

Ayurvedic Drug Misails for Target Disease

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ABSTRACT

Cancer cells in a part of the body start to grow out of control and produce many kinds of cancer, but they all start because of this out-of-control growth of cancerous cells and they divided too quickly as compare to normal cell. Cancer cell growth is different from normal cell growth hence, it is difficult to detect cancer cell than normal cell. Some plant species mentioned in the Indian systems of ayurvedic medicines which treated various types of cancer. The modern scientific approaches of the ayurvedic plant preparation showed best health care of patients. In case of cancer treatment, the Pancha Karma was the best for treat an early stage of cancer. Ayurvedic dosage form like Gutica, vati, bhasma, churna, lepa, swarasa, paka, leha, kashaya is also important for treating cancer.

Keywords: Cancer, Ayurveda, Ayurvedic plant

INTRODUCTION

Cancer [1] is one of the most dreaded diseases of the 20th century and spreading further with continuance and increasing incidence in 21st century. In the United States, 5% deaths of persons occur due to cancer. It is considered as an adversary of modernization and advanced pattern of socio-cultural life dominated by Western medicine. Multidisciplinary scientific investigations are making best efforts to combat this disease, but the sure-shot, perfect cure is yet to be brought into world medicine. Recently, a greater emphasis has been given towards the researches on complementary and alternative medicine that deals with cancer management. Several studies have been conducted on herbs under a multitude of ethno botanical grounds. For example, Hartwell has collected data on about 3000 plants, those of which possess anticancer properties and subsequently been used as potent anticancer drugs.

Ayurveda, a traditional Indian medicine of plant drugs has been successful from very early times in using these natural drugs and preventing or suppressing various tumors using various lines of treatment. The broad aim of this article is to provide a general outline on descriptions of cancers and their management from an ayurvedic practitioner's perspective underlying its scientific principles involved in treating these conditions with the use of natural products. This article reviews the available literature regarding researches on anticancerous ayurvedic herbs and also includes a summary of treatment strategies for various cancers. It is written with an intention to raise awareness and encourage implementation of ayurvedic therapies for combating cancer and suggesting

an integrated approach in tumor management and treatment [2]. According to a report by the World Health Organization (WHO), cancer remains a major cause of death worldwide. In 2008 there were 7.6 million deaths (approximately 13% of all deaths). In addition, approximately 70% of all cancer deaths in 2008 occurred in countries of low and middle-income status [3].

Marie (1867–1934) and Pierre Curie (1859–1906) discovered radium in 1895, which became the basis for application of radiotherapy in cancer patients. In 1898, William Bradley Coley (1862–1936), the American pioneer of cancer immunotherapy suggested a treatment modality based on provoking an immune response to a sterilized bacterial extract in cases of lymphoma and sarcoma [4].

TYPES OF CANCER

1. Anal cancer
2. Bladder cancer
3. Brain cancer

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4. Carcinoid tumor, gastrointestinal
5. Gallbladder cancer
6. Gastric (stomach) cancer
7. Liver cancer (primary)
8. Lung cancer, non-small cell
9. Lung cancer, small cell
10. Pancreatic cancer
11. Renal cell carcinoma (kidney cancer)
12. Salivary gland cancer
13. Small intestine cancer
14. Vaginal cancer

ORIGIN OF AYURVED

'Ayurveda' is generally understood as 'Science of life' translating 'Ayuh(r)' as life and 'Veda' as science. However, the word 'science' in its conventional meaning is not an appropriate equivalent for 'Veda' [5]. Ayurvedic medicine (also called Ayurveda) is one of the world's oldest medical systems. It originated in India and has evolved there over thousands of years. In the United States, Ayurvedic medicine is considered complementary and alternative medicine (CAM)-more specifically, a CAM whole medical system. Many therapies used in Ayurvedic medicine are also used on their own as CAM-for example, herbs, massage, and specialized diets. Ayurvedic medicine, as practiced in India, is one of the oldest systems of medicine in the world. Many Ayurvedic practices predate written records and were handed down by word of mouth. Two ancient books, written in Sanskrit more than 2,000 years ago, are considered the main texts on Ayurvedic medicine-Charaka Samhita and Sushruta Samhita. Ayurvedic medicine uses a variety of products and techniques to cleanse the body and restore balance. Some of these products may be harmful if used improperly or without the direction of a trained practitioner. For example, some herbs can cause side effects or interact with conventional medicines [6]. Charaka and Sushrutasamhitas, two well-known Ayurvedic classics, describe cancer as inflammatory or non-inflammatory swelling and mention them as either Granthi (minor neoplasm) or Arbuda (major neoplasm). Ayurvedic literature defines three body-control systems, viz., the nervous system (Vata or air), the venous system (Pitta or fire), and the arterial system (Kapha or water) which mutually coordinate to perform the normal function of the body. In benign neoplasm (Vataja, Pittaja or Kaphaja) one or two of the three bodily systems are out of control and is not too harmful because the body is still trying to coordinate among these systems. Malignant tumors (Tridosaja) are very harmful because all the three major bodily systems lose mutual coordination and thus cannot prevent tissue damage, resulting in a deadly morbid condition [7].

