

## CURCUMA LONGA AND CURCUMIN: A REVIEW ARTICLE

M. AKRAM<sup>1</sup>, SHAHAB-UDDIN<sup>1</sup>, AFZAL AHMED<sup>2</sup>, KHAN USMANGHANI<sup>3</sup>,  
ABDUL HANNAN<sup>3</sup>, E. MOHIUDDIN<sup>4</sup>, M. ASIF<sup>5</sup>

Turmeric is a spice derived from the rhizomes of *Curcuma longa*, which is a member of the ginger family (*Zingiberaceae*). Rhizomes are horizontal underground stems that send out shoots as well as roots. The bright yellow color of turmeric comes mainly from fat-soluble, polyphenolic pigments known as curcuminoids. Curcumin, the principal curcuminoid found in turmeric, is generally considered its most active constituent. Other curcuminoids found in turmeric include demethoxycurcumin and bisdemethoxycurcumin. In addition to its use as a spice and pigment, turmeric has been used in India for medicinal purposes for centuries. More recently, evidence that curcumin may have anti-inflammatory and anticancer activities has renewed scientific interest in its potential to prevent and treat the disease.

*Key words:* Anti-Inflammatory, Anti-H. pylori, curcumin.

Curcumin is the principal curcuminoid of the popular Indian spice turmeric, which is a member of the ginger family (*Zingiberaceae*). The other two curcuminoids are desmethoxycurcumin and bis-desmethoxycurcumin. The curcuminoids are polyphenols and are responsible for the yellow color of turmeric. Curcumin can exist in at least two tautomeric forms, keto and enol. The enol form is more energetically stable in the solid phase and in solution. Curcumin can be used for boron quantification in the so-called curcumin method. It reacts with boric acid forming a red colored compound, known as rosocyanine. Curcumin is brightly yellow colored and may be used as a food coloring. As a food additive, its E number is E100.

**Properties of Curcumin.** Curcumin has antioxidant, anti-inflammatory, antiviral and antifungal actions. Studies have shown that curcumin is not toxic to humans. Curcumin exerts anti-inflammatory activity by inhibition of a number of different molecules that play an important role in inflammation. Turmeric is effective in reducing post-surgical inflammation. Turmeric helps to prevent atherosclerosis by reducing the formation of blood clumps. Curcumin inhibits the

---

<sup>1</sup> Hamdard University, Faculty of Eastern Medicine, Department of Basic Medical Sciences, Pakistan, Corresponding author address Dr. Muhammad Akram E-mail: makram\_0451@hotmail.com.

<sup>2</sup> Hamdard University, Faculty of Eastern Medicine, Department of Medicine and Allied Sciences, Pakistan.

<sup>3</sup> Hamdard University, Faculty of Eastern Medicine, Department of Preclinical Sciences, Pakistan.

<sup>4</sup> Hamdard University, Faculty of Eastern Medicine, Department of Preclinical Sciences, Pakistan.

<sup>5</sup> Islamia University Bahawalpur, Department of Conventional Medicine, Pakistan.

growth of *Helicobacter pylori*, which causes gastric ulcers and has been linked with gastric cancers. Curcumin can bind with heavy metals such as cadmium and lead, thereby reducing the toxicity of these heavy metals. This property of curcumin explains its protective action to the brain. Curcumin acts as an inhibitor for cyclooxygenase, 5-lipoxygenase and glutathione S-transferase. It is a common spice, known mostly for its use in Indian dishes as a common ingredient in curries and other ethnic meals. Turmeric has also been used for centuries in Ayurvedic medicine, which integrates the medicinal properties of herbs with food. This extraordinary herb has found its way into the spotlight in the west because of its wide range of medicinal benefits. Turmeric is a potent antioxidant.

Curcumin, its main active constituent, is as powerful and antioxidant as vitamins C, E and Beta-Carotene, making turmeric usage a consumer choice for cancer prevention, liver protection and premature aging. Several published studies also show that turmeric inhibits the growth of several different types of cancer cells. In addition, turmeric is a powerful anti-inflammatory, easing conditions such as bursitis, arthritis and back pain. Turmeric's anti-inflammatory action is likely due to a combination of three different properties.

First, turmeric lowers the production of inflammation-inducing histamine. Secondly, it increases and prolongs the action of the body's natural anti-inflammatory adrenal hormone, cortisol, and finally, turmeric improves circulation, thereby flushing toxins out of small joints where cellular wastes and inflammatory compounds are frequently trapped. Research has also confirmed the digestive benefits of turmeric. Turmeric acts as a cholagogue, stimulating bile production, thus, increasing the bodies' ability to digest fats, improving digestion and eliminating toxins from the liver.

