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## **HerbClip**<sup>TM</sup>

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File: ■ Turmeric (*Curcuma longa*)
■ Curcumin
■ Potential Therapeutic Uses

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**RE: Curcumin Exhibits Potential in the Treatment of Various Chronic Diseases** 

Gupta SC, Patchva S, Koh W, Aggarwal BB. Discovery of curcumin, a component of golden spice, and its miraculous biological activities. *Clin Exp Pharmacol Physiol*. 2012 Mar;39(3):283-299.

The medicinal properties of turmeric (*Curcuma longa*), the source of curcumin, have been known for thousands of years, but advancements in modern science have provided a scientific basis for using curcumin therapy against numerous human diseases. In this article, the authors discuss the discovery and key biological activities of curcumin, particularly its activities at the molecular and cellular levels, as well as in animals and humans.

The discovery of curcumin dates back 2 centuries ago when Vogel and Pelletier<sup>1</sup> isolated "yellow colouring matter" from the rhizomes of turmeric and named it curcumin (the word curcumin stems from the Persian word "kurkum" [curcuma], which means saffron). Later, the substance was found to be a mixture of resin and turmeric oil. For several decades, chemists reported possible structures of curcumin; however, in 1910, the chemical structure of curcumin was identified as diferuloylmethane or 1,6-heptadiene-3,5-dione-1,7-bis(4-hydroxy-3-methoxyphenyl)-(1E,6E).

Although turmeric has been used as a dietary spice and a cure for human ailments for thousands of years in Asian countries, the biological characteristics of curcumin were not identified scientifically until the mid-20<sup>th</sup> century. In a paper<sup>2</sup> published in *Nature* in 1949, the authors reported that curcumin was active against strains of *Staphylococcus aureus*, *Salmonella paratyphi*, *Trichophyton gypseum*, and *Mycobacterium tuberculosis*. In the 1970s, the cholesterol-lowering, antidiabetic, anti-inflammatory, and antioxidant activities of curcumin were reported, and in the 1980s, its anticancer activity in both in vitro and in vivo models was reported.<sup>3</sup> By 1995, anti-inflammatory activity and the mechanisms involved were elucidated by the authors.<sup>4</sup>

Although curcumin is effective against many human ailments, it has poor bioavailability, apparently due to its poor absorption, rapid metabolism, and rapid systemic elimination. Efforts have been made to improve its bioavailability by improving these factors. In spite

of these limitations in bioavailability, as little as 150 mg curcumin administered twice a day orally in humans (n=23) was sufficient to lower serum levels of inflammatory biomarkers such as endothelin-1, interleukin-6, tumor necrosis factor- $\alpha$  and malondialdehyde levels.<sup>5</sup>

Curcumin is now regarded as a "new drug" with great potential and is being used as a supplement and marketed in several forms, including capsules, tablets, ointments, energy drinks, soaps, and cosmetics.

Curcumin has been shown to modulate various signaling molecules, including inflammatory molecules, transcription factors, enzymes, protein kinases, protein reductases, carrier proteins, cell survival proteins, drug resistance proteins, adhesion molecules, growth factors, receptors, cell cycle regulatory proteins, chemokines, DNA, RNA, and metal ions. Curcumin may cause up- or downregulation, depending on the target and cellular context. These targets fall into 2 categories: those to which curcumin binds directly and those whose activity is modulated indirectly by curcumin; both of these areas are extensively covered in review articles. Among the most important targets of curcumin are the pro-inflammatory transcription factors, which regulate the expression of genes that contribute to tumorigenesis, cell survival, cell proliferation, invasion, and angiogenesis. Another important target is the protein kinases.

Because it modulates multiple signaling molecules, curcumin has been reported to possess pleiotropic activities. In vitro, it has been shown to possess anti-inflammatory, antioxidant, pro-apoptotic, chemopreventive, chemotherapeutic, antiproliferative, wound healing, antinociceptive, antiparasitic, and antimalarial properties.

Curcumin has also been shown to have activity as an antioxidant and free radical scavenger in several in vitro studies. This activity is due to either the hydroxyl group or the methylene group of the  $\alpha$ -diketone (heptadiene-dione) moiety. It is also cytotoxic to cancer cells, via the mechanism of apoptosis, and has wide antimicrobial activity.

In animal studies, curcumin has been found to be active against diabetes, obesity, neurological and psychiatric disorders (such as depression, Alzheimer's disease, and Parkinson's disease), and cancer (particularly colon cancer) and cancer-related symptoms such as fatigue, neuropathic pain, and cognitive deficit. It has also shown potential against chronic illnesses affecting the eyes, lungs, liver, kidneys, and gastrointestinal and cardiovascular systems.

About 50 clinical trials have been completed on curcumin's potential in humans. Most have suggested that curcumin is safe and effective to treat a number of diseases, with the most promising effects observed with cancer, inflammatory conditions, skin, eye and neurological disorders, diabetic nephropathy, and pain. Other trials have investigated the role of curcumin in improving body weight and reducing lipid levels in patients with acute coronary syndrome. At the time of this article's preparation, curcumin was undergoing investigation in more than 30 clinical trials, say the authors, adding, "We expect that the completion of these clinical trials will provide further credence to the already established positive effects of curcumin."

Curcumin has not yet been approved to treat any human diseases, say the authors, who suggest that more extensive and well-controlled human studies are needed to demonstrate its safety and efficacy. "Future research should be focused on bringing this

fascinating molecule to the forefront of therapeutic agents for the treatment of human diseases," they write.

—Shari Henson

## References

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<sup>2</sup>Schraufstatter E, Bernt H. Antibacterial action of curcumin and related compounds. *Nature*. 1949:164(4167):456-457.

<sup>3</sup>Kuttan R, Bhanumathy P, Nirmala K, George MC. Potential anticancer activity of turmeric (*Curcuma longa*). *Cancer Lett.* 1985;29(2):197-202.

<sup>4</sup>Singh S, Aggarwal BB. Activation of transcription factor NF-kappa B is suppressed by curcumin (diferuloylmethane) [corrected] [published erratum appears in *J Biol Chem.* 1995;270(50):30235]. *J Biol Chem.* 1995;270(42):24995-25000.

<sup>5</sup>Usharani P, Mateen AA, Naidu MU, Raju YS, Chandra N. Effect of NCB-02, atorvastatin and placebo on endothelial function, oxidative stress and inflammatory markers in patients with type 2 diabetes mellitus: a randomized, parallel-group, placebo-controlled, 8-week study. *Drugs R D.* 2008;9(4):243-250.

The American Botanical Council has chosen not to reprint the original article.