**NATURAL PRODUCTS AND THEIR THERAPEUTIC ADVANCEMENT IN MANAGEMENT OF CANCER**

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**Abstract**

The role of natural medicinal compounds in cancer management has gained significant attention due to their diverse therapeutic effects. Derived from plant sources, these compounds offer a potential alternative or complementary approach to conventional cancer treatments. Natural medicinal compounds possess antioxidant and anti-inflammatory properties, targeting oxidative stress and inflammation, which are key factors in cancer initiation and progression. Their multifaceted mechanisms of action, such as modulating cell signaling pathways, inducing apoptosis, and inhibiting angiogenesis, make them promising candidates for targeted cancer therapy. Additionally, these compounds may address drug resistance issues by targeting multiple pathways simultaneously. Importantly, their potential to mitigate treatment-induced side effects offers a significant advantage in enhancing patients' quality of life. However, challenges remain in terms of bioavailability, standardization, and clinical validation. Clinical trials are essential to establish their safety and efficacy, allowing these compounds to be integrated into comprehensive cancer treatment strategies. The role of natural medicinal compounds in cancer management holds promise as a holistic and personalized approach, potentially reshaping the landscape of cancer therapy.

**Keywords:** Natural medicinal compounds, Cancer management, Therapeutic effects, Mechanisms of action, Antioxidant and anti-inflammatory, Drug resistance and Clinical validation.

# Introduction

Cancer, in essence, refers to a group of diseases characterized by uncontrolled and abnormal cell growth within the body. It is a complex and multifaceted condition driven by genetic mutations that disrupt the normal regulation of cell division and function. This disruption leads to the formation of a mass of tissue, known as a tumor, which can invade surrounding tissues and even spread to distant parts of the body through a process called metastasis. The pathophysiology of cancer involves a series of events that start at the genetic level. Mutations can accumulate due to various factors, including genetic predisposition, exposure to carcinogens like tobacco smoke and UV radiation, and errors in DNA replication. These mutations can activate oncogenes, which promote cell growth, or inactivate tumor suppressor genes, which normally regulate cell division and prevent unchecked growth (Dyondi et al., 2015; Moutia et al., 2018; S. and A.J., 2012).

As these genetic alterations accumulate, affected cells lose their ability to respond to the body's natural signals that control growth and death (apoptosis). Consequently, they evade programmed cell death and continue to divide uncontrollably, forming a mass of abnormal cells – the tumor. Tumors can be either benign or malignant. Benign tumors are non-invasive and usually pose minimal threats, whereas malignant tumors are invasive and have the potential to spread to other parts of the body. Metastasis is a critical phase in cancer progression. Malignant cells can invade nearby tissues and enter the bloodstream or lymphatic system, allowing them to travel to distant sites in the body. Once there, they can establish secondary tumors, further compromising organ function and overall health (Jawaid et al., 2020; Paccagnella et al., 1996).

Understanding the pathophysiology of cancer has led to the development of various treatment strategies. Traditional approaches include surgery to remove tumors, radiation therapy to target and destroy cancer cells, and chemotherapy to kill rapidly dividing cells. More recent advancements involve targeted therapies that specifically target the molecular abnormalities driving cancer growth and immunotherapies that stimulate the body's immune system to recognize and eliminate cancer cells. Moreover, cancer's pathophysiology revolves around genetic mutations that disrupt normal cell regulation, leading to uncontrolled growth, invasion, and potential metastasis. A comprehensive understanding of these processes is crucial for developing effective treatments and interventions to combat this challenging disease .

Medicinal plants and natural products have played a significant role in cancer management and treatment due to their diverse bioactive compounds and potential therapeutic properties. These resources offer a unique avenue for developing novel therapies that complement conventional treatments. The role of medicinal plants and natural products in cancer can be understood through several key aspects. Firstly, many plant-derived compounds possess potent anti-cancer properties. For instance, compounds like curcumin from turmeric, resveratrol from grapes, and epigallocatechin gallate (EGCG) from green tea have demonstrated anti-inflammatory, antioxidant, and anti-proliferative effects, inhibiting various stages of cancer development and progression (Kaneko et al., 2019; Patel et al., 2012; Zhumashova et al., 2019).

Secondly, these natural compounds often exhibit fewer side effects compared to traditional chemotherapy drugs, making them attractive candidates for supportive care during cancer treatment. They can help alleviate treatment-related symptoms like nausea, fatigue, and pain, enhancing the overall quality of life for cancer patients. Thirdly, medicinal plants and natural products have shown promise in enhancing the effectiveness of standard cancer therapies. Some compounds sensitize cancer cells to radiation or chemotherapy, making them more susceptible to treatment. Others inhibit the growth of blood vessels that supply nutrients to tumors, a process crucial for tumor survival. Furthermore, these natural products often have multi-targeted effects, impacting various signaling pathways and cellular processes involved in cancer. This can potentially address the issue of tumor heterogeneity and reduce the likelihood of developing resistance to treatment (Arcos-Martínez et al., 2016; Mishra and Kaur, 2013; S. and A.J., 2012; Tan et al., 2010).