AYURVEDIC PREPARATION

1) *Azadirachta indica* (Neem)

It is a member of the *Meliaceae* family, is a fast-growing tropical evergreen tree with a highly branched and stout, solid stem. All parts of this tree, particularly the leaves, bark, seed-oil and their purified products are widely used for treatment of cancer. In those 60 different types constituents of biochemical's including terpenoids and steroids have been purified from this plant. Pre-clinical research work done on the *Azadirachta Indica* of the anticancer properties of the crude and purified products from this plant. The anticancer properties of the plant were studied largely in terms of its preventive, protective, tumor-suppressive, immunomodulatory and apoptotic effects against various types of cancer and their molecular mechanisms [8].

2) *Basellarubra* (Malabar spinach)

It is also known as in English Malabar spinach, Indian spinach and Hindi *lalbachlu* belongs to family *Basellaceae*. It contains phytochemical constituents such as Betacyanins, Carotenoids and Organic acids Triterpene oligoglycosides, Basellasaponins A, B, C, and D having a dioxolane type substituent, Alkaloids, Tannins, Phenols, Proteins and amino acids, Steroids and triterpenoids, Carbohydrates. The plant powder and leaves powder are triturated with sour buttermilk (takra) with salt for preparing a poultice and used to treat arbuda. *Basellarubra* extract is given in dose of 500 mg/kg. *Basellarubra* (B. Alba var. *Cordifolia*) is a traditional Indian medicine used to treat cancer [9].

3) *Andrographis paniculata* (Kalmegh)

Andrographis paniculata commonly known as 'King of Bitters' belongs to family *Acanthaceae*. It is distributed in tropical Asian countries. *Andrographis paniculata* phytoconstituents contains andrographolide and a compound named kalmeghin, diterpenes, lactones, flavonoids, alkanes, ketones, and aldehydes. The extract contains andrographiside and neoandrographolide from these plants are proved to benefit against tumorigenesis. Andrographolide reduces the adhesion of gastric cancer cells which block E-selection expression. The effect of ethanolic extract and andrographolide on cell-mediated immune responses in normal and tumor-bearing control animals was reported [10].

4) *Allium sativum* (Garlic)

Allium sativum commonly known as garlic belongs to family *Amaryllidaceae*. In that more than two hundred Chemical Constituents present volatile oil with sulfur containing compounds like Ajoene (4,5,9-trithiadodeca-1,6,11-triene-9-oxide), Alliin and Allicin, enzymes like peroxidase, allinase, myrosinase. The water and lipid soluble allyl sulfur compounds are effective in blocking a myriad of chemically induced tumors. These plants block in nitrosamine formation and metabolism. So, nitrosamine

blockage in the initiation and promotion phases of the carcinogenicity of various compounds, including polycyclic hydrocarbons, provides evidence that garlic and its constituents can alter several phase I and II enzymes. The higher consumption of garlic to treat the stomach cancer and prostate cancer [11]

5) *Aloe barbadensis* (Aloe vera)

It also known as kumari, Chinese aloe, indian aloe, true aloe, barbados aloe belongs to family *Liliaceae*. The active chemical constituents of Aloins, barbaloins a-barbaloins fatty acids, cholesterol, campesterol and β -sitosterol. In case clinical trials of the use of aloe vera mild soap and aloe vera gel against incidence of radiation therapy induced skin reactions, after two weeks apart from five weeks was taken to show any skin changes in the aloe/soap treatment versus three weeks in the soap only treatment. Aloe soap is beneficial for treating the skin which is damaged due to radiation exposure [12]. In another clinical trial involving patients with advanced solid tumours, for whom no other standard effective therapy was available, combination of pineal indole melatonin (MLT) plus Aloe vera extracts produced some therapeutic benefits, at least in terms of stabilization of disease and survival when compared to MLT alone treatment [13].

6) *Barleriapronitis* (Vajradanti)

Barleriapronitis also known as the porcupine flower is a species of plants in the family *Acanthaceae*. The chemical constituent of plant contains flavonoid type phenolic compounds, apigenin, quercetin, naringenin, luteolin and apigenin glucuronide. The *Barleriapronitis* oil prepared with whole plant is indicated for external application during acute stages of cyst in blood vessels [14].

