

## Review Of Medicinal Plants And Their Application In Cancer Therapy

**Dr. Gaviraj.E.N<sup>1</sup>**

Professor & HoD, Department of Pharmacognosy,  
BLDEAs SSM College of Pharmacy and Research Center, Vijayapur, Karnataka State  
[kleraj2009@gmail.com](mailto:kleraj2009@gmail.com)

**Dr.D.Jayarajan<sup>2</sup>**

Head, Dept of Medical Lab Technology, Divine Mother College,  
Korkadu, Puducherry-605110  
email: [asairaj123@gmail.com](mailto:asairaj123@gmail.com),

**Prof. Dr. Jayesh Gujarathi<sup>3</sup>**

Professor in Chemistry, Pratap College Amalner Autonomous  
[jayeshgujarathi496@gmail.com](mailto:jayeshgujarathi496@gmail.com)

**M.Krishnaveni<sup>4</sup>**

Assistant Professor in Biochemistry, Nadar Saraswathi College of Arts & Science, Theni.  
[venibio87@gmail.com](mailto:venibio87@gmail.com)

### ABSTRACT

Cancer affects individuals worldwide. Novel treatments are needed to treat and prevent this deadly disease. Science is studying natural chemicals since they have fewer adverse effects than chemotherapy. Plant secondary metabolites may be used to develop novel anticancer drugs. These compounds, which have become cancer treatments, are advancing the science. Most chemotherapy drugs restrict dose, develop resistance, and are non-selective to normal cells. Making effective cancer treatments is therefore still a clinical challenge. However, plants are rich in naturally occurring chemicals that are physiologically active and may have commercial value or be utilized to generate modified derivatives with increased activity and/or reduced toxicity in cancer treatment. The potential efficacy, accessibility, low cost, lack of toxic side effects, and high use rates of herbal medicines means that they are being seriously considered as anticancer therapy. Science has accelerated. Traditional medicines are safe and effective, hence the WHO recommends them. We reviewed several anticancer herbs.

**Keywords:** Medicinal plants, Anticancer, Cell line, Traditional medicine, Phyto compounds

## INTRODUCTION

A lot of progress has been made in the treatment and prevention of cancer, and it has been a continual fight everywhere. The condition is characterized by an uncontrollable or irreversible cell proliferation in the human body. Resulting in the growth of cancerous tumours that have the potential to spread [1]. Chemotherapy, radiation, and medications made from chemicals are now used as therapies. Chemotherapy may stress and harm patients. Thus, complementary and alternative cancer treatments are prioritized [2]. Herbal treatments have long been the backbone of medical care in developing countries. Plants in medicine have antimicrobial properties. Researchers are looking at extracts from land plants to find ways to treat cancer with nanomaterial [3]. Several types of plants can treat or stop cancer. Several studies have found that some plant species used in poor herbal therapies have anticancer properties [4,5,6,7,8]. Plant kingdom chemicals required for plant survival and "housekeeping" may inhibit malignant cell proliferation and induce apoptosis. This article summarizes plant-derived anticancer drugs.

### Epigenetic characteristics

Alterations to epigenetic processes and their dysregulation have a role in the development of cancer [9]. Cancer cells have dysregulated control over the hypermethylation of tumor-suppressor genes on CpG islands. Tumor-suppressor genes may become inactive as a consequence of this [10]. Under recent years, medications that may prevent or undo epigenetic changes have been in development.

Chemically generated epigenetic drugs including SAHA (Vorinostat, Zolinza), FK228 (Romidespin, Istodax), and 5-aza-2-deoxycytidine (decitabine, Dacogen) have been investigated. DNMT and HDAC inhibitors, respectively. However, making a chemical treatment that selectively kills cancer cells but not normal ones is difficult. As a result, there is a growing need for anticancer drugs derived from organic sources, with a focus on plant species and their chemical compounds [11]. All cancers share susceptibility to signals that inhibit cell growth and produce an unending cycle of replication. Cancer cells persist because apoptosis is never produced and angiogenesis is maintained [12]. Plant-derived compounds inhibit cancer cell proliferation and induce apoptosis.