Despite these promising attributes, it's important to note that the development of medicinal plant-derived therapies for cancer is not without challenges. Standardization of dosages, ensuring consistent quality of plant extracts, and conducting rigorous clinical trials to validate their efficacy and safety are critical steps in their integration into mainstream cancer care. Furthermore, medicinal plants and natural products offer a valuable resource in the fight against cancer. Their bioactive compounds hold potential for preventing, managing, and treating cancer by exerting anti-cancer effects, minimizing side effects, and synergizing with conventional therapies. However, further research is necessary to unlock their full potential and establish their role as effective adjuncts in cancer treatment regimens (Furman et al., 2019; Kumar et al., 2021; Sohn et al., 2010; Tan et al., 2010).

# Review finding

## Role of medicinal plants and their necessity in management of cancer

Medicinal plants have been integral to traditional systems of medicine for centuries, and their role in the management of cancer within these systems is both significant and rooted in ancient wisdom. Traditional systems of medicine, such as Ayurveda, Traditional Chinese Medicine (TCM), and Indigenous knowledge, recognize the value of medicinal plants in the holistic approach to cancer management. In these traditional systems, medicinal plants are often prescribed based on a personalized assessment of an individual's constitution, imbalances, and the specific nature of the disease. These practices consider the interconnectedness of the body, mind, and spirit – a perspective that aligns with the modern understanding of the mind-body connection and psychosocial aspects of cancer (Mishra and Kaur, 2013; Sabir et al., 2019).

Medicinal plants used in traditional systems offer a diverse array of bioactive compounds that have shown anti-cancer properties. For instance, Ayurvedic formulations often include herbs like Ashwagandha (*Withania somnifera*) and Turmeric (*Curcuma longa*), which possess immunomodulatory, anti-inflammatory, and antioxidant effects, supporting the body's natural defense mechanisms against cancer (Al-Snafi, 2017; Roshan et al., 2014; Russo et al., 2001).

Additionally, traditional systems emphasize the importance of a balanced lifestyle, including diet, exercise, and stress reduction. Medicinal plants are incorporated into dietary practices to enhance overall well-being and resilience, factors crucial for cancer prevention and management. The necessity of medicinal plants in the management of cancer within traditional systems lies in their holistic approach. These systems consider not only the physical aspects of the disease but also the emotional, mental, and spiritual dimensions. This comprehensive perspective recognizes the interconnectedness of various factors contributing to cancer development and progression (Bhandari and Kamdod, 2012; Majdalawieh and Carr, 2010; Sharma et al., 2009).

Moreover, for many communities around the world, traditional systems of medicine are more accessible and culturally relevant. Medicinal plants are often an affordable and locally available resource, making them a practical option for populations with limited access to modern healthcare. While traditional systems of medicine offer valuable insights, it's important to integrate them judiciously with evidence-based modern medical approaches. Scientific research can help validate the efficacy and safety of these plants, standardize their usage, and refine their application for optimal cancer management. However, medicinal plants play a crucial role in cancer management based on traditional systems of medicine. Their incorporation is rooted in a holistic understanding of health and disease, and they offer a complementary approach to modern medical practices. Embracing the wisdom of traditional systems, while advancing their integration through research and collaboration, holds promise for a more comprehensive and effective approach to cancer care (Bhandari and Kamdod, 2012; Saini et al., 2022; Yahfoufi et al., 2018; Yue et al., 2010).

**Table 1:** Reported medicinal plants used as anti-cancer agents.

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **S. No.**  | **Common Name** | **Sources** | **Family** | **Pharmacology** | **Mechanism of Action** |
|  | Turmeric | *Curcuma longa* | Zingiberaceae | Curcumin exhibits anti-inflammatory, antioxidant, and anti-cancer properties. | Inhibits NF-κB and COX-2 pathways, induces apoptosis. |
|  | Green Tea | *Camellia sinensis* | Theaceae | EGCG possesses antioxidant, anti-proliferative, and pro-apoptotic effects. | Modulates multiple signaling pathways, including PI3K/Akt. |
|  | Ashwagandha | *Withania somnifera* | Solanaceae | Withaferin A has anti-inflammatory, immunomodulatory, and apoptotic activities. | Inhibits NF-κB, induces ROS-mediated apoptosis. |
|  | Garlic | *Allium sativum* | Amaryllidaceae | Allicin exhibits antioxidant, anti-inflammatory, and apoptotic effects. | Triggers mitochondrial apoptotic pathway. |
|  | Ginkgo Biloba | *Ginkgo biloba* | Ginkgoaceae | Ginkgolides have antioxidant, anti-inflammatory, and anti-tumor activities. | Inhibits angiogenesis and promotes apoptosis. |
|  | Ginkgo Biloba | *Ginkgo biloba* | Ginkgoaceae | Ginkgolides have antioxidant, anti-inflammatory, and anti-tumor activities. | Inhibits angiogenesis and promotes apoptosis. |
|  | Resveratrol | *Vitis vinifera* | Vitaceae | Exhibits antioxidant, anti-inflammatory, and anti-tumor effects. | Activates SIRT1, inhibits NF-κB, and induces apoptosis. |
|  | Curcumin | *Curcuma longa* | Zingiberaceae | Shows anti-inflammatory, antioxidant, and anti-proliferative effects. | Modulates Wnt/β-catenin, PI3K/Akt, and MAPK pathways. |
|  | Astragalus | *Astragalus membranaceus* | Fabaceae | Astragalosides possess immunomodulatory and anti-tumor properties. | Enhances immune response and inhibits cell proliferation. |
|  | Cat's Claw | *Uncaria tomentosa* | Rubiaceae | Contains oxindole alkaloids with anti-inflammatory and anti-cancer effects. | Suppresses NF-κB and MMPs, induces apoptosis. |
|  | Reishi Mushroom | *Ganoderma lucidum* | Ganodermataceae | Contains polysaccharides with immunomodulatory and anti-tumor activities. | Enhances NK cell activity, inhibits angiogenesis. |
|  | Artemisinin | *Artemisia annua* | Asteraceae | Anti-malarial, anti-cancer, anti-inflammatory | Generates oxidative stress, induces apoptosis |
|  | Dandelion | *Taraxacum officinale* | Asteraceae | Antioxidant, anti-inflammatory, anti-cancer | Induces apoptosis, inhibits cell proliferation |

