vitro. *Ind Crop Prod* 25:129-135
 14. Eigner D, Scholz D 1999. *Ferula asafoetida* and *Curcuma longa* in traditional medical treatment and diet in Nepal. *J Ethnopharmacol* 67:1-6
 15. Gilani N, Basharat H and Qureshi H, Curcumin—a review on multipotential phytochemical. *J Coastal Life Med* 5:455-458 (2017). <https://doi.org/10.12980/jclm.5.2017j7-115>.
 16. Huang HC, Jan TR, Yeh SF 1992. Inhibitory effect of curcumin, an anti-inflammatory agent, on vascular smooth muscle cell proliferation. *Eur J Pharmacol* 221:381-384
 17. Huang MT, Lysz T, Ferraro T, Abidi TF, Laskin JD, Conney AH 1991. Inhibitory effects of curcumin on in vitro lipoxygenase and cyclooxygenase activities in mouse epidermis. *Cancer Res* 51:813-819
 18. Huang MT, Smart RC, Wong CQ, Conney AH 1988. Inhibitory effect of curcumin, chlorogenic acid, caffeic acid and ferulic acid on tumor promotion in mouse skin by 12-O-tetradecanoylphorbol-13-acetate. *Cancer Res* 48:5941-5946

19. Hussain HEM 2002. Hypoglycemic, hypolipidemic and antioxidant properties of combination of curcumin from *Curcuma longa* Linn and partially purified product from *Abroma augusta* Linn in streptozotocin induced diabetes. *Indian J Clin Biochem* 17:33-43
20. Inano H, Onoda M 2002. Prevention of radiation-induced mammary tumors. *Int J Radiat Oncol Biol Phys* 52:212-223
21. Jang HD, Chang KS, Huang YS, Hsu CL, Lee SH, Su MS 2007. Principal phenolic phytochemicals and antioxidant activities of three Chinese medicinal plants. *Food Chem* 103:749-756
22. Jaruga E, Bielak-Zmijewska A, Sikora E, Skierski J, Radziszewska E, Pioweka K, Bartoz G 1998. Glutathione-independent mechanism of apoptosis inhibited by curcumin in rat thymocytes. *Biochem Pharmacol* 56:961-965
23. Jayaprakasha GK, Rao LJ, Sakariah KK 2006. Antioxidant activities of curcumin, demethoxycurcumin and bisdemethoxy curcumin. *Food Chem* 98:720-724
24. Johnson JJ, Mukhtar H 2007. Curcumin for chemoprevention of colon cancer. *Can Lett* 255:170-181
25. Kang HC, Nan JX, Park PH 2002. Curcumin inhibits collagen synthesis and hepatic stellate cell activation in vivo and in vitro. *J Pharm Pharmacol* 54:119-126
26. Khafif A, Hurst R, Kyker K, Fliss DM, Gil Z, Medina JE 2005. Curcumin: a new radio-sensitizer of squamous cell carcinoma cells. *Otolaryngol Head Neck Surg* 132:317-321
27. Krishnaswamy, K. (2008). Traditional Indian spices and their health significance. *Asia Pac J Clin Nutr.*, 17(1), 265-268.
28. Kunwar A, Narang H, Priyadarsini KI, Krishna M, Pandey R, Sainis KB 2007. Delayed activation of PKCdelta and NFkappa B and higher radioprotection in splenic lymphocytes by copper (II)-curcumin (1:1) complex as compared to curcumin. *J Cell Biochem* 102:1214-1224
29. Kuo ML, Huang TS, Lin JK 1996. Curcumin an antioxidant and anti-tumor promoter, induces apoptosis in human leukemia cells. *Biochem Biophys Acta* 15:95-100
30. Kurien BT, Scofield RH 2007. Curcumin/turmeric solubilized in sodium hydroxide inhibits HNE protein modification-An in vitro study. *J Ethnopharmacol* 110:368-373
31. Lal B, Kapoor AK, Agarwal PK, Asthana OP, Srimal RC 2000. Role of curcumin in idiopathic inflammatory orbital pseudotumours. *Phytother Res* 14:443-447
32. Lee SL, Huang WJ, Lin WW, Lee SS and Chen CH 2005. Preparation and anti-inflammatory activities of diarylheptanoid and diarylheptylamine analogs.