### Medicinal plants having anticancer potential

#### *Heracleum persicum*

The *Agrobacterium tumefaciens* that caused potato disc crown gall tumours was stopped by a methanol and petroleum ether extract from the root and fruit of the *Heracleum persicum* (Apiaceae) plant. *H. persicum* essential oils have 2.24 mg/mL IC<sub>50</sub> anticancer activity [13].

#### *Ophiorrhiza mungos*

The *Agrobacterium tumefaciens* caused crown gall tumours on potato discs were stopped by a methanol and petroleum ether extract from the root and fruits of *Heracleum persicum* (Apiaceae). H. The IC50 for how well persicum essential oils fight cancer was 2.24 mg/mL.

### ***Rhizophora apiculata***

Luteolin-7-O-glucoside (LUT7G) and camptothecin were taken from the methanolic extract of the leaves and roots of *Ophiorrhiza mungos* (Rubiaceae). They were tested for their anticarcinogenic ability in 4 cancer cell lines (COLO 320 DM, AGS, MCF-7, and A549) and the healthy VERO cell line. The antiradical activity, DNA fragmentation, -catenin expression, and chemo preventive efficacy in vivo were used to measure LUT7apoptosis-causing G's ability. 20 mg/kg lowers the activity of tumour cells [14,15].

### ***Peristrophe bicalyculata***

In lab tests, 2.5–22.3 g/ml of *P. bicalyculata* (Acanthaceae) oil killed human breast cancer cells MCF-7 and MDA-MB-468. A lot of beta-caryophyllene (33.9%), alpha-zingiberene (10.4%), germacrene D (5.0%), and globulol (5.0%) were found in *P. bicalyculata* oil. [16]

### ***Perilla frutescens***

*Perilla frutescens* (Lamiaceae) methanolic extracts were tested on human non-small cell lung A549 carcinoma cells to see if they stopped the cells from dividing. The methanolic extract of the stalk had some effects against cell growth. The stalks of *P. frutescens* may help fight cancer and act as antioxidants. At 10 g/ml, the stalk's hot water extract had moderate DPPH radical scavenging activity of 54.8%, while the leaf and seed extracts had only 5.5 and 6.7%, respectively. Cancer can be stopped by phytochemicals, polyphenols, and flavanoids [17].

### ***Rubia cordifolia***

The methanol extract was made by letting 80% methanol flow through powdered *Rubia cordifolia* roots (Rubiaceae). Many secondary plant chemicals, like two new dimers and four new naphthohydroquinones, were found in *R. roots of cordifolia*. These COX-2 inhibitors might be good places to start looking for ways to prevent cancer with chemoprevention. R. The new anticancer bicyclic hexapeptides RA-VI, VIII, XV, XVI, and XVII were made from the roots of *Cordifolia*. RA-IX and -X were found in *R. cordifolia*, are bicyclic hexapeptides with glutamic acid residues that fight cancer very well. RA-XI, RA-XII, RA-XIII, and RA-XIV were all taken from *R. cordifolia*, showed that it was very good at fighting cancer against P-388 [18].

### ***Radix sophorae***

Leachianone A from *Radix Sophorae* (Leguminosae) causes apoptosis in the human hepatoma cell line HepG2 in vitro through both extrinsic and intrinsic pathways with an IC50 of 3.4 g/ml after 48 hours of treatment. In human hepatoma HepG2 cells, an extract from the

root of *Radix Sophorae* caused mitochondrial dysfunction, apoptosis, and the production and loss of reactive oxygen species (ROS) [19].

### *Taxus yunnanensis*

High amounts of the anticancer taxane diterpenoids paclitaxel and dihalocephalomannine were found in an extract of the roots of the *T. yunnanensis* (Taxaceae) plant. When the leafy parts of *Taxus yunnanensis* were extracted with ethanol, they gave off 14 taxoids and two new taxane diterpenes called dantaxusin C and D. Twelve taxane diterpenes that had already been made from *T. yunnanensis* or *T. chinensis* aerial parts in EtOH extract were tested for their cytotoxicity against KB-VIN and KB-7d cancer cells that were resistant to drugs. The cytotoxicity of seven taxane diterpenes from an Ethanol extract of *T. chinensis* aerial parts was tested on nine human cell lines, including one that was resistant to paclitaxel. Taxane-type diterpenes like 10-deacetyl cephalomannine and 10-deacetyltaxol were found in *T. yunnanensis* wood extract in amounts that stop growth. Antiproliferative diterpenes [20].