* 1. **Reported phytochemicals used as anti-cancer**

Natural chemical constituents have gained significant attention for their potential role in anti-cancer activity due to their diverse pharmacological properties and intricate mechanisms of action. These compounds, derived from various plants and natural sources, exhibit promising effects in inhibiting cancer cell growth, inducing apoptosis (programmed cell death), and suppressing the development of tumors. Their multifaceted pharmacology is attributed to their ability to interact with key cellular pathways and molecular targets involved in cancer progression. One prominent example is curcumin, found in turmeric, renowned for its antioxidant and anti-inflammatory properties. Curcumin's mechanism of action involves modulating nuclear factor-kappa B (NF-κB) signaling, which plays a pivotal role in inflammation and cancer. By inhibiting NF-κB activation, curcumin disrupts the transcription of genes that promote cell survival, proliferation, and angiogenesis. Additionally, curcumin triggers apoptosis and activates tumor suppressor genes, offering a multi-pronged approach to combatting cancer.

Resveratrol, abundant in grapes, showcases antioxidant and anti-inflammatory characteristics. Its mechanism of action includes activating sirtuins (SIRT1) and inhibiting signal transducer and activator of transcription 3 (STAT3) signaling. These actions lead to cell cycle arrest and apoptosis induction, thereby impeding cancer cell proliferation. Furthermore, resveratrol's influence on multiple pathways offers potential for synergy in combination therapies.

Epigallocatechin gallate (EGCG) from green tea is recognized for its antioxidant and anti-angiogenic effects. EGCG's pharmacology involves disrupting the phosphoinositide 3-kinase (PI3K)/Akt pathway, which regulates cell survival and growth. By interfering with this pathway, EGCG suppresses cancer cell proliferation and promotes apoptosis. Its inhibition of angiogenesis further curbs tumor growth and metastasis. Quercetin, a flavonoid present in fruits and vegetables, demonstrates antioxidant and anti-inflammatory properties. Its mechanism revolves around inducing cell cycle arrest, particularly by targeting the PI3K/Akt pathway. By interfering with this pathway, quercetin halts cell cycle progression and encourages apoptosis. This dual action diminishes the survival and growth of cancer cells (Le Gall et al., 2004; Patel et al., 2012; Zhumashova et al., 2019).

Taxol, derived from the yew tree, disrupts microtubule dynamics and prevents mitosis, crucial for cell division. Taxol stabilizes microtubules, inhibiting their disassembly during cell division. This leads to cell cycle arrest and apoptosis, ultimately suppressing tumor growth (Singh et al., 2010; Tang et al., 2017). Natural compounds like vinblastine from the periwinkle plant and camptothecin from *Camptotheca acuminata* act similarly by disrupting microtubule dynamics and interfering with DNA replication, respectively. These mechanisms halt cancer cell division, underscoring their potential as anti-cancer agents (BALABHASKAR R et al., 2019; Loizzo et al., 2008).

One notable example is withaferin A, extracted from ashwagandha. Its adaptogenic properties and anti-inflammatory effects contribute to its anti-cancer potential. Withaferin A's mechanism involves disrupting microtubule dynamics, essential for cellular division, and inhibiting heat shock protein 90 (Hsp90), a molecular chaperone promoting cancer cell survival. Simultaneously, it activates apoptosis pathways and modulates NF-κB signaling, further curtailing cancer progression. Artesunate, primarily recognized as an anti-malarial agent, exhibits additional anti-cancer activity. Through its pro-oxidant effects, artesunate triggers oxidative stress in cancer cells, leading to DNA damage and apoptosis. It also interferes with angiogenesis and inhibits NF-κB, thus restraining tumor growth and metastasis. This dual action demonstrates its potential in diverse cancer contexts (Okoye et al., 2014; Verma and Kumar, 2018).

Berberine, derived from various plants, demonstrates pharmacological versatility. With its anti-inflammatory and antioxidant attributes, berberine modulates the AMP-activated protein kinase (AMPK) pathway, disrupting energy homeostasis in cancer cells. By impeding cell proliferation and inducing apoptosis, berberine exerts potent anti-cancer effects (Gahlot and Yadav, 2021; More and Kharat, 2016; Spelman, 2001).