7) *Commiphoramukul* (guggul)

Commiphoramukul also known as Indian bdellium-tree is a flowering plant in the family *Burseraceae*. The chemical constituents of plant contain myrecene, dimyrecene polymyrecene, guggulsterone, E-Guggulsterone, Guggulsterone-I, II, III cholesterol. Guggulsterone [4, 17(20)-pregnadiene-3, 16-dione] is a plant sterol derived from the gum resin (guggulu) of the tree *Commiphoramukul*. The active constituent guggulsterones inhibited the growth of wide variety of human tumor cells such as leukemia, head and neck carcinoma, multiple myeloma, lung carcinoma, melanoma, breast carcinoma, and ovarian carcinoma. The cancer cell resistance produces particular drug in this case they show activity against the drug-resistant cancer cells. This correlated with the enhanced apoptosis induced by TNF and chemotherapeutic agents [15].

8) *Amorphophallus campanulatus* (Elephant foot)

It also known as is a "Suran" or "Jimmikand". *Amorphophallus* belongs to family *Araceae*. *Amorphophallus campanulatus* contains phytoconstituents

like alkaloids, steroids, fats, fixed oil, flavonoids, tannins, proteins and carbohydrates. *Amorphophallus* species can be investigated further to achieve lead molecules in the search of novel herbal drugs. *Amorphophallus campanulatus* traditionally used for the treatment of abdominal tumors; Epidemiological evidence suggests that dietary factors play an important role in human health as per availability and consumption rate use in the treatment of certain chronic diseases including cancer. The old plant parts burned and then mixed with butter and produce paste applied for tumor destruction [10].

9) *Agrimoniapilosa*

Agrimoniapilosa belongs to family *Rosaceae*, is commonly known as Agrimony. It is perennial herbaceous flowering plant native to temperate regions of the northern hemisphere. The methanolic extract of *Agrimoniapilosa* it gives anti tumour activity may be due to host-mediated actions so that the extract stimulates macrophages and induces cytotoxic macrophages [15].

10) *Acoruscalamus* (Araceae)

Acoruscalamus (Araceae) belongs to family *Acoraceae* and it is found in eastern countries and indigenous to the marshes of the mountains of India. The chemical components present in *Acoruscalamus* α and β -asarone, calamenene, asaronaldehyde, acorenone, calamenone, n-heptanic acid, calanendiol and sesquiterpenes. In that α -asarone showed anti-cancer activity against human carcinoma cells and β -asarone is also responsible for anti-carcinogenic activity [16].

11) *Boswelliaserrata* (Kundur)

Boswelliaserrata also known as Shallkiguggul belongs to family *Burseraceae* it contains active component is Boswellic acid. It showed antitumour effects in addition to its anti-inflammatory effects. *Boswelliaserrata* is beneficial for treating brain tumours, leukemic cells, colon cancer cells, metastatic melanoma and fibrosarcoma cells, and hepatoma [17-19].

12) *Baliospermummontanum* (Jamalgota)

Baliospermummontanum is also known as Danti belongs to family *Euphorbiaceae*. The *Baliospermummontanum*, *Plumbagozeylanica*, *Euphorbia neriifolia*, *Calotropisprocera*, jaggery, *Semecarpusanacardium* all the plants make powder produce a paste applied over the tumours [9].

13) *Crocus sativus* (Saffron)

Crocus sativus also known as Kumkuma. Saffron is a small perennial plant of oriental origin. It grows from a bulb and flowers in the fall with white, lilac or violet flowers. Saffron contains chemical constituents is carotenoid pigments it showed antioxidant properties and flower contains vitamins like Riboflavin and Thiamine. The biomedical properties of saffron and its potential use in cancer therapy and prevent

the dividing of cancer cells. In case of in vitro and in vivo investigations focused on the anticancer activity of saffron (*Crocus sativus L.*) and its principal ingredients [20-21].

14) *Curcuma domestica* (Turmeric)

It also known as turmeric belongs to family *Zingiberaceae*. The active constituents of turmeric are Zingiberene, curcumin, curcuim, alpha- & betaturmerone, zedoarondiol, alpha- & delta- atlantones, bisaboladienones, bisabolenes, bisacumol, bisacurone, curlone, curdinone, curcumins and derivatives, curcumenone, curcumenol, caryophyllenes, curzerenones, germacron derivatives, beta-sesquiphellandrene, alpha-turmerine, turmeronols, beta-turmerone, borneol, isoborneol, camphene, camphor. The produce powder in combination of turmeric, *Symplocos racemosa*, *Soymida febrifuga* and grinding well is mixed with honey produce a paste this is used as an external remedy. Curcumin (diferuloylmethane) is an active component of turmeric (*Curcuma longa*) used as a spice and as an Ayurvedic medicine for centuries to treat various diseases. Curcumin showed carcinogenesis activity of the skin, liver, lung, colon, stomach and breast. It also inhibits the growth of a wide variety of tumour cells in culture and to promote apoptosis through Bid cleavage, cytochrome c release, caspase-9 activation and then caspase-3 activation [22-44]. In vitro anticancer efficacy of CurcuEmulsomes when combined with PiperineEmulsomes highlight the potential of the system for future in vivo and clinical studies. It is essential to emphasize that the present study is the first report disclosing emulsomes as the nanocarrier system achieving combinational cancer therapy with piperine and curcumin on HCT116 CRC cell line [45].