- Bioorganic Med Chem 13:6175-6181
33. Lin JK, Pan MH, Lin-Shiau SY 2000. Recent studies on the biofunctions and biotransformations of curcumin. *Biofactors* 13:153-158
34. Masuda T, Hidaka K, Shinohara A, Maikawa T, Takeda Y, Yamaguchi H 1999. Chemical studies on antioxidant mechanism of curcuminoid: analysis of radical reaction products from curcumin. *J Agric Food Chem* 47:71-77
35. Mukhophadhyay A, Basu N, Ghatak N, Gujral PK 1982. Anti-inflammatory and irritant activities of curcumin analogues in rats. *Agents Actions* 12:508-515
36. Murugan P, Pari L 2007. Influence of tetrahydrocurcumin on erythrocyte membrane bound enzymes and antioxidant status in experimental type 2 diabetic rats. *J Ethnopharmacol* 113:479-486
37. Nair KPP, *The Agronomy and Economy of Turmeric and Ginger the Invaluable Medicinal Spice Crops*. Elsevier, Amsterdam, pp. 1–544 (2013)
38. Nair, K. P. (2013). *The agronomy and economy of turmeric and ginger: the invaluable medicinal spice crops*. Newnes.
39. Ozaki K, Kawata Y, Amano S, Hanazawa S 2000. Stimulatory effect of curcumin on osteoclast apoptosis. *Biochem Pharmacol* 59:1577-1581
40. Pan K, Luo Y, Gan Y, Baek SJ and Zhong Q, pH-driven encapsulation of curcumin in self-assembled casein nanoparticles for enhanced dispersibility and bioactivity. *Soft Matter* 10:6820–6830 (2014). <https://doi.org/10.1039/c4sm00239c>.
41. Panchatcharam M, Miriyala S, Gayathri VS, Suguna L 2006. Curcumin improves wound healing by modulating collagen and decreasing reactive oxygen species. *Mol Cell Biochem* 290:87-96
42. Panchatcharam M, Miriyala S, Gayathri VS, Suguna L 2006. Curcumin improves wound healing by modulating collagen and decreasing reactive oxygen species. *Mol Cell Biochem* 290:87-96
43. Phan TT, See P, Lee ST, Chan SY 2001. Protective effects of curcumin against oxidative damage on skin cells in vitro: its implication for wound healing. *J Trauma* 51:927-931
44. Phan TT, See P, Lee ST, Chan SY 2001. Protective effects of curcumin against oxidative damage on skin cells in vitro: its implication for wound healing. *J Trauma* 51:927-931
45. Prasad, S. & Aggarwal B.B. (2011). *Turmeric, the Golden Spice: From Traditional Medicine to Modern Medicine*. In Benzie I.F.F, Wachtel-Galor S, (eds) *Herbal Medicine: Biomolecular and Clinical Aspects*. 2nd edition. Boca Raton (FL): CRC Press/Taylor & Francis. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK92752/> • Rahimi, H. R., Nedaeinia, R., Sepehri, S. A

46. Ramsewak RS, De Witt DL, Nair MG 2000. Cytotoxicity, antioxidant and anti-inflammatory activities of curcumins I-III from *Curcuma longa*. *Phytomed* 7:303-308
47. Ratanshi, S. (2017). Application of Turmeric. Retrieved from <http://www.shahrkturmeric.com/turmeric/properties-a-applications>
48. Sa, G., & Das, T. (2008). Anti-cancer effects of curcumin: cycle of life and death. *Cell division*, 3(1), 14.
49. Sabale, P., Modi, P., & Sabale, V. (2013). *Curcuma longa* Linn. A Phytochemical and Phytopharmacological Review. *Research Journal of Pharmacognosy and Phytochemistry*, 5(2), 59- 68.