**Active Constituents.** The active constituents of turmeric are the flavonoid curcumin (diferuloylmethane) and various volatile oils, including tumerone, atlantone, and zingiberone. Other constituents include sugars, proteins, and resins. The best-researched active constituent is curcumin, which comprises 0.3–5.4 percent of raw turmeric.

**Pharmacokinetics.** Pharmacokinetic studies in animals have demonstrated that 40-85 percent of an oral dose of curcumin passes through the gastrointestinal tract unchanged, with most of the absorbed flavonoid being metabolized in the intestinal mucosa and liver. Due to its low rate of absorption, curcumin is often formulated with bromelain for increased absorption and enhanced anti-inflammatory effect.

#### MECHANISMS OF ACTION

**Antioxidant Effects.** Water- and fat-soluble extracts of turmeric and its curcumin component exhibit strong antioxidant activity, comparable to vitamins C

and E. A study of ischemia in the feline heart demonstrated that curcumin pretreatment decreased ischemia-induced changes in the heart. An *in vitro* study measuring the effect of curcumin on endothelial heme oxygenase-1, an inducible stress protein, was conducted utilizing bovine aortic endothelial cells. Incubation (18 hours) with curcumin resulted in enhanced cellular resistance to oxidative damage.

**Hepatoprotective Effects.** Turmeric has been found to have a hepatoprotective characteristic similar to silymarin. Animal studies have demonstrated turmeric's hepatoprotective effects from a variety of hepatotoxic insults, including carbon tetrachloride (CCl<sub>4</sub>), galactosamine, acetaminophen (paracetamol), and Aspergillus aflatoxin. Turmeric's hepatoprotective effect is mainly a result of its antioxidant properties, as well as its ability to decrease the formation of pro-inflammatory cytokines. In rats with CCl<sub>4</sub>-induced acute and subacute liver injury, curcumin administration significantly decreased liver injury in test animals compared to controls. Turmeric extract inhibited fungal aflatoxin production by 90 percent when given to ducklings infected with Aspergillus parasiticus. Turmeric and curcumin also reversed biliary hyperplasia, fatty changes, and necrosis induced by aflatoxin production. Sodium curcumin, a salt of curcumin, also exerts choleric effects by increasing biliary excretion of bile salts, cholesterol, and bilirubin, as well as increasing bile solubility, therefore possibly preventing and treating cholelithiasis.

**Anti-inflammatory Effects.** The volatile oils and curcumin of *Curcuma longa* exhibit potent anti-inflammatory effects. Oral administration of curcumin in instances of acute inflammation was found to be as effective as cortisone or phenylbutazone, and one-half as effective in cases of chronic inflammation. In rats with Freund's adjuvant-induced arthritis, oral administration of *Curcuma longa* significantly reduced inflammatory swelling compared to controls. In monkeys, curcumin inhibited neutrophil aggregation associated with inflammation. *C. longa*'s anti-inflammatory properties may be attributed to its ability to inhibit both biosynthesis of inflammatory prostaglandins from arachidonic acid, and neutrophil function during inflammatory states. Curcumin may also be applied topically to counteract inflammation and irritation associated with inflammatory skin conditions and allergies, although care must be used to prevent staining of clothing from the yellow pigment.

**Anticarcinogenic Effects.** Animal studies involving rats and mice, as well as *in vitro* studies utilizing human cell lines, have demonstrated curcumin's ability to inhibit carcinogenesis at three stages: tumor promotion, angiogenesis, and tumor growth. In two studies of colon and prostate cancer, curcumin inhibited cell proliferation and tumor growth. Turmeric and curcumin are also capable of suppressing the activity of several common mutagens and carcinogens in a variety of cell types in both *in vitro* and *in vivo* studies. The anticarcinogenic effects of turmeric and curcumin are due to direct antioxidant and free-radical scavenging

effects, as well as their ability to indirectly increase glutathione levels, thereby aiding in hepatic detoxification of mutagens and carcinogens, and inhibiting nitrosamine formation.

**Antimicrobial Effects.** Turmeric extract and the essential oil of *Curcuma longa* inhibit the growth of a variety of bacteria, parasites, and pathogenic fungi. A study of chicks infected with the caecal parasite *Eimeria maxima* demonstrated that diets supplemented with 1-percent turmeric resulted in a reduction in small intestinal lesion scores and improved weight gain. Another animal study, in which guinea pigs were infected with either dermatophytes, pathogenic molds, or yeast, found that topically applied turmeric oil inhibited dermatophytes and pathogenic fungi, but neither curcumin nor turmeric oil affected the yeast isolates. Improvements in lesions were observed in the dermatophyte- and fungi-infected guinea pigs, and at seven days post-turmeric application the lesions disappeared. Curcumin has also been found to have moderate activity against *Plasmodium falciparum* and *Leishmania major* organisms.