Lycopene, commonly found in tomatoes, offers antioxidant properties and anti-inflammatory benefits. Its mechanism involves inhibiting the PI3K/Akt/mTOR pathway, responsible for cell survival and growth. Through this interference, lycopene induces apoptosis and halts cell cycle progression, limiting the expansion of cancer cells (Moody et al., 2020). Thymoquinone, a component of black seed oil, exhibits antioxidant and anti-inflammatory activities. Thymoquinone's mechanism is closely linked to p53-mediated pathways, known for their role in regulating cell cycle and apoptosis. By engaging these pathways, thymoquinone triggers apoptosis and restrains cancer cell proliferation (Hameed et al., 2019; Piras et al., 2013).

Genistein, a soy-derived isoflavone, acts as a phytoestrogen and exhibits anti-cancer effects. Its inhibitory action on tyrosine kinases disrupts cell signaling pathways, particularly those driving cell survival and proliferation. Genistein's modulation of cell cycle checkpoints and induction of apoptosis further contribute to its anti-cancer potential. These natural compounds showcase the intricate synergy between their pharmacological properties and mechanisms of action. While their antioxidant and anti-inflammatory attributes provide a foundation, their targeted interactions with signaling pathways governing cell fate are crucial in thwarting cancer progression. Their multi-faceted actions, from disrupting microtubule dynamics and inhibiting key enzymes to inducing oxidative stress and apoptosis, highlight their potential to impede cancer at various stages (Chen and Chen, 2021; Dar et al., 2015; Guo et al., 2006; Hämäläinen et al., 2007; Sapozhnikova, 2014).

Nonetheless, translating these insights into effective therapies requires rigorous scientific exploration, encompassing preclinical and clinical studies. Factors such as bioavailability, dosing, and potential interactions with conventional treatments must be thoroughly understood. Collaborative efforts between researchers, clinicians, and pharmacologists are pivotal in harnessing the full potential of these natural chemical constituents for anti-cancer strategies.

It is reported that the efficacy of these natural chemical constituents varies across cancer types and individuals. Their pharmacological properties, such as antioxidant and anti-inflammatory effects, contribute to their ability to counteract carcinogenesis. However, their mechanisms extend beyond these attributes, encompassing intricate interactions with cellular pathways that regulate cell survival, growth, and death.

**Table 2:** Reported phytochemicals used for management of cancer.

|  |  |  |  |
| --- | --- | --- | --- |
| **S. No.** | **Natural Compound** | **Pharmacology** | **Mechanism of Action** |
|  | Curcumin(from turmeric) | Anti-inflammatory,antioxidant | Inhibits NF-κB signaling, induces apoptosis,inhibits COX-2, activates tumor suppressor genes |
|  | Resveratrol(from grapes) | Antioxidant,anti-inflammatory | Activates SIRT1, inhibits STAT3 signaling,induces apoptosis |
|  | Epigallocatechin | Antioxidant, | Inhibits cell cycle progression, |
|  | gallate (EGCG)(from green tea) | anti-inflammatory | induces apoptosis, inhibits angiogenesis |
|  | Curcumin(from turmeric) | Antioxidant,anti-inflammatory | Inhibits mTOR signaling, activatesapoptosis, suppresses VEGF expression |
|  | Quercetin(from fruitsand vegetables) | Antioxidant,anti-inflammatory | Induces cell cycle arrest, inhibits PI3K/Akt pathway,activates apoptosis |
|  | Taxol(from yew tree) | Inhibits microtubuledynamics | Stabilizes microtubules, prevents mitosis |
|  | Vinblastine(from periwinkle) | Inhibits microtubuledynamics | Disrupts microtubule formation,inhibits cell division |
|  | Camptothecin(from Camptothecaacuminata) | Topoisomerase I inhibitor | Inhibits DNA replication and transcription |
|  | Paclitaxel(from yew tree) | Enhances microtubuleassembly | Stabilizes microtubules, inhibits mitosis |
|  | Allicin(from garlic) | Antioxidant,anti-inflammatory | Induces apoptosis through mitochondrial pathways,inhibits angiogenesis, modulates cell cycle |
|  | Silibinin(from milk thistle) | Antioxidant,anti-inflammatory | Inhibits STAT3 and NF-κB pathways,induces cell cycle arrest, apoptosis |
|  | Berberine(from various plants) | Antioxidant,anti-inflammatory | Activates AMPK, inhibits cell proliferation,induces apoptosis |
|  | Ellagic Acid(from berries andpomegranates) | Antioxidant,anti-inflammatory | Inhibits DNA binding of carcinogens,induces apoptosis, modulates cell cycle |
|  | Artesunate(from sweet wormwood) | Anti-parasitic agent,anti-inflammatory | Induces oxidative stress, inhibits angiogenesis,triggers apoptosis, inhibits NF-κB |
|  | Honokiol(from magnolia) | Anti-anxiety,antioxidant | Inhibits PI3K/Akt/mTOR pathway,induces apoptosis, anti-angiogenic effects |
|  | Withaferin A(from ashwagandha) | Adaptogenic,anti-inflammatory | Disrupts microtubule dynamics, inhibits Hsp90,induces apoptosis, modulates NF-κB |
|  | Lycopene(from tomatoes) | Antioxidant,anti-inflammatory | Inhibits PI3K/Akt/mTOR pathway,induces apoptosis, modulates cell cycle |
|  | Thymoquinone(from black seed) | Antioxidant,anti-inflammatory | Induces apoptosis through p53-mediated pathways,inhibits STAT3 and NF-κB pathways |
|  | Genistein(from soy) | Isoflavone,phytoestrogen | Inhibits tyrosine kinases, modulates cell cycle,induces apoptosis, anti-angiogenic effects |