15) *Flavopiridol* (*Dysoxylum binctariferum*)

Flavopiridol is a semisynthetic flavonoid closely related to a compound originally isolated from the stem bark of *Dysoxylum binctariferum* (also called rohitukine from *Amoorohituka*), a plant indigenous to India and described in Ayurveda. The tyrosine phosphorylation of CDK 2 is also inhibited by this flavone. Through inhibition of CDKs, flavopiridol induces arrest of cell growth at the G1 and G2 phases of the cell cycle. Because of its ability to suppress the growth of breast carcinoma, lung carcinoma, chronic B cell leukaemia and lymphoma, multiple myeloma and head and neck squamous cell carcinoma, flavopiridol is currently in clinical trials for the treatment of several cancers [45-58].

16) *Ficus bengalensis* (Phagweri)

Ficus bengalensis is commonly known as phagwari belongs to family *Urticaceae*. *Ficus bengalensis* and *Saussurea lappa* make powder produce a mixture and applied on infected bone area where is tumour growth [22].

17) *Flacourtiaromantchi* (Bhanber)

Flacourtiaromantchi commonly known as baichi or Katai belongs to family *Flacourtiaceae*. It is an indigenous plant

widely distributed in Bangladesh and India. The *Flacourtiaromantchi*, *Cassia fistula*, *Capparis sepia* all drug make powder produce a paste and applied it used to treatment for kaphaja tumours [8,46].

18) *Moringaoleifera* (Mungna)

Moringaoleifera commonly known as mungna belongs to family *Moringaceae*. The plant of *Moringaoleifera* seeds, *Solanum xanthocarpum*, *Sinapis dichotoma*, *Holarrhena antidysenterica* and *Nerium odoratum* roots produce a powder mix with buttermilk make a paste. It is used for arbuda tumors [22].

19) *Madhuca indica*

Madhuca indica belongs to family *Sapotaceae*. *Madhuca indica*, *Syzygium cumini*, *arjuna Terminalia arjuna* and *Salix caprea* barks of all the plant parts make powder and produce a paste prescribed for local application [9].

20) *Oxoxylum indicum* (podaval)

Oxoxylum indicum also called tetu it belongs to family *Bignoniaceae*. The drug *Oxoxylum indicum* used for treatment of granthi cancer [9].

21) *Pandanus odoratissimus* (kewada)

Pandanus odoratissimus its common name kewada belongs to family *Pandanaceae*. The plant *Pandanus odoratissimus* make paste with sugar was applied externally [9].

22) *Phyllanthus niruri/amarus* (Leafflower)

Phyllanthus niruri/amarus its common name is Buinowala belongs to family *Euphorbiaceae*. An aqueous extract of *Phyllanthus amarus* was applied locally to treat the tumor. It plays a major role in disruption of HBsAg mRNA transcription and post-transcription which could be beneficial against viral carcinogenesis [59-60].

23) *Prosopis cineraria* (chikura)

Prosopis cineraria its common name is chikura belongs to family *Mimosaceae*. This paste made up of *Prosopis cineraria* seeds, *Raphanus sativa*, *Moringaoleifera*, barley and mustard with sour buttermilk was applied locally for disintegrating cysts [61].

24) *Pterospermum acerifolium*

Pterospermum acerifolium its common name is kanakcha maka belongs to family *Sterculiaceae*. The flowers of *Pterospermum acerifolium* make paste with sugar was applied locally [62]

25) *Raphanus sativus*

Raphanus sativus belongs to family *Brassicaceae* its common name is black radish. The plant *Raphanus sativus* produce powder then makes paste with the radish ash and locally applied against kaphaja arbuda [62].

26) *Vitisvinifera*

Vitisvinifera belongs to family *vitaceae* is also known as vine. The powder of Terminalia chebula, grape juice and sugar cane juice all this mix well and it used to treat cancer. Resveratrol it is a natural product derivative from grape juice. The resveratrol act as cancer chemo preventive activity and anticancer properties, as suggested by its ability to it decreases growth variety of tumor cells, including lymphoid and myeloid cancers, multiple myeloma, cancers of the breast, prostate, stomach, colon, pancreas and thyroid, melanoma, head and neck squamous cell carcinomas, ovarian carcinoma, and cervical carcinoma [63-64].