**Cardiovascular Effects** Turmeric's protective effects on the cardiovascular system include lowering cholesterol and triglyceride levels, decreasing susceptibility of low density lipoprotein (LDL) to lipid peroxidation, and inhibiting platelet aggregation. These effects have been noted even with low doses of turmeric. A study of 18 atherosclerotic rabbits given low-dose (1.6–3.2 mg/kg body weight daily) turmeric extract demonstrated decreased susceptibility of LDL to lipid peroxidation, in addition to lower plasma cholesterol and triglyceride levels. The higher dose did not decrease lipid peroxidation of LDL, but cholesterol and triglyceride level decreases were noted, although to a lesser degree than with the lower dose. Turmeric extract's effect on cholesterol levels may be due to decreased cholesterol uptake in the intestines and increased conversion of cholesterol to bile acids in the liver. Inhibition of platelet aggregation by *C. longa* constituents is thought to be *via* potentiation of prostacyclin synthesis and inhibition of thromboxane synthesis.

**Gastrointestinal Effects.** Constituents of *Curcuma longa* exert several protective effects on the gastrointestinal tract. Sodium curcumin ate inhibited intestinal spasm and p-tolymethylcarbinol, a turmeric component, increased gastrin, secretin, bicarbonate, and pancreatic enzyme secretion. Turmeric has also been shown to inhibit ulcer formation caused by stress, alcohol, indomethacin, pyloric ligation, and reserpine, significantly increasing gastric wall mucus in rats subjected to these gastrointestinal insults.

**Curcumin enhances immunity.** Curcumin can also help the body fight off cancer should some cells escape apoptosis. When researchers looked at the lining of the intestine after ingestion of curcumin, they found that CD4+ T-helper and B type immune cells were greater in number. In addition to this localized immune stimulation, curcumin also enhances immunity in general. Researchers in India

have documented increased antibodies and more immune action in mice given curcumin.

**Curcumin blocks NF- $\kappa$ B and the motogenic response in *Helicobacter pylori*-infected epithelial cells** Studies indicate that infection of epithelial cells by the microbial pathogen *Helicobacter pylori* leads to activation of the transcription factor nuclear factor  $\kappa$ B (NF- $\kappa$ B), the induction of pro-inflammatory cytokine/chemokine genes, and the motogenic response (cell scattering). It has been investigated that *H. pylori*-induced NF- $\kappa$ B activation and the subsequent release of interleukin 8 (IL-8) are inhibited by curcumin (diferuloylmethane), a yellow pigment in turmeric (*Curcuma longa* L.). It has been demonstrated that curcumin inhibits I $\kappa$ B $\alpha$  degradation, the activity of I $\kappa$ B kinases  $\alpha$  and  $\beta$  (IKK $\alpha$  and  $\beta$ ), and NF- $\kappa$ B DNA-binding. The mitogen-activated protein kinases (MAPK), extracellular signal-regulated kinases 1/2 (ERK1/2) and p38, which are also activated by *H. pylori* infection, are not inhibited by curcumin. It is studied that *H. pylori*-induced motogenic response is blocked by curcumin. It has been concluded that curcumin, due to inhibition of NF- $\kappa$ B activation and cell scattering, should be considered as a potential therapeutic agent effective against pathogenic processes initiated by *H. pylori* infection.

**Pregnancy and Lactation.** Although there is no evidence that dietary consumption of turmeric as a spice adversely affects pregnancy or lactation, the safety of curcumin supplements in pregnancy and lactation has not been established.