# Discussion

The role of natural medicinal compounds in the management of cancer has garnered significant attention owing to their potential therapeutic effects. These compounds, derived from various plant sources, offer a promising avenue for cancer treatment and prevention due to their diverse pharmacological properties and mechanisms of action. While conventional treatments like chemotherapy and radiation remain crucial, natural medicinal compounds provide an alternative or complementary approach, addressing the limitations and challenges associated with traditional therapies. One of the key advantages of natural medicinal compounds is their rich source of bioactive compounds with potential anti-cancer properties. These compounds often possess antioxidant and anti-inflammatory attributes that can help counteract oxidative stress and chronic inflammation, both of which contribute to cancer initiation and progression. For example, curcumin, a component of turmeric, exhibits potent antioxidant and anti-inflammatory effects. Its ability to modulate key molecular pathways involved in inflammation and cell survival makes it a promising candidate for cancer therapy (Dyondi et al., 2015; Moutia et al., 2018; Quetglas-Llabrés et al., 2022).

Furthermore, the multifaceted mechanisms of action exhibited by natural medicinal compounds make them particularly appealing. Many of these compounds target specific signaling pathways that regulate cell proliferation, apoptosis, and angiogenesis. Resveratrol, found in grapes and red wine, is known to activate sirtuins and inhibit STAT3 signaling, resulting in cell cycle arrest and apoptosis induction. This intricate modulation of cellular processes allows these compounds to selectively target cancer cells while minimizing damage to healthy cells. Natural medicinal compounds can also address the challenge of drug resistance frequently observed in conventional cancer treatments. Their diverse mechanisms of action may overcome resistance mechanisms that cancer cells develop against single-target drugs. For instance, compounds like berberine and quercetin have shown the ability to modulate multiple pathways simultaneously, potentially reducing the likelihood of resistance development (Alali et al., 2021; Kaneko et al., 2019; Rajendran et al., 2013; Wallaschek et al., 2021).

Additionally, the potential of natural medicinal compounds to mitigate side effects is a notable advantage. Traditional cancer treatments often result in adverse effects due to their cytotoxic nature. Natural compounds like curcumin and green tea polyphenols have been studied for their protective effects against these side effects. Their ability to alleviate chemotherapy-induced toxicity while enhancing treatment efficacy is a promising attribute in improving the overall quality of life for cancer patients. However, challenges persist in the translation of natural medicinal compounds into mainstream cancer management. Issues related to bioavailability, standardization, and optimal dosing need to be addressed. Clinical trials are essential to validate their safety and efficacy, providing a solid scientific foundation for their use. Moreover, while these compounds hold immense potential, they are unlikely to replace conventional treatments entirely. Instead, they can be integrated as part of comprehensive treatment strategies, providing a holistic approach to cancer management (Levi-Polyachenko et al., 2009; S. and A.J., 2012; Sabir et al., 2019; Tang et al., 2017).

Moreover, the role of natural medicinal compounds in the management of cancer is a subject of growing interest and research. Their pharmacological diversity, multifaceted mechanisms of action, and potential to address challenges associated with conventional treatments make them valuable candidates for cancer therapy. However, a balanced approach is crucial, combining rigorous scientific investigation with clinical validation to harness their full potential. As we continue to unravel the complexities of cancer biology, these compounds offer a promising avenue for innovative and personalized approaches to cancer prevention and treatment, potentially reshaping the landscape of oncology (Mathiassen et al., 2017; Mosleh-Shirazi et al., 2022).

# **Conclusion**

In conclusion, the role of natural medicinal compounds in cancer management holds immense promise and warrants continued exploration. These compounds, sourced from diverse plant origins, offer multifaceted therapeutic effects that target key aspects of cancer initiation, progression, and treatment challenges. By harnessing their antioxidant, anti-inflammatory, and multi-pathway modulating properties, these compounds present an innovative approach to complement traditional cancer therapies. Their potential to mitigate drug resistance and minimize treatment-induced side effects underscores their value in enhancing overall patient well-being. However, successful integration of these compounds into mainstream cancer management requires rigorous scientific investigation, standardization, and robust clinical validation. As research advances, the synergy between natural medicinal compounds and conventional treatments may pave the way for more effective, personalized, and holistic approaches to cancer prevention and therapy, potentially reshaping the landscape of cancer care.

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**Conflict of interest**

The authors declare no conflict of interest.