27) *Withaniasomnifera* (Ashwagandha)

Withaniasomnifera belongs to family *Solanaceae*. It is used as traditional medicine for centuries for the treatment of various disease. The various parts of *Withaniasomnifera* and its constituents are effective in prevention and treatment of different kinds of cancer like colon cancer, lung cancer, blood cancer, skin cancer, breast cancer, renal cancer, fibrosarcoma, prostate cancer, pancreatic cancer. *Withaniasomnifera* act as prevention and treatment of different forms of cancer like prostate and lung cancers, it prevents cancer in last stage also and this wonder medicinal herb is found to be beneficial in many cancer patients [65].

28) *Zingiberofficinale Rosc.* (Ginger)

Zingiberofficinale Rosc it also known as *assunth* belongs to family *Zingiberaceae*. *Zingiber officinale Rosc*, which is widespread in Southeast Asia. The active constituent of plant is Zerumbone (2,6,9,9-tetramethyl-[2E,6E,10E]-cycloundeca-2,6,10-trien-1-one) was first isolated in 1956 from the essential oil of the rhizomes of a wild ginger. Zerumbone decreases the proliferation of colon cancer and breast cancer, with minimal effects on normal cells. The pungent principle of rhizome produces anticancer activity and its powder is given to treat cancer [66].

29) *Bacopamonniera*

Bacopa is a medicinal herb used in Ayurveda, where it is also known as "Brahmi" belongs to family *Plantaginaceae*. The constituent most studied has been bacoside A, bacoside A3, bacoside II, bacosaponin C, and a jujubogenin isomer of bacosaponin C. The whole plant of *bacopamonniera* used to produce extract, and bacoside concentrations may vary depending upon which part of the plant extracted. The traditional use of *bacopamonniera* Ayurvedic treatment for brain cancer [67].

30) *Piper longum* (pippali)

Piper longum also called as Indian long pepper belongs to the family *Piperaceae*. The active chemical constituent is aromatic oil, piperine, alkaloids, sesamin and pipalestrol. The roots of this plant contain piperin, pippalartin, piperleguminin, sterols and glycosides, piperlongumine or

piplartine and dihydrostigmasterol. The active constituents piperine it showed the antitumour activity in the cancer therapy [68].

AYURVEDIC TREATMENT

When we live according to our constitution, daily and seasonal rhythm and every once in a while, receive a Pancha Karma treatment - to prevent or restore the imbalance of doshas and dhatus - a number of conditions can be relieved at an early stage. The following suggestions can be used to maintain a healthy body and mind and to prevent the development of a tumor in the best possible way. A large part of the suggestions is aimed at prevention.

AYURVEDIC DOSAGE FORM

1. Cap. Phyllanthus Niruri - 2 twice daily
2. Cap. Pitta Balance - 1 twice daily
3. Syp. Amlycure DS - 20 ml twice daily
4. Syp. Nirocil - 10 ml twice daily
5. Maharishi Amrit - Kalash - 1 teaspoonful twice daily, tablets 2 twice daily
6. Swamla compound -1 teaspoonful twice daily
7. Cap. Cruel - 2 twice daily
8. Tab. Kachnaar Guggul - 2 twice daily
9. Cap. Curcumin - 2 twice daily

CONCLUSION

Cancer is a serious health problem due to this increased death rate. GIT cancer is one of the common cancers in most people. Ayurveda the science of life is a traditional medicine system used to treat cancer. All the above plant products have been studied well and described for their medicinal properties. With these plants other supportive medicines were used according to ayurvedic principles patient achieved complete disease treating without any side effects. Medicinal property of ayurvedic plant preparation inducing cell proliferation and self-renewal of damaged proliferating tissues, and replenishing them by eliminating damaged or mutated cells with fresh cells. Medicinal plants have contributed a rich health to human beings.

REFERENCES

1. Harry N. Abrams, Lyons AS, Petrucelli RJ (2010) The History of Cancer. American Cancer Society. pp: 1-16.
2. Patel D, Mansoori AM (2012) Cancer-an Ayurveda Perspective. Int J Adv Res Pharm Bio Sci 2: 179-195.
3. Kramer J (2015) Cancer Facts & Figures. American Cancer Society. pp: 1-52.
4. Azizi MH, Bahadori M, Azizi F (2013) History of Cancer in Iran. Arch Iran Med 16: 613-622.