### *Vaccinium macrocarpon*

Triterpene cinnamates 3-O-p-hydroxycinnamoyl ursolic acid, which was found in *Vaccinium macrosporan* (Ericaceae) ethyl acetate extracts, had GI50 values of 20 M in most tumour cell lines that were a little bit more active. HepG2 liver cancer cells and MCF-7 breast cancer cells were used to test the antiproliferative effects of parts of cranberry extract. Ursolic acid, quercetin, and 3,5,7,3',4'-pentahydroxyflavonol-3-O-beta-D-glucopyranoside all had EC50 values of 87.4 2.7, 40.9 1.1, and 49.2 4.9 M, respectively [21,22].

## Plant chemicals having anticancer properties

Industrialized nations consume many plants for their health benefits. Asia and Africa have used medicinal herbs for millennia. According to the WHO, some nations continue utilize plant-based medicines, while developing countries are taking advantage of the medicinal benefits of natural substances [23]. Terrestrial plant polyphenols, brassinosteroids, and taxols have anticancer properties.

### Polyphenols

Cancer-fighting chemicals include flavonoids, tannins, curcumin, resveratrol, and gallacatechins [24]. Resveratrol is found in wine, grapes, and peanuts. Gallacatechins are found in green tea. Polyphenols are natural antioxidants that may be good for your health and lower your risk of cancer [25]. Many cancer cells are killed by polyphenols, which are antioxidants.

### Flavonoids

Polyphenolic flavonoids are secondary plant chemicals with 10,000 different parts. They are chemicals in plants that affect how the body works, and their health benefits are being looked

into [26,27]. Flavonoid concentration and cancer cell effects have been examined in several plants, including ferns and traditional Chinese therapies like litchi leaf [28]. Anthocyanins, flavones, flavonols, chalcones, and others are found in the seed [22]. In 2013, Coa et al. found flavonoids and tested them on human lung cancer cells (A456 cell line) from the fern *Dryopteris erythrosora*. Flavonoids kill cancer cells and get rid of free radicals. Hepatoma (Hep-G2), cervical cancer (Hela), and breast cancer (MCF-7) are all resistant to the anticancer effects of pure flavonoids [29].

### Brassinosteroids

Brassinosteroids (BRs) are plant hormones that regulate cell growth and differentiation, stem and root cell elongation, and disease and stress resistance. BRs regulate plant senescence [30]. Plant growth requires them. BRs are another natural cancer treatment. In research on cancerous cells, two natural BRs have been shown to have anticancer properties. At micromolar levels, 28-homocasterone (28-homoCS) and 24-epibrassinolide (24-epiBL) stop cancer cells from growing [31,32]. Cancer cells cannot apoptosis and proliferate endlessly. BRs interact with the cell cycle to restrict proliferation and induce apoptosis [33].

