

Importance of Epigallocatechin and its Health Benefits

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ABSTRACT

Natural products have a wide range of diversity of multidimensional chemical structures which play a vital role which show the important nature as golden source for achieving the herbal drug discovery. Literature survey was accomplished using multiple databases including PubMed, Science Direct, ISI web of knowledge and Google Scholar. Epigallocatechin-3-gallate is the most abundant tea polyphenol, followed by other polyphenols, namely, catechin, epicatechin, epicatechin-3-gallate and epigallocatechin. The most important pharmacological activities of EGCG are antineoplastic, HIV infection, hypertension and associated complications, type II diabetes mellitus, its usage as cardioprotective, hepatoprotective, nephroprotective and its application in Alzheimer, Parkinson and Osteoporosis. Natural products have played a key role in drug discovery and development in modern days.

Key words: Natural Compounds, Medicinal Plants, Phloretin, Epigallocatechin.

INTRODUCTION

Traditional herbal medicines have been considered as a source of curative remedy, because chemical components of plants are used to promote health and prevent diseases¹ and plants are invaluable sources of new drugs.¹⁻¹³ Epigallocatechingallate (EGCG), which is also known as epigallocatechin-3-gallate, a type of catechin and it is the ester of epigallocatechin and gallic acid. EGCG is the most abundant in tea and used in many dietary supplements and beneficial to affect human health and disease. The aim of this mini-review article is survey on the most important pharmacological benefits of Epigallocatechin.

EPIGALLOCATECHIN

Epigallocatechin-3-gallate is the most abundant tea polyphenol, followed by other polyphenols, namely, catechin, epicatechin, epicatechin-3-gallate and epigallocatechin.¹⁴ Epigallocatechingallate (EGCG), also known as epigallocatechin-3-gallate, is a polyphenolic flavonoid from tea (*Camellia sinensis*) possess various pharmacological activities such as anticancer, antimicrobial and antioxidant.¹⁵ Its neuroprotective effect against neural injuries and neurodegenerative diseases is also reported.¹⁶ It can also ameliorate protein and lipid damage induced by hepatotoxin, ethanol.¹⁷ EGCG carriers also appropriate for its wide application in food industry.¹⁸ The interaction mechanism of EGCG and natural α -glucosidase (SCG) can be beneficial to the development of functional foods to prevent diabetes,¹⁹ or develop active and environmental friendly packing materials for food industry.²⁰ Its importance in cancer treatment is because of natural origin, safety and low

cost, but EGCG challenge is its low bioavailability with various major limitations in EGCG studies which are study design, experimental bias, inconsistent results and reproductivity among different study cohorts.²¹ It can increase the potency of several chemotherapeutics such as doxorubicin, cisplatin and tamoxifen, *in vivo* and *in vitro*, in many cancers,²² and a suitable adjuvant to potentiate anti-glioma therapies.²³ EGCG may help in alleviating Methyl parathion-induced oocyte abnormalities.²⁴ Wu *et al.*²⁵ showed that epicatechingallate is a potential inhibitor of α -amylase and α -glucosidase, which indicates its importance as a nutrient supplement for the prevention of diabetes mellitus. Its fatty acid derivatives use for the prevention and treatment of viral infections.²⁶ Ling *et al.*²⁷ introduced EGCG as a novel and safe chemopreventive agent for influenza A infection. EGCG has found appropriate as cryopreservation procedures in stallions with low-quality sperm and possibly equine, to avoid or minimize DNA damages and preserve sperm plasma membrane integrity and mitochondrial activity.²⁸ EGCG may improve growth performance and alleviate the oxidant damage by modulating the antioxidant properties of broilers.²⁹ The most important pharmacological impacts of EGCG are presented in Table 1.

CONCLUSION

Epigallocatechingallate is a teacatechin. The most important pharmacological activities of EGCG are antineoplastic, HIV infection, hypertension and associated complications, type II diabetes mellitus,

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Table 1: The most important pharmacological effects of EGCG.