5. Manyam B, Khalsa B (2009) Ayurvedic Medicine: An Introduction. U.S. Department of Health and Human Services. pp: 1-8.
6. Balachandran P, Rajgopal G (2005) Cancer-an ayurvedic perspective. *Pharmacol Res* 51: 19-30.
7. Bhisagrattha KL. *Sushrutasamhita* (1991) Varanasi: Choukhamba Orientalia.
8. Milner JA (2001) A Historical Perspective on Garlic and Cancer. *J Nutr* 131: 1027S-1031S.
9. Paul R, Murari P, Sah NK (2011) Anticancer biology of *Azadirachta indica* L (neem). *Cancer Biol Ther* 12: 467-476.
10. Glaser T, Winter S, Groscurth P (1999) Boswellic acids and malignant glioma: induction of apoptosis but no modulation of drug sensitivity. *Br J Cancer* 80: 756-765.
11. Olsen DL, Bradley C, Johnson M, Macias JL, Love V, et al. (2001) The effect of Aloe vera gel/mild soap versus mild soap alone in preventing skin reactions in patients undergoing radiation therapy. *Oncol Nurs Forum* 28: 543.
12. Lissoni P, Giani L, Zerbin S, Trabattoni P, Rovelli F (1998) Biotherapy with the pineal immunomodulating hormone melatonin versus melatonin plus aloe vera in untreatable advanced solid neoplasms. *Nat Immunol* 16: 27-33.
13. Kinjavadekara RS (1998) *Astangasangraha*. New Delhi: Uppal Publishing House.
14. Shishodia S, Aggarwal BB (2004) Guggulsterone inhibits NF-kappaB and IkappaBalpha kinase activation, suppresses expression of anti-apoptotic gene products, and enhances apoptosis. *J Biol Chem* 279: 47148-47158.
15. Winking M, Sarikaya S, Rahmanian A (2000) Boswellic acids inhibit glioma growth: A new treatment option. *J Neuro-Oncol* 46: 97-103.
16. Jing Y, Nakajo S, Xia L (1999) Boswellic acid acetate induces differentiation and apoptosis in leukemia cell lines. *Leukemia Res J* 23: 43-50.
17. Shao YC, Ho T, Chin CK (1998) Inhibitory activity of boswellic acids from *Boswelliaserrata* against human leukemia HL-60 cells in culture. *Planta Medica* 64: 328-331.
18. Liu JJ, Nilsson A, Oredsson S (2002) Boswellic acids trigger apoptosis via a pathway dependent on caspase-8 activation but independent on Fas/Fas ligand interaction in colon cancer HT-29 cells. *Carcinogenesis* 23: 2087-2093.
19. Zohara YB (2012) Contribution of Selected Medicinal Plants for Cancer Prevention and Therapy. *Scientific J Faculty Med Naissensis* 29: 117-123.
20. Abdullaev FL, Espinosa-Aquire LL (2004) Biomedical properties of saffron and its potential use in cancer therapy and chemoprevention trials. *Cancer Detect Prevent J* 8: 426-432.
21. Murthy KRS (2001) *Bhavaprakasa of bhavamisra Madhya and UttaraKhanda*. Varanasi: Krishnadas Academy. pp: 2.
22. Bharti AC, Donato N, Aggarwal BB (2003) Curcumin (diferuloylmethane) inhibits constitutive and IL-6-inducible STAT3 phosphorylation in human multiple myeloma cells. *J Immunol* 171: 3863-3871.
23. Anto RJ, Mukhopadhyay A, Denning K, Aggarwal BB (2002) Curcumin (diferuloylmethane) induces apoptosis through activation of caspase-8, BID cleavage and cytochrome c release: its suppression by ectopic expression of Bcl-2 and Bcl-xl. *Carcinogenesis* 23: 143-150.
24. Mukhopadhyay A, Bueso RC, Chatterjee D (2001) Curcumin down regulates cell survival mechanisms in human prostate cancer cell lines. *Oncogene* 20: 7597-7609.
25. Mukhopadhyay A, Banerjee S, Stafford LJ (2002) Curcumin-induced suppression of cell proliferation correlates with down-regulation of cyclin D1 expression and CDK4-mediated retinoblastoma protein phosphorylation. *Oncogene* 21: 8852-8861.
26. Aggarwal BB, Kumar A, Bharti AC (2003) Anticancer potential of curcumin: Preclinical and clinical studies. *Anti-Cancer Res* 23: 363-398.
27. Shishodia S, Potdar P, Gairola CG, Aggarwal B (2003) B. Curcumin (diferuloylmethane) downregulates cigarette smoke-induced NFkappaB activation through inhibition of IkappaBalpha kinase in human lung epithelial cells: correlation with suppression of COX-2, MMP-9 and cyclin D1. *Carcinogenesis* 24: 1269-1279.
28. Bharti AC, Takada Y, Aggarwal B (2004) B. Curcumin (diferuloyl methane) inhibits receptor activator of NF-kappa B ligand-induced NF-kappa B activation in osteoclast precursors and suppresses osteoclastogenesis. *J Immunol* 172: 5940-5947.
29. Bharti AC, Shishodia S, Reuben JM (2004) Nuclear factor-kappa B and STAT3 are constitutively active in CD138+ cells derived from multiple myeloma patients, and suppression of these transcription factors leads to apoptosis. *Blood* 103: 3175-3184.
30. Bharti AC, Donato N, Singh S, Aggarwal B (2003) B. Curcumin (diferuloylmethane) down-regulates the