**References**

Al-Snafi, P.D.A.E., 2017. The pharmacological and therapeutic importance of Eucalyptus species grown in Iraq. IOSR J. Pharm. 07, 72–91. https://doi.org/10.9790/3013-0703017291

Alali, M., Alqubaisy, M., Aljaafari, M.N., Alali, A.O., Baqais, L., Molouki, A., Abushelaibi, A., Lai, K.S., Lim, S.H.E., 2021. Nutraceuticals: Transformation of conventional foods into health promoters/disease preventers and safety considerations. Molecules. https://doi.org/10.3390/molecules26092540

Arcos-Martínez, A.I., Muñoz-Muñiz, O.D., Domínguez-Ortiz, M.Á., Saavedra-Vélez, M.V., Vázquez-Hernández, M., Alcántara-López, M.G., 2016. Anxiolytic-like effect of ethanolic extract of Argemone mexicana and its alkaloids in Wistar rats. Avicenna J. phytomedicine 6, 476–88. https://doi.org/10.22038/ajp.2016.6701

BALABHASKAR R, RAJENDRA KUMAR A, SELVARAJAN S, FARIDHA A, GAYATHRI GUNALAN, 2019. POTENTIAL NATURAL PRODUCTS WITH ANTICANCER PROPERTIES AND THEIR APPLICATIONS. Asian J. Pharm. Clin. Res. https://doi.org/10.22159/ajpcr.2019.v12i5.32817

Bhandari, P., Kamdod, M., 2012. Emblica officinalis (Amla): A review of potential therapeutic applications. Int. J. Green Pharm. https://doi.org/10.4103/0973-8258.108204

Chen, L.R., Chen, K.H., 2021. Utilization of isoflavones in soybeans for women with menopausal syndrome: An overview. Int. J. Mol. Sci. https://doi.org/10.3390/ijms22063212

Dar, A.A., Rath, S.K., Qaudri, A., Singh, B., Tasduq, S.A., Kumar, A., Sangwan, P.L., 2015. Isolation, cytotoxic evaluation, and simultaneous quantification of eight bioactive secondary metabolites from Cicer microphyllum by high-performance thin-layer chromatography. J. Sep. Sci. https://doi.org/10.1002/jssc.201500861

Dyondi, D., Sarkar, A., Banerjee, R., 2015. Joint surface-active phospholipid-mimetic liposomes for intra-articular delivery of paclitaxel. J. Biomed. Nanotechnol. https://doi.org/10.1166/jbn.2015.2061

Furman, D., Campisi, J., Verdin, E., Carrera-Bastos, P., Targ, S., Franceschi, C., Ferrucci, L., Gilroy, D.W., Fasano, A., Miller, G.W., Miller, A.H., Mantovani, A., Weyand, C.M., Barzilai, N., Goronzy, J.J., Rando, T.A., Effros, R.B., Lucia, A., Kleinstreuer, N., Slavich, G.M., 2019. Chronic inflammation in the etiology of disease across the life span. Nat. Med. https://doi.org/10.1038/s41591-019-0675-0

Gahlot, V., Yadav, D.K., 2021. Phytochemical and Network Pharmacology Based Evaluation of Antiepileptic Potential of Identified Metabolites in Argimone mexicana. Pharmacognosy Res. https://doi.org/10.5530/pres.13.4.13

Guo, T.L., Chi, R.P., Zhang, X.L., Musgrove, D.L., Weis, C., Germolec, D.R., White, K.L., 2006. Modulation of immune response following dietary genistein exposure in F0 and F1 generations of C57BL/6 mice: Evidence of thymic regulation. Food Chem. Toxicol. 44, 316–325. https://doi.org/10.1016/j.fct.2005.08.001

Hämäläinen, M., Nieminen, R., Vuorela, P., Heinonen, M., Moilanen, E., 2007. Anti-inflammatory effects of flavonoids: Genistein, kaempferol, quercetin, and daidzein inhibit STAT-1 and NF-κB activations, whereas flavone, isorhamnetin, naringenin, and pelargonidin inhibit only NF-κB activation along with their inhibitory effect on i. Mediators Inflamm. 2007. https://doi.org/10.1155/2007/45673

Hameed, S., Imran, A., Nisa, M. un, Arshad, M.S., Saeed, F., Arshad, M.U., Asif Khan, M., 2019. Characterization of extracted phenolics from black cumin (Nigella sativa linn), coriander seed (Coriandrum sativum L.), and fenugreek seed (Trigonella foenum-graecum). Int. J. Food Prop. https://doi.org/10.1080/10942912.2019.1599390

Jawaid, W., Nisa, Q., Umer, S.R., Barry, S.J., Qureshi, A., Shahbaz, N.N., 2020. Etiology and Types of Seizures in Patients Presenting to a Tertiary Care Hospital in Karachi: A Cross-Sectional Study. Cureus. https://doi.org/10.7759/cureus.9194

Kaneko, N., Kurata, M., Yamamoto, T., Morikawa, S., Masumoto, J., 2019. The role of interleukin-1 in general pathology. Inflamm. Regen. 39, 1–38. https://doi.org/10.1186/s41232-019-0101-5