50. Saeed M, Arain MA, Ali Fazlani S, Marghazani IB, Umar M, Soomro J et al., A comprehensive review on the health benefits and nutritional significance of fucoidan polysaccharide derived from brown seaweeds in human, animals and aquatic organisms. *Aquacult Nutr* 27:633–654 (2021). <https://doi.org/10.1111/anu.13233>
51. Selvem R, Subramaniam L, Gayathri R, Angayarkanni N 1995. The antioxidant activity of turmeric (*Curcuma longa*). *J Ethnopharmacol* 47:59-67
52. Sidhu GS, Singh AK, Thaloor D, Banaudha KK, Patniak GK, Srimal RC, Maheshwari RK 1998. Enhancement of wound healing by curcumin in animals. *Wound Repair Reg* 6:167-177
53. Sidhu GS, Singh AK, Thaloor D, Banaudha KK, Patniak GK, Srimal RC, Maheshwari RK 1998. Enhancement of wound healing by curcumin in animals. *Wound Repair Reg* 6:167-177
54. Soudamini, Kuttan R 1988. Cytotoxic and tumor reducing properties of curcumin. *Indian J Pharmacol* 20:95-101
55. Sreejayan N, Rao MN 1994. Curcuminoids as potent inhibitors of lipid peroxidation. *J Pharm Pharmacol* 46:1013-1016
56. Sreejayan N, Rao MN 1996. Free radical scavenging activity of curcuminoids. *Arzneimittelforschung* 46:169-171
57. Sreejayan N, Rao MN 1997. Nitric acid oxide scavenging by curcuminoids. *J Pharmacol* 49:105-107
58. Stano J, Grancai D, Neubert K, Kresanek J 2000. Curcumin as a potential antioxidant. *Ceska Slov Farm* 49:168-170
59. Tapal A and Tiku PK, Complexation of curcumin with soy protein isolate and its implications on solubility and stability of curcumin. *Food Chem* 130:960–965 (2012). <https://doi.org/10.1016/j.foodchem.2011.08.025>.
60. Thangapazham RL, Sharma A, Maheshwari RK 2007. Beneficial role of curcumin in skin diseases. *Adv Exp Med Biol* 595:343- 357
61. Thangapazham RL, Sharma A, Maheshwari RK 2007. Beneficial

- role of curcumin in skin diseases. *Adv Exp Med Biol* 595:343- 357
62. Thresiamma KC 1996. Protective effect of curcumin, ellagic acid and bixin on radiation induced toxicity. *Indian J Exp Biol* 34:845-847
63. Unnikrishnan MK, Rao MN 1995. Inhibition of nitrite induced oxidation of hemoglobin by curcuminoids. *Pharmazie* 50:490-492
64. Varadkar P, Dubey P, Krishna M, Verma NC 2001. Modulation of radiation induced protein kinase and activity by phenolics. *J Radio Prot* 21:261-370
65. Venkatesan P, Rao MN 2000. Structure activity relationships for the inhibition of lipid peroxidation and the scavenging of free radicals by synthetic symmetric curcumin analogues. *J Pharm Pharmacol* 52:1123-1128
66. Wahl H, Tan L, Griffith K, Choi M, Liu JR 2007. Curcumin enhances Apo2L/TRAIL induced apoptosis in chemoresistant ovarian cancer cells *Gynecol Oncol* 105:104–112
67. Woo HM, Kang JH, Kawada T, Yoo H, Sung MK, Yu R 2007. Active spice-derived components can inhibit inflammatory responses of adipose tissue in obesity by suppressing inflammatory actions of macrophages and release of monocyte chemoattractant protein-1 from adipocytes. *Life Sci* 80:926–931
68. Wu NC 2003. Safety and anti-inflammatory activity of curcumin: A component of turmeric (*Curcuma longa*). *J Alter Comp Med* 9:161-168
69. Xu PH, Long Y, Dai F, Liu ZL 2007. The relaxant effect of curcumin on porcine coronary arterial ring segments. *Vasc Pharmacol* 47:25–30
70. Xu YX, Pindolia KR, Janakiraman N, Noth CJ, Chapman RA, Gautam SC 1997. Curcumin, a compound with anti-inflammatory and antioxidant properties, downregulates Chemokine expressions in bone marrow stromal cells. *Exp Hematol* 25:413-422
71. Yamamoto H, Mizutani T, Nomura H 1997. Inhibitory effects of curcumin on mammalian phospholipase D activity. *FEBS Lett* 417:196-198
72. Yu, Z. F., Kong, L. D., & Chen, Y. (2002). Antidepressant activity of aqueous extracts of *Curcuma longa* in mice. *J Ethnopharmacol.*, 83, 161–5.
73. Zhang HG, Kim H, Liu C, Yu S, Wang J, Grizzle WE, Kimberly RP, Barnes S 2007. Curcumin reverses breast tumor exosomes mediated immune suppression of NK cell tumor cytotoxicity. *Biochim Biophys Acta* 1773:1116–1123