- constitutive activation of nuclear factor-kappa B and I-kappaB kinase in human multiple myeloma cells, leading to suppression of proliferation and induction of apoptosis. *Blood* 101: 1053-1062.
31. Aggarwal S, Takada Y, Singh S (2004) Inhibition of growth and survival of human head and neck squamous cell carcinoma cells by curcumin via modulation of nuclear factor-kappaB signaling. *Int J Cancer* 111: 679-692.
 32. Aggarwal BB, Takada Y, Oommen OV (2004) From chemoprevention to chemotherapy: Common targets and common goals. *Expert Opin Investig Drugs* 13: 1327-1338.
 33. Li L, Aggarwal BB, Shishodia S (2004) Nuclear factor-kappa B and I-kappa B kinase are constitutively active in human pancreatic cells, and their down-regulation by curcumin (diferuloylmethane) is associated with the suppression of proliferation and the induction of apoptosis. *Cancer* 101: 2351-2362.
 34. Dorai T, Aggarwal BB (2004) Role of chemo preventive agents in cancer therapy. *Cancer Lett* 215: 129-140.
 35. Takada Y, Bhardwaj A, Potdar P, Aggarwal BB (2004) Nonsteroidal anti-inflammatory agents differ in their ability to suppress NF-kappaB activation, inhibition of expression of cyclooxygenase-2 and cyclin D1, and abrogation of tumor cell proliferation. *Oncogene* 23: 9247-9258.
 36. Aggarwal BB, Shishodia S (2004) Suppression of the nuclear factor-kappa B activation pathway by spice-derived phytochemicals: reasoning for seasoning. *Ann N Y Acad Sci* 1030: 434-441.
 37. Bharti AC, Takada Y, Aggarwal BB (2005) Cleavage and caspase activity to assess chemosensitivity. *Methods Mol Med* 111: 69-78.
 38. Siwak DR, Shishodia S, Aggarwal BB, Kurzrock R (2005) Curcumin-induced antiproliferative and proapoptotic effects in melanoma cells are associated with suppression of I-kappaB kinase and nuclear factor kappaB activity and are independent of the B-Raf/mitogen activated/ extracellular signal-regulated protein kinase pathway and the Akt pathway. *Cancer* 104: 879-890.
 39. Shishodia S, Amin HM, Lai R, Aggarwal B (2005) B. Curcumin (diferuloylmethane) inhibits constitutive NF-kappaB activation, induces G1/S arrest, suppresses proliferation, and induces apoptosis in mantle cell lymphoma. *Biochem Pharmacol* 70: 700-713.
 40. Yan C, Jamaluddin MS, Aggarwal B (2005) Gene expression profiling identifies activating transcription factor 3 as a novel contributor to the proapoptotic effect of curcumin. *Mol Cancer Ther* 4: 233-241.
 41. Shishodia S, Gethi G, Aggarwal B (2005) B. Curcumin: Getting back to the roots. *Ann N Y Acad Sci* 1056: 206-217.
 42. Aggarwal BB, Kumar A, Bharti AC (2004) Therapeutic potential of curcumin derived from turmeric (*Curcuma longa*). Marcel Dekker, New York.
 43. Aggarwal BB, Kumer S, Aggarwal S, Shishodia S (2005) Curcumin derived from turmeric (*Curcuma longa*): A spice for all seasons. *Phytochemicals in Cancer Chemoprevention*. Available online at: https://curcumin.co.nz/pdf/Curcumin_A_Spice_For_All_Seasons.pdf
 44. Carlson BA, Dubay MM, Sausville EA (1996) Flavopiridol induces G1 arrest with inhibition of cyclin-dependent kinase (CDK) 2 and CDK4 in human breast carcinoma cells. *Cancer Res* 56: 2973-2978.
 45. Losiewicz MD, Carlson BA, Kaur G (1994) potent inhibition of CDC2 kinase activity by the flavonoid L86-8275. *Biochem Biophys Res Commun* 201: 589-595.
 46. Azevedo WF, Mueller-Dieckmann HJ, Schulze GU (1996) Structural basis for specificity and potency of a flavonoid inhibitor of human CDK2, a cell cycle kinase. *Proc Natl Acad Sci USA* 93: 2735-2740.
 47. Worland PJ, Kaur G, Stetler-SM (1993) Alteration of the phosphorylation state of kinase by the flavone in breast carcinoma cells. *Biochem Pharmacol* 46: 1831-1840.
 48. Kaur G, Stetler SM, Sebers S (1992) Growth inhibition with reversible cell cycle arrest of carcinoma cells by flavones. *J Natl Cancer Inst* 84: 1736-1740.
 49. Bible KC, Kaufmann SH (1996) Flavopiridol: a cytotoxic flavone that induces cell death in noncycling human lung carcinoma cells. *Cancer Res* 56: 4856-4861.
 50. Konig A, Schwartz GK, Mohammad RM (1997) The novel cyclin-dependent kinase inhibitor flavopiridol down regulates and induces growth arrest and apoptosis in chronic B-cell leukemia lines. *Blood* 90: 4307-4312.
 51. Arguello F, Alexander M, Sterry JA (1998) Flavopiridol induces apoptosis of normal lymphoid cells, causes immunosuppression, and has potent antitumor activity In vivo against human leukemia and lymphoma xenografts. *Blood* 91: 2482-2490.
 52. Byrd JC, Shinn C, Waselenko JK (1998) Flavopiridol induces apoptosis in chronic lymphocytic leukemia cells via activation of caspase-3 without evidence of modulation or dependence on functional group. *Blood* 92: 3804-3816.
 53. Gojo I, Zhang B, Fenton RG (2002) The cyclin-dependent kinase inhibitor flavopiridol induces