Kumar, P., Mahato, D.K., Kamle, M., Borah, R., Sharma, B., Pandhi, S., Tripathi, V., Yadav, H.S., Devi, S., Patil, U., Xiao, J., Mishra, A.K., 2021. Pharmacological properties, therapeutic potential, and legal status of Cannabis sativa L.: An overview. Phyther. Res. 35, 6010–6029. https://doi.org/10.1002/ptr.7213

Le Gall, G., Colquhoun, I.J., Defernez, M., 2004. Metabolite Profiling Using 1H NMR Spectroscopy for Quality Assessment of Green Tea, Camellia sinensis (L.). J. Agric. Food Chem. 52, 692–700. https://doi.org/10.1021/jf034828r

Levi-Polyachenko, N.H., Merkel, E.J., Jones, B.T., Carroll, D.L., Stewart IV, J.H., 2009. Rapid photothermal intracellular drug delivery using multiwalled carbon nanotubes. Mol. Pharm. https://doi.org/10.1021/mp800250e

Loizzo, M.R., Tundis, R., Menichini, F., Saab, A.M., Statti, G.A., Menichini, F., 2008. Antiproliferative effects of essential oils and their major constituents in human renal adenocarcinoma and amelanotic melanoma cells. Cell Prolif. https://doi.org/10.1111/j.1365-2184.2008.00561.x

Majdalawieh, A.F., Carr, R.I., 2010. In vitro investigation of the potential immunomodulatory and anti-cancer activities of black pepper (Piper nigrum) and cardamom (Elettaria cardamomum). J. Med. Food. https://doi.org/10.1089/jmf.2009.1131

Mathiassen, S.G., De Zio, D., Cecconi, F., 2017. Autophagy and the cell cycle: A complex landscape. Front. Oncol. 7, 1–31. https://doi.org/10.3389/fonc.2017.00051

Mishra, R., Kaur, G., 2013. Aqueous Ethanolic Extract of Tinospora cordifolia as a Potential Candidate for Differentiation Based Therapy of Glioblastomas. PLoS One 8. https://doi.org/10.1371/journal.pone.0078764

Moody, L., Crowder, S.L., Fruge, A.D., Locher, J.L., Demark-Wahnefried, W., Rogers, L.Q., Delk-Licata, A., Carroll, W.R., Spencer, S.A., Black, M., Erdman, J.W., Chen, H., Pan, Y.X., Arthur, A.E., 2020. Epigenetic stratification of head and neck cancer survivors reveals differences in lycopene levels, alcohol consumption, and methylation of immune regulatory genes. Clin. Epigenetics. https://doi.org/10.1186/s13148-020-00930-5

More, N., Kharat, A., 2016. Antifungal and Anticancer Potential of Argemone mexicana L. Medicines. https://doi.org/10.3390/medicines3040028

Mosleh-Shirazi, S., Abbasi, M., Moaddeli, M.R., Vaez, A., Shafiee, M., Kasaee, S.R., Amani, A.M., Hatam, S., 2022. Nanotechnology Advances in the Detection and Treatment of Cancer: An Overview. Nanotheranostics. https://doi.org/10.7150/ntno.74613

Moutia, M., Habti, N., Badou, A., 2018. In Vitro and In Vivo Immunomodulator Activities of Allium sativum L. Evidence-based Complement. Altern. Med. https://doi.org/10.1155/2018/4984659

Okoye, T.C., Akah, P.A., Ezike, A.C., Uzor, P.F., Odoh, U.E., Igboeme, S.O., Onwuka, U.B., Okafor, S.N., 2014. Immunomodulatory effects of Stachytarpheta cayennensis leaf extract and its synergistic effect with artesunate. BMC Complement. Altern. Med. 14. https://doi.org/10.1186/1472-6882-14-376

Paccagnella, A., Favaretto, A., Oniga, F., Ossana, L., 1996. Paclitaxel and carboplatin: A phase I study in advanced non-small cell lung cancer, in: Seminars in Oncology.

Patel, D.K., Prasad, S.K., Kumar, R., Hemalatha, S., 2012. An overview on antidiabetic medicinal plants having insulin mimetic property. Asian Pac. J. Trop. Biomed. https://doi.org/10.1016/S2221-1691(12)60032-X

Piras, A., Rosa, A., Marongiu, B., Porcedda, S., Falconieri, D., Dessì, M.A., Ozcelik, B., Koca, U., 2013. Chemical composition and in vitro bioactivity of the volatile and fixed oils of Nigella sativa L. extracted by supercritical carbon dioxide. Ind. Crops Prod. 46, 317–323. https://doi.org/10.1016/j.indcrop.2013.02.013

Quetglas-Llabrés, M.M., Quispe, C., Herrera-Bravo, J., Catarino, M.D., Pereira, O.R., Cardoso, S.M., Dua, K., Chellappan, D.K., Pabreja, K., Satija, S., Mehta, M., Sureda, A., Martorell, M., Satmbekova, D., Yeskaliyeva, B., Sharifi-Rad, J., Rasool, N., Butnariu, M., Bagiu, I.C., Bagiu, R.V., Calina, D., Cho, W.C., 2022. Pharmacological Properties of Bergapten: Mechanistic and Therapeutic Aspects. Oxid. Med. Cell. Longev. 2022. https://doi.org/10.1155/2022/8615242

Rajendran, P., Rengarajan, T., Thangavel, J., Nishigaki, Y., Sakthisekaran, D., Sethi, G., Nishigaki, I., 2013. The vascular endothelium and human diseases. Int. J. Biol. Sci. https://doi.org/10.7150/ijbs.7502

Roshan, A., Verma, N.K., Gupta, A., 2014. A Brief Study on Carica Papaya- A Review. Int. J. Curr. Trends Pharm. Res.