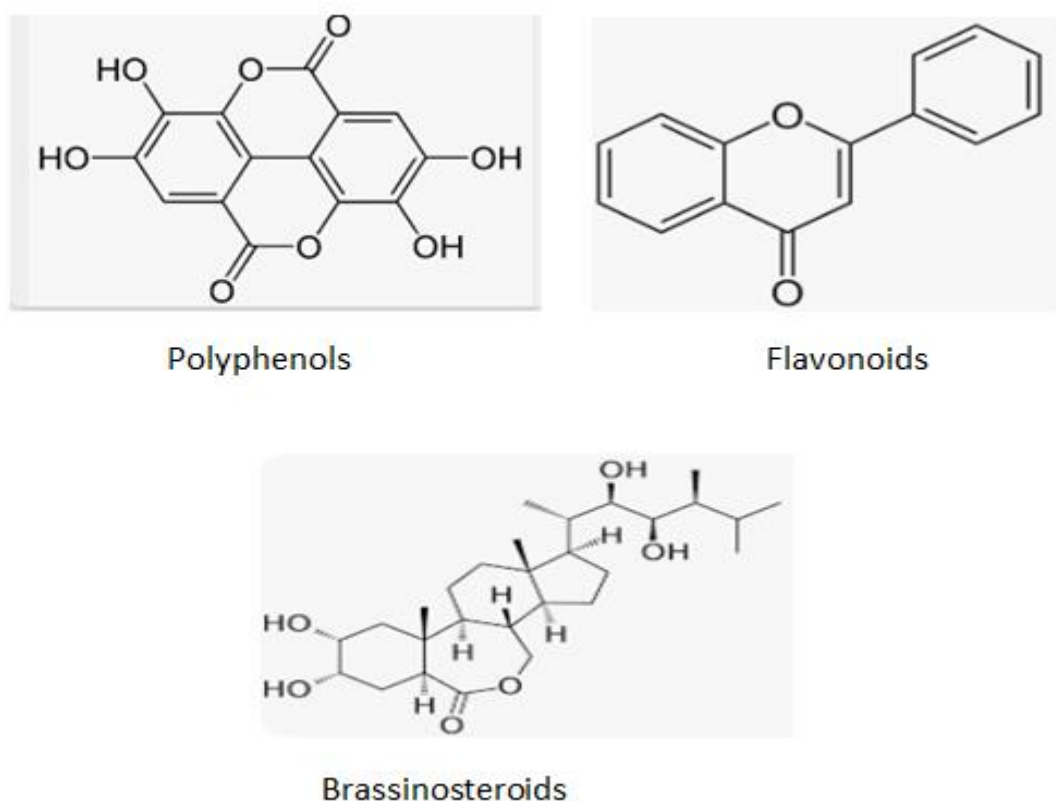


Fig: 1 Plant derived compounds

### **Plant-based medicines that fight cancer**

Plant-based anticancer drugs are safe and accessible. Oral administration is simple [34]. They're also better for healthy cells since they're plant-based [35]. Taxanes, lectins, saponins, lignans, and cyanogenetic glycosides are exceptions [36,37] Plant-based drugs may go into clinical trials if studies show that they are selective, are safe for healthy cell lines, and kill cancer cells. Plant-derived drugs are methyltransferase inhibitors, DNA damage preventatives, histone deacetylase (HDAC) inhibitors, and mitotic disruptors.

### **Enhancing pharmaceutical administration**

Natural medicine developments are enabling new anticancer chemical application and dose approaches. A new drug must be administered well to replace chemotherapy. Nanotechnology is enabling the delivery of drugs using nanoparticles (NPs). Anticancer drugs that need high concentrations may have limited clinical development. NPs enhanced the anticancer effects of *Ananas comosus* bromelain [38].

### **Demands for medicinal plants**

A clinical trial favors plant-based drugs due of their effectiveness. Their cytotoxic effects on cancer cells and non-toxicity to healthy cells make them desirable. Most of the species studied were from poor African and Asian countries where herbal remedies and medicinal plants constitute the backbone of healthcare. The World Health Organization estimated 2007 plant-based drug sales at \$100 billion. Commerce might reach \$5 trillion USD by 2050. In rising countries, medicinal herb demand strains plant populations. For informal trade, some therapeutic plants are produced from wild populations, although this cultivation is uncontrolled [39]. The protection of medicinal plants is a concern that needs attention due to population increase, deforestation, and expanding urbanization [40]. High-value medicinal plants could go extinct if they are used too much because demand is going up. It is very important to protect these plants. Only some portions of wild medicinal plants are utilized as medicine, such as the bark of trees or the bulbs and tubers of bulbous and tuberous plants.