EGCG	Function	Reference
Autoxidation effect	<p>a. EGCG may lead to the formation of reactive oxygen species.</p> <p>b. Its mechanism is based on the formation of EGCG quinone, EGCG dimer quinone and other related compounds and the formation of autoxidized products may contribute to the inhibition of fibrillation.</p> <p>c. EGCG are found to form covalent adducts with cysteinylthiol residues in proteins through autoxidation to subsequently modulate protein function, which can be applied to treat human gastric cancer.</p> <p>d. The stability and autoxidation of EGCG are related to pH, temperature, metal ion, antioxidant levels, oxygen levels, concentration of EGCG and other ingredients in tea.</p>	30
Its effects for treatment of Alzheimers disease (AD), Parkinson's disease (PD) and Huntington's disease (HD)	<p>a. EGCG effects AD through oxidative stress, neurogenesis alteration and neuroinflammation.</p> <p>b. EGCG suppressed the production of Aβ and reduced inflammation, oxidative stress and cell apoptosis.</p> <p>c. EGCG can also increase in the ey autophagy adaptor proteins NDP52 and p62.</p> <p>d. EGCG reduce the Aβ levels by enhancing andogenous APP proteolysis and decrease nuclear translocation of c-Abl.</p> <p>e. EGCG induce an increase in the key autophay adaptor proteins NDP52 and P62.</p> <p>f. EGCG reulates the iron-export ferroporting in substantianigra, reduce oxidative stress and exert a neurorescue effect against 1-methyl-4-phenyl-1,2,3,6-tetrahydropyridine (MPTP)-induced functional and neurochemical deficits.</p> <p>g. AD, PD and HD are led by protein misfolding resulting from amyloid protein, which is a fibros polymer rich in β sheets, formed by the self-assembly of proteins of different sequences, structures and functions.</p>	31
Antitumor activity	<p>a. Introduction of 6-methoxycabonyl to Epigallocatechin-3-gallate (EGCG) is effective against gefitinib-resistant HCC827-Gef cells which can enhance its antitumor activities.</p>	32
Antiviral activity	<p>a. It has antiviral acitvities for many viruses such as human immunodeficiency virus (HIV), herpes simplex virus (HSV), influenza virus (Flu) and hepatitis C virus (HCV) by inhibiting the entry the virus into the host cell.</p> <p>b. EGCG can block the binding HIV-1 gp 120 to CD4 receptor and suppress the macrophage infiltration/activation in the rectal mucosa od macaques. Thus, EGCG can be considered as a novel, safe and cost-effective microbiocide for preventing sexual transmission of HIV-1.</p>	33
Antifibrosis effect	<p>a. EGCG can inhibit the activation and proliferation of hepatic stellate cells and synthesis of collagen, in a rat model.</p> <p>b. EGCG can reduce MMP-2 activity and possess the antifibrosis effect via downregulation of the expresson of MMP-2 mRNA.</p> <p>c. ECGCG can also inhibit the NF-κB to reverse the peritoneal fibrosis process.</p>	34,35
Antimicrobial activity	<p>a. Rinsing with EGCG solution may reduce the levels of mutans streptococci and lactobacilli in the oral cavity of children.</p> <p>b. It may increase the DPPH radical scavenging activity and the enzyme inhibitory activity against α-amylase and α-glucosidase which shows its antibacterial effects.</p> <p>c. The application of EGCG may play important roles in cell behavior and beneficial for regenerative endodontic therapy due to the antibacterial cross-linking agent of EGCG and differentiation of human dental pulp cells (hDPCs) cultured in collagen scaffolds.</p>	36
EGCC against fungi	<p>a. It has fungicialacticities against <i>Trichophyton mentagrophytes</i>, <i>T. rubrum</i>, <i>Cryptococcus neoformans</i> and <i>C. albicans</i>.</p> <p>b. It has activity against <i>T. mentagrophytes</i>.</p> <p>c. EGCG synergistically increase the antigunfal potential of azola drugs which maybe helpful in preventing the development of drug resistance, in lowering the drug dosage and minimizing adverse effects.</p>	37,38
EGCG against bacteria	<p>a. EGCG has antimicrobial effects against bacteria causing food-borne disease, the most recognized bacterium are <i>Escherichia coli</i>, <i>Helicobacter pylori</i>, <i>Bacillus stearothermophilus</i>, <i>Clostridium thermoaceticum</i>, <i>Helicobacter pylori</i>, <i>Salmonella typhi</i> and <i>Bacillus cereus</i>.</p>	39,40
EGC against viruses	<p>a. HCV, HIV1, HBV, HSV-1/HSV-2, EBV, Adenovirus, Influenza virus, Enterovirus.</p> <p>b. For HCV, inhibitory effect is viral entry by interference with binding to target cells.</p> <p>c. For HIV-1, inhibitory effects are inhibition of integrase, inhibition of RT, destruction of virions by bidning to envelope and binding CD4 and interference with gp 120 binding.</p> <p>d. For HBV, inhibitory effects are antigen expression, extracellular HBV DNA and cccDNA.</p> <p>e. For HSV-1/HSV-2, inhibitory effects are damage and inactivation of virions probably by binding to envelope proteins.</p> <p>f. For EBV, inhibitory effects are inhibition of transcription of immediate-early genes Rta, Zta and EA-D.</p> <p>g. For Adenovirus, inhibitory effects are inactivation of virus pariticles, inhibition of intracellular virus growth and viral protease.</p> <p>h. For influenza virus, inhibitory effects are alteration of physical integrity of virus particles, inhibiting of entry by binding to haemagglutinin.</p> <p>i. For enterovirus, inhibitory effects are suppression of viral replication via modulation of cellular redox milieu.</p>	41