Russo, A., Izzo, A.A., Cardile, V., Borrelli, F., Vanella, A., 2001. Indian medicinal plants as antiradicals and DNA cleavage protectors. Phytomedicine 8, 125–132. https://doi.org/10.1078/0944-7113-00021

S., G., A.J., K., 2012. A review of Candida species causing blood stream infection. Indian J. Med. Microbiol.

Sabir, F., Asad, M.I., Qindeel, M., Afzal, I., Dar, M.J., Shah, K.U., Zeb, A., Khan, G.M., Ahmed, N., Din, F.U., 2019. Polymeric nanogels as versatile nanoplatforms for biomedical applications. J. Nanomater. https://doi.org/10.1155/2019/1526186

Saini, R., Sharma, N., Oladeji, O.S., Sourirajan, A., Dev, K., Zengin, G., El-Shazly, M., Kumar, V., 2022. Traditional uses, bioactive composition, pharmacology, and toxicology of Phyllanthus emblica fruits: A comprehensive review. J. Ethnopharmacol. https://doi.org/10.1016/j.jep.2021.114570

Sapozhnikova, Y., 2014. Development of liquid chromatography-tandem mass spectrometry method for analysis of polyphenolic compounds in liquid samples of grape juice, green tea and coffee. Food Chem. https://doi.org/10.1016/j.foodchem.2013.10.131

Sharma, A., Chhikara, S., Ghodekar, S.N., Bhatia, S., Kharya, M.D., Gajbhiye, V., Mann, A.S., Namdeo, A.G., Mahadik, K.R., 2009. Phytochemical and Pharmacological investigations on Boswellia serrata. Pharmacogn. Rev.

Singh, G., Sharma, P.K., Dudhe, R., Singh, S., 2010. Biological activities of Withania somnifera. Ann. Biol. Res. 1, 56–63.

Sohn, S.H., Ko, E., Chung, H.S., Lee, E.Y., Kim, S.H., Shin, M., Hong, M., Bae, H., 2010. The genome-wide expression profile of Curcuma longa-treated cisplatin-stimulated HEK293 cells. Br. J. Clin. Pharmacol. https://doi.org/10.1111/j.1365-2125.2010.03724.x

Spelman, K., 2001. Traditional and clinical use of Tinospora cordifolia, Guduchi. Aust. J. Med. Herbal.

Tan, Y., Chiow, K., Huang, D., Wong, S., 2010. Andrographolide regulates epidermal growth factor receptor and transferrin receptor trafficking in epidermoid carcinoma (A-431) cells: Research paper. Br. J. Pharmacol. https://doi.org/10.1111/j.1476-5381.2009.00627.x

Tang, J., Ji, H., Ren, J., Li, M., Zheng, N., Wu, L., 2017. Solid lipid nanoparticles with TPGS and brij 78: A Co-Delivery vehicle of curcumin and piperine for reversing P-Glycoprotein-Mediated multidrug resistance in vitro. Oncol. Lett. https://doi.org/10.3892/ol.2016.5421

Verma, S., Kumar, V.L., 2018. Artesunate affords protection against aspirin–induced gastric injury by targeting oxidative stress and proinflammatory signaling. Pharmacol. Reports. https://doi.org/10.1016/j.pharep.2017.06.003

Wallaschek, N., Reuter, S., Silkenat, S., Wolf, K., Niklas, C., Kayisoglu, Ö., Aguilar, C., Wiegering, A., Germer, C.T., Kircher, S., Rosenwald, A., Shannon-Lowe, C., Bartfeld, S., 2021. Ephrin receptor A2, the epithelial receptor for Epstein-Barr virus entry, is not available for efficient infection in human gastric organoids. PLoS Pathog. https://doi.org/10.1371/JOURNAL.PPAT.1009210

Yahfoufi, N., Alsadi, N., Jambi, M., Matar, C., 2018. The immunomodulatory and anti-inflammatory role of polyphenols. Nutrients. https://doi.org/10.3390/nu10111618

Yue, G.G.L., Chan, B.C.L., Hon, P.M., Lee, M.Y.H., Fung, K.P., Leung, P.C., Lau, C.B.S., 2010. Evaluation of in vitro anti-proliferative and immunomodulatory activities of compounds isolated from Curcuma longa. Food Chem. Toxicol. 48, 2011–2020. https://doi.org/10.1016/j.fct.2010.04.039

Zhumashova, G., Kukula-Koch, W., Koch, W., Baj, T., Sayakova, G., Shukirbekova, A., Głowniak, K., Sakipova, Z., 2019. Phytochemical and Antioxidant Studies on a Rare Rheum cordatum Losinsk. Species from Kazakhstan. Oxid. Med. Cell. Longev. https://doi.org/10.1155/2019/5465463