## **CONCLUSION**

Cancer is becoming more popular worldwide. The WHO claimed in 2007 that 7.6 million people died from cancer in 2005, most of them in poor countries. Cancer killed 1 in 4 In 2010, 1.5 million new cases were reported in the United States [41]. According to Cancer Research UK, 8.2 million people died from cancer in 2012, while 14.1 million were diagnosed. As a result, cancer treatment and prevention are critical. Other cancer treatments and chemical-based medicines exist. Chemotherapy may injure non-target tissues, worsening health problems. Thus, complementary treatments using naturally occurring anticancer compounds, preferably from plants, are needed. Due to increased demand for plant-based medicines, high-value medicinal plants are threatened. Rising populations, urbanisation, and deforestation threaten species in developing countries. Cryopreservation, tissue culture, and



plant part replacement are needed to conserve these species 46. Large-scale medicinal plant cultivation and industrial utilization of raw byproducts may help conservation [42]. Plant-derived anticancer drugs inhibit cancer cell lines and are popular. Controlling these agents' exploitation helps satisfy demand and stay sustainable.

## REFERENCE

1. Ochwang<sup>†</sup>I, D.O., Kimwele, C.N., Oduma, J.A., Gathumbi, P.K., Mbaria, J.M. and Kiama, S.G. (2014) „Medicinal plants used in treatment and management of cancer in Kakamega County Kenya“, *Journal of Ethnopharmacology*, 151, pp. 1040-1055. 2.
2. Cancer Research UK (2014) What is cancer? Available at: <http://www.cancerresearchuk.org/about-cancer/what-iscancer> (Accessed 23 January 2015).
3. Sivaraj, R., Rahman, P.K.S.M., Rajiv P, Vanathi, P., Venckatesh R. 2014. Biosynthesis and characterization of *Acalypha indica* mediated copper oxide nanoparticles and evaluation of its antimicrobial and anticancer activity. *Spectrochimica Acta Part A: Molecular and Biomolecular Spectroscopy*, 129: 255-258
4. Freiburghaus, F. Kaminsky, R., Nkunya, M.H.H. and Brun, R. (1996) „Evaluation of African plants for their in vitro trypanocidal activity“, *Journal of Ethnopharmacology*, 55, pp.1-11.
5. Costa-Lotuf, L.V., Khan, M.T.H., Ather, A., Wilke, D.V., Jimenez, P.C., Pessoa, C., Amaral de Moraes, M.E. and Odorico de Moraes, M. (2005) „Studies of the anticancer potential of plants used in Bangladeshi folk medicine“, *Journal of Ethnopharmacology*, 99, 21-30.
6. Cai, Y.Z., Sun, M., Xing, J., Luo, Q. and Corke, H. (2006) „Structure-radical scavenging activity relationships of phenolic compounds from traditional Chinese medicinal plants“, *Life Sciences*, 78, pp. 2872-2888.
7. Fouche, G., Cragg, G.M., Pillay, P., Kolesnikova, N., Maharaj, V.J. and Senabe, J. (2008) „In vitro anticancer screening of South African plants“, *Journal of Ethnopharmacology*, 119, pp. 455-461.
8. . Kamatou, G.P.P., Van Zyl, R.L., Davids, H., Van Heerden, F.R., Lourens, A.C.U. and Viljoen, A.M. (2008) „Antimalarial and anticancer activities of selected South African *Salvia* species and isolated compounds from *S. radula*“, *South African Journal of Botany*, 74, pp. 238-243
9. Schneckeburger, M., Dicato, M. and Diederich, M. (2014) „Plant-derived epigenetic modulators for cancer treatment and prevention“, *Biotechnology Advances*, 32, pp.1123- 1132.
10. Esteller, M. (2007) „Epigenetic gene silencing in cancer: the DNA hypermethylome“, *Human Molecular Genetics*, 16 (1), pp. 50-59.
11. Seidel, C., Florean, C., Schneckeburger, M., Dicato, M. and Diederich, M. (2012) „Chromatin-modifying agents in anti-cancer therapy“, *Biochimie*, 94, pp. 2264-2279.