Anti-inflammatory effect	a. EGCG may ameliorate OVA-induced airway inflammation by increasing the production of IL-10, the number of CD4 ⁺ CD25 ⁺ Foxp3 ⁺ Treg cells and expression of Foxp3 mRNA in the lung tissue and it can be recommend for treating asthma. b. EGCG inhibits the transfection of NF-κB and AP-1 to downregulate the expression of iNOS and COX-2 mainly by scavenging NO, peroxynitrite and other ROS/RNS and decreases the production of inflammatory factors to show the anti-inflammatory effects. c. EGCG may inhibit species and it has been pointed out that EGCG may inhibit the IL-8 production of respiratory passage epithelium cell, which may reduce the severity of respiratory passage inflammatory response.	42
Anti-angiogenesis	a. EGCG may inhibit tumor growth and angiogenesis which is possibly involved with the signaling intervention of MAPK/ERK1/2 and PI3K/AKT/HIF-1α/VEGF pathways which is supposed to be a potential therapeutic reagent for anti-angiogenesis treatment of solid tumors. b. The modification of the 3' position methylation of EGCG (MethylEGCG) may reduce cell growth effects at a low concentration <i>in vivo</i> . c. EGCG may prevent most of the IR-induced cellular and molecular events.	43,44
Antiobesity	a. Epigallocatechingallate binds with human peroxisome proliferator-activated receptors gamma (PPAR) gamma at its active site and block its activity and this mode of action may be helpful for antiobesity development.	45,46
Its effects against breast cancer	a. Peptide-conjugated formulations possess great cytotoxicity and great survivability and reduction in tumour volume in mice treated with peptide-conjugated shows EGCG as a novel drug delivery system in breast cancer therapy.	47
Anti Zika virus effect	a. EGCG may inhibit of ZIKV entry into the host cells	
Spermatogenesis activity	a. EGCG at the concentration of 5μM has influence on hBMSCs.	48

its usage as cardioprotective, hepatoprotective, nephroprotective and its application in Alzheimer, Parkinson and Osteoporosis. Its importance in cancer treatment is because of natural origin, safety and low cost, but the main problem is its low bioavailability with various major limitations in EGCG studies.

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CONFLICT OF INTEREST

The authors declare that they have no potential conflicts of interest.

ABBREVIATIONS

EGCG: Epigallocatechingallate; **SCG:** Natural α-glucosidase; **AD:** Alzheimer's disease; **PD:** Parkinson's disease; **HD:** Huntington's disease; **MPTP:** 1-methyl-4-phenyl-1,2,3,6-tetrahydropyridine; **HIV:** Human immunodeficiency virus; **HSV:** Herpes simplex virus; **HCV:** Hepatitis C virus; **hDPCs:** Human dental pulp cells; **Methyl EGCG:** Methylation of EGCG; **PPAR:** Peroxisome proliferator-activated receptors.

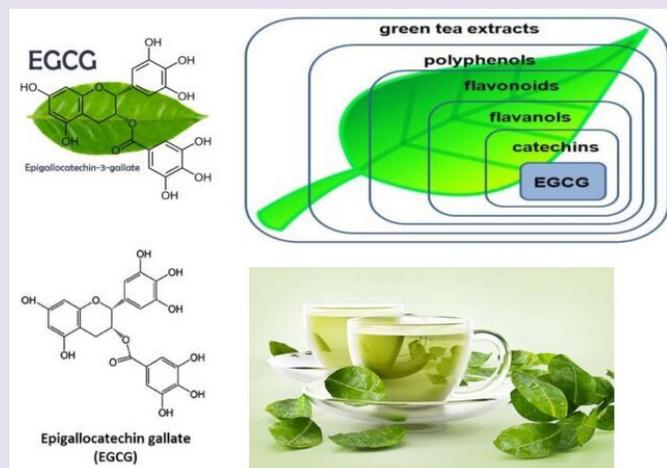
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GRAPHICAL ABSTRACT



SUMMARY

- Epigallocatechingallate is a tea catechin.
- The most important pharmacological activities of EGCG are antineoplastic, HIV infection, hypertension and associated complications, type II diabetes mellitus, its usage as cardioprotective, hepatoprotective, nephroprotective, and its application in Alzheimer, Parkinson, and Osteoporosis.
- Its importance in cancer treatment is because of natural origin, safety, and low cost, but the main problem is its low bioavailability with various major limitations in EGCG studies.

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Prof. Dr. Qi Cheng is a professor of Biotechnology and his researches have connected with agrobiotechnology. Presently, he is interested to traditional Chinese medicine and molecular researches.

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