#### REFERENCES

1. Bush J.A., Cheung K.J. Jr, Li G., 2001, Curcumin induces apoptosis in human melanoma cells through a Fas receptor/caspase-8 pathway independent of p53. *Exp Cell Res*, **271**: 305-314.
2. Cheng A.L., Hsu C.H., Lin J.K., 2001, Phase I clinical trial of curcumin, a chemopreventive agent, in patients with high-risk or pre-malignant lesions. *Anticancer Res*, **21**: 2895-2900.
3. Hanif R., Qiao L., Shiff S.J., Rigas B., 1997, Curcumin, a natural plant phenolic food additive, inhibits cell proliferation and induces cell cycle changes in colon adenocarcinoma cell lines by a prostaglandin-independent pathway. *J Lab Clin Med*, **130**: 576-584.
4. Hour T.C., Chen J., Huang C.Y., 2002, Curcumin enhances cytotoxicity of chemotherapeutic agents in prostate cancer cells by inducing p21 (WAF1/CIP1) and C/EBP $\beta$  expressions and suppressing NF- $\kappa$ B activation. *Prostate*, **51**: 211-218.
5. Kawamori T., Lubet R., Steele V.E., 1999, Chemopreventative effect of curcumin, a naturally occurring anti-inflammatory agent, during the promotion/progression stages of colon cancer. *Cancer Res*, **59**: 597-601.
6. Lal B., Kapoor A.K., Asthana O.P., 1999, Efficacy of curcumin in the management of chronic anterior uveitis. *Phytother Res*, **13**: 318-322.
7. Limtrakul P., Lipigorngoson S., Namwong O., 1997, Inhibitory effect of dietary curcumin on skin carcinogenesis in mice. *Cancer Lett*, **116**: 197-203.
8. Mehta R.G., Moon R.C., 1991, Characterization of effective chemopreventive agents in mammary gland *in vitro* using an initiation-promotion protocol. *Anticancer Res*, **11**: 593-596.
9. Mukhopadhyay A., Basu N., Ghatak N., 1982, Anti-inflammatory and irritant activities of curcumin analogues in rats. *Agents Actions*, **12**: 508-515.

10. Munzenmaier A., *et al.*, 1997, A secreted/shed product of *Helicobacter pylori* activates transcription factor nuclear factor-kappa B. *J Immunol*, **159**: 6140-47.
11. Park EJ, *et al.*, 2000, Protective effect of curcumin in rat liver injury induced by carbon tetrachloride. *J Pharm Pharmacol*, **52**: 437-40.
12. Pendurthi U.R., *et al.*, 1997, Inhibition of tissue factor gene activation in cultured endothelial cells by curcumin. Suppression of activation of transcription factors Egr-1, AP-1, and NF-kappa B. *Arterioscler Thromb Vasc Biol.*, **17**: 3406-13.
13. Ramachandran C., Fonseca H.B., Jhabvala P., *et al.*, 2002, Curcumin inhibits telomerase activity through human telomerase reverse transcriptase in MCF-7 breast cancer cell line. *Cancer Lett*, **184**: 1-6.
14. Reddy B.S., Rao C.V., 2002, Novel approaches for colon cancer prevention by cyclooxygenase-2 inhibitors. *J Environ Pathol Toxicol Oncol*, **21**: 155-164.
15. Shao Z.M., Shen Z.Z., Liu C.H., *et al.*, 2002, Curcumin exerts multiple suppressive effects on human breast carcinoma cells., *Int J Cancer*, **98**: 234-240.
16. Sharma A., *et al.*, 2000, Spice extracts as dose-modifying factors in radiation inactivation of bacteria. *J Agric Food Chem*, **48**: 1340-44.
17. Simon A, *et al.*, 1998. Inhibitory effect of curcuminoids on MCF-7 cell proliferation and structure-activity relationships. *Cancer Lett*, **129**:111-16.
18. Somasundaram S., Edmund N.A., Moore D.T., *et al.*, 2002, Dietary curcumin inhibits chemotherapy-induced apoptosis in models of human breast cancer. *Cancer Res*, **62**: 3868-3875.
19. Sreejayan, N. & Rao, M.N., 1994, Curcuminoids as potent inhibitors of lipid peroxidation. *J. Pharm.Pharmacol*, **46**: 1013.
20. Sreejayan, N., & Rao, M.N. 1996, Free radical scavenging activity of curcuminoids. *Arzneimittelforschung*, **46** (2): 169-171.
21. Srivastava R., Puri V., Srimal R.C., Dhawan B.N. 1986, Effect of curcumin on platelet aggregation and vascular prostacyclin synthesis. *Arzneimittelforschung*, **36**: 715-717.
22. Thaloor D., Singh A.K., Sidhu G.S., *et al.*, 1998, Inhibition of angiogenic differentiation of human umbilical vein endothelial cells by curcumin. *Cell Growth Differ*, **9**: 305-312.
23. Venkatesan N., 2000, Pulmonary protective effects of curcumin against paraquat toxicity. *Life Sci* **66**(2):PL21-28.
24. Verma S.P., *et al.*, 1997, Curcumin and genistein, plant natural products, show synergistic inhibitory effects on the growth of human breast cancer MCF-7 cells induced by estrogenic pesticides. *Biochem Biophys Res Comm*, **233**: 692-96.
25. Verma S.P., *et al.*, 1998, The inhibition of the estrogenic effects of pesticides and environmental chemicals by curcumin and isoflavonoids. *Environ Health Perspect*, **106**: 807-812.