12. Kumar, S., Pathania, A.S., Saxena, A.K., Vishwakarma, R.A., Ali, A. and Bhunshan, S. (2013) „The anticancer potential of flavonoids isolated from the stem bark of *Erythrina suberosa* through induction of apoptosis and inhibition of STAT signalling pathway in human leukaemia HL-60 cells“, *Chemico-Biological Interactions*, 205, pp. 128-137
13. Noudeh GD, Sharififar F, Noodeh AD, Moshafi MH, Afzadi MA, Behravan E, et al. Antitumor and antibacterial activity of four fractions from *Heracleum persicum* Desf. and *Cinnamomum zeylanicum* Blume. *J Med Plants Res*. 2010;4(21):2176-80
14. Raveendran VV, Vijayan FP, Padikkala J. Antitumor Activities of an Anthraquinone Fraction Isolated from in Vitro Cultures of *Ophiorrhiza rugosa* var *decumbens*. *Integr Cancer Ther*. 2012;12(2):120-8
15. Vinod PV, Guruvayoorappan C. Evaluation of immunostimulant activity and chemoprotective effect of mangrove *Rhizophora apiculata* against cyclophosphamide induced toxicity in BALB/c mice. *Immunopharmacol Immunotoxicol*. 2012;34(4):608-15
16. Ogunwande IA, Walker TM, Bansal A, Setzer WN, Essien EE. Essential oil constituents and biological activities of *Peristrophe bicalyculata* and *Borreria verticillata*. *Nat Prod Commun*. 2010;5(11):1815-8.
17. Lin ES, Chou JH, Kuo PL, Huang YC. Antioxidant and antiproliferative activities of methanolic extracts of *Perilla frutescens*. *J Med Plants Res*. 2010;4(6):477- 83
18. Patel P, Nagar A, Patel R, Rathod D, Patel V. In vitro Anticancer Activity of *Rubia Cordifolia* against Hela and Hep2 Cell Lines. *Int J Pharm Pharm Sci*. 2011;3(2):70-1.
19. Long G, Wang G, Ye L, Lin B, Wei D, Liu L, Yang L. Important Role of TNF- $\alpha$  in Inhibitory Effects of *Radix Sophorae flavescentis* Extract on Vascular Restenosis in a Rat Carotid Model of Balloon Dilatation Injury. *Planta Medica*. 2009;75(12):1293-9
20. Shinozaki Y, Fukamiya N, Fukushima M, Okano M, Nehira T, Tagahara K, Zhang SX, Zhang DC, Lee KH. Dantaxusins C and D, Two Novel Taxoids from *Taxus yunnanensis*. *J Nat Prod*. 2002;65(3):371-4
21. He X, Liu RH. Cranberry phytochemicals: Isolation, structure elucidation, and their antiproliferative and antioxidant activities. *J Agric Food Chem*. 2006;54(19):7069-74.
22. Kondo M, MacKinnon SL, Craft CC, Matchett MD, Hurta RA, Neto CC. Ursolic acid and its esters: occurrence in cranberries and other *Vaccinium* fruit and effects on matrix metalloproteinase activity in DU145 prostate tumor cells. *J Sci Food Agric*. 2011;91(5):789-96.
23. Rajeswara Rao, B.R, Singh, K., Sastry, K.P., Singh, C.P., Kothari, S.K., Rajput, D.K. and Bhattacharya, A.K. (2007) „Cultivation Technology for Economically Important Medicinal Plants“, in Reddy, K.J., Bahadur, B., Bhadracharya, B. and Rao, M.L.N. (ed.) *Advances in Medicinal Plants*. Hyderabad: University Press, pp. 112-122
24. Azmi, A.S., Bhat, S.H., Hanif, S. and Hadi, S.M. (2006) „Plant polyphenols mobilize endogenous copper in human peripheral lymphocytes leading to oxidative DNA