- apoptosis in multiple myeloma cells through transcriptional repression and down-regulation of Mcl-1. *Clin Cancer Res* 8: 3527-3538.
54. Patel V, Senderowicz AM, Pinto D (1998) Flavopiridol, a novel cyclin-dependent kinase inhibitor, suppresses the growth of head and neck squamous cell carcinomas by inducing apoptosis. *J Clin Invest* 102: 1674-1681.
55. Shapiro GI, Supko JG, Patterson A (2001) A Phase II trial of the cyclin-dependent kinase inhibitor flavopiridol in patients with previously untreated stage IV non-small cell lung cancer. *Clin Cancer Res* 7: 1590-1599.
56. Senderowicz AM, Sausville EA (2000) Preclinical and clinical development of cyclin-dependent kinase modulators. *J Natl Cancer Inst* 92: 376-387.
57. Karp JE, Ross DD, Yang W (2003) Timed sequential therapy of acute leukemia, with flavopiridol: In vitro model for a Phase I clinical trial. *Clin Cancer Res* 9: 307-315.
58. Rajeshkumar NV, Kuttan R (2000) *Phyllanthus amarus* extract administration increases the life span of rats with hepatocellular carcinoma. *J Ethno pharmacol* 73: 215.
59. Lee CD, Thyagarajan SP, Shafritz DA, Burk RD (1996) *Phyllanthus amarus* down-regulates hepatitis B virus mRNA transcription and replication. *Eur J Clin Invest* 26: 1069-1076.
60. Kinjavadekara RS. *Astangasangraha*, New Delhi: Uppal Publishing House 1998.
61. Balachandran B, Govindarajan R (2005) Cancer-an ayurvedic perspective. *Pharmacol Res* 51: 19-30.
62. Aggarwal BB, Ichikawa H, Garodia P (2006) Identification of therapeutic Targets for suppression of inflammation and cancer. *Expert Opin Ther Targets* 10: 87-118.
63. Jang M, Cai L, Udeani GO (1997) Cancer chemopreventive activity of Resveratrol, a natural product derived from Grapes. *Science* 275: 218-220.
64. Singh N, Verma P, Pandey BR, Gilca M (2011) Role of *Withania somnifera* in Prevention and Treatment of Cancer: An Overview. *Int J Pharm Sci Drug Res* 3: 274-279.
65. Kitayama T, Okamoto T, Hill RK (1999) Chemistry of zerumbone. Simplified isolation, conjugate addition reactions, and a unique ring contracting transannular reaction of its dibromide. *J Org Chem* 64: 2667-2672.
66. Dev S. Zermbone (1956) A monocyclic sesquiterpene ketone. *Chem Ind.* pp: 1051.
67. Srivastava S, Mishra N, Misra U (2009) Bacopamonniera -A Future Perspective. *Int J Pharm Sci Drug Res* 1: 154-157.
68. Srivastava P (2014) Therapeutic potential of *Piper longum* L. for disease management - A review. *Int J Pharm Sci* 4: 692-696.
69. Hartwell JL (1969) Plants used against cancer. A survey *Lloydia* 32: 78-107.
70. Bolat ZB, Islek Z, Demir BN, Yilmaz EN, Sahin F, et al. (2020) Curcumin- and Piperine-Loaded Emulsomes as Combinational Treatment Approach Enhance the Anticancer Activity of Curcumin on HCT116 Colorectal Cancer Model. *Front Bioeng Biotechnol* 8: 50.