- breakage: A putative mechanism for anticancer Properties", FEBS Letters, 580, pp. 533-538.
25. Apostolou, A., Stagos, D., Galitsiou, E., Spyrou, A., Haroutounian, S., Portesis, N., Trizoglou, I., Hayes, A.W., Tsatsakis, A.M. and Kouretas, D. (2013) „Assessment of polyphenolic content, antioxidant activity, protection against ROS-induced DNA damage and anticancer activity of Viti vinifera stem extracts“, Food and Chemical Toxicology, 61, pp. 60-68.
  26. Cao, J., Xia, X., Chen, X., Xiao, J. and Wang, Q. (2013) „Characterization of flavonoids from Dryopteris erythrosora and evaluation of their antioxidant, anticancer and acetylcholinesterase inhibition activities“, Food and Chemical Toxicology, 51, pp. 242-250.
  27. Agati, G., Azzarello, E., Pollastri, S. and Tattini, M. (2012) „Flavonoids as antioxidants in plants: Location and functional significance“, Plant Science, 196, pp. 67-76.
  28. Huntely, A.L. (2009) „The health benefits of berry flavonoids for menopausal women: Cardiovascular disease, cancer and cognition“, Maturitas, 63, pp. 297-301.
  29. Wen, L., Wu, D., Jiang, Y., Prasad, K.N., Lin, S., Jiang, G., He, J., Zhao, M., Luo, W. and Yang, B. (2014) „Identification of flavonoids in litchi (Litchi chinensis Soon.) leaf and evaluation of anticancer activities“, Journal of Functional Foods, 6, pp. 555-563
  30. Bishop, G.J and Koncz, C. (2002) „Brassinosteroids and Plant Steroid Hormone signaling“, The Plant Cell, supplement 2002, pp. 97-110.
  31. Malíková, J., Swaczynová, J., Kolář, Z. and Strnad, M. (2008) „Anticancer and antiproliferative activity of natural brassinosteroids“, Phytocchemistry, 69, pp. 418-426.
  32. Steigerová J., Oklešťková, J., Levková, L., Kolář, Z. and Strnad, M. (2010) „Brassinosteroids cause cell cycle arrest and apoptosis of human breast cancer cells“, ChemicoBiological Interactions, 188, pp. 487-496.
  33. Steigerová J., Rárová, L., Oklešťková, J., Křížová, K., Levková, M., Šváchová, M., Kolář, Z. and Strnad, M. (2012) „Mechanisms of natural brassinosteroid-induced apoptosis of prostate cancer cells“, Food and Chemical Toxicology, 50, pp. 4068-4076
  34. Cornblatt, B.S., Ye, L., Dinkova-Kostova, A.T., Erb, M., Fahey, J.W., Singh, K., Chen, M.A., Stierer, T., GarrettMayer, E., Argani, P., Davidson, N.E., Talalay, P., Kensler, T.W. and Visvanathan, K. (2007) „Preclinical and clinical evaluation of sulforaphane for chemoprevention in the breast“, Carcinogenesis, 28 (7), pp. 1485-1490.
  35. Amin, A., Gali-Muhtasib, H., Ocker, M. and SchneiderStock, R. (2009) „Overview of Major Classes of Plant Derived Anticancer Drugs“, International Journal of Biomedical Science, 5 (1), pp. 1-11.
  36. Unnati, S., Ripal, S., Sanjeev, A. and Niyati, A. (2013) „Novel anticancer agents from plant sources“, Chinese Journal of Natural Medicines, 11 (1), pp. 0016-0023.

37. Phillipson, J.D. (1999) „Medicinal Plants“, Journal of Biological Education (Society of Biology), 31 (2), pp. 109
38. Bhatnagar, P., Pant A.B., Shukla, Y., Chaudhari, B., Kumar, P. and Gupta, K.C. (2015) „Bromelain nanoparticles protect against 7,12-dimethylbenz[a] anthracene induced skin carcinogenesis in mouse model“, European Journal of Pharmaceutics and Biopharmaceutics, 91, pp. 35-46
39. Zschocke, S., Rabe, T., Taylor, J.L.S, Jäger, A.K. and van Staden, J. (2000) „Plant part substitution – a way to conserve endangered medicinal plants?“, Journal of Ethnopharmacology, 71, pp. 281-292.
40. Parveen, S., Jan, U. and Kamili, A. (2013) „Importance of Himalayan medicinal plants and their conservation strategies“, Australian Journal of Herbal Medicine, 25 (2), pp. 63-67
41. Jemal. A., Siegel, R., Xu, J. and Ward, E. (2010) „Cancer Statistics, 2010“, CA: A Cancer Journal for Clinicians, 60, pp. 277-300.
42. Cancer Research UK (2014) World cancer statistics. Available at: <http://www.cancerresearchuk.org/cancerinfo/cancerstats/world/> (Accessed 23 January 2015)