

Nutritional and Biochemical Composition of Amla (*Emblica officinalis*) and its Therapeutic Impact: A Review

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Abstract

From ancient times, plants played a vital role in the development of mankind and was an exceptional source of natural medicine. The difficulty of the production of chemical medications, their side effects on safety, and the expense of rebellion have led researchers worldwide to concentrate on therapeutic plant science. Pakistan has a large collection of various plant species, with around 5,000 plants claiming to have important medicinal values. The papers published in recent decades on medicinal plants relate to the function of various plant bioactive composites commonly used in the cure of numerous human diseases. Bioactive compounds such as ascorbic acids, flavonoids, tannins, terpenoids, saponins, and several other components are reported to have been verified to have various medicinal activities, such as anti-inflammatory, antioxidant, antimicrobial, radiological, antitoxic, hepatoprotective, immune-modulatory, hypolipidemic and several other actions. The drug also has anti-cancer, antidepressant, antidiabetic, injury healing, antiulcerogenic, and so on. The present review article sums up the phytochemical components and pharmacology of the *Emblica officinalis* plant and its traditional application. Amla phyto-chemicals, including ellagic acid, emblicanin A, emblicanin B, Gallic acid, phyllanthin, quercetin and phyllanthidin were found to have various biological activities such as antioxidants, antimicrobials, anti-inflammatory, antidiabetic, anti-radiation protection, chemopreventive and healing of wounds.

Keywords: Amla; Applications; Traditional Uses and Medicinal Effects

Introduction

Emblica officinalis (EO) holds a sacred place in Ayurveda, an Indian indigenous medicine system. EO is the first tree grown in the world, according to Indian mythology. The other names of this tree are Amla, Indian Glassberry, or *Phyllanthus emblica*. It belongs to the family *Euphorbiaceae*. It is observed that intuitive species of

Indian amla grow in both tropical and subtropical zones of China, Pakistan, Sri Lanka, South Asia, and Malaysia. The plant, leaves and fruit of amla have been listed in the figure 1. Ayurveda believed that the fruit of EO increase defense against numerous diseases such as cancer, ulcer, anemia, heart diseases, liver, and diabetes, for that reason they extensively used EO fruits. It also acts as an antioxi-

dant, antitussive, gastroprotective, immunomodulatory, analgesic, and antipyretic. Besides, it is useful to control the cholesterol level and also boost memory in ophthalmic disorders. It is also beneficial in snake venom neutralization and as an antimicrobial.

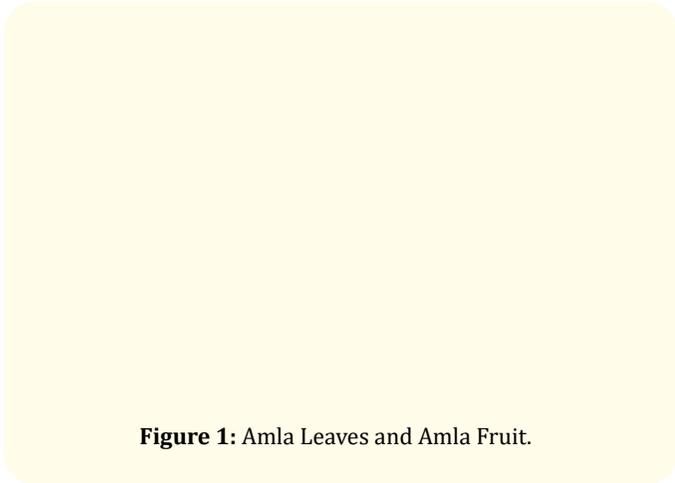


Figure 1: Amla Leaves and Amla Fruit.

Traditional uses

Amla considered a powerful Rejuvenator and valuable in delaying the senescence and degenerative process. Chinese, Sir Lankan, Siddha, and Unani Tibetan utilize *E. Officinalis* for medicine purposes. This helps improve lifespan, digestion, and relieve constipation. According to the Ayurvedic Medicine System EO significantly decreases fever, inflammation, blood cleansing, heart health improving, benefits the vision, stimulates hair growth, strengthens the body, relieves asthma, and improves the intellect ability. Fruits that are astringent used in many folk medicines for the treatment of ophthalmic disorders, hyperacidity, osteoporosis, gastritis, premature hair graying, dyspepsia, colitis, hemorrhoids, constipation, cough, anemia, hematuria, diabetes, asthma, and tiredness. *E. officinalis* probed to have anti-inflammatory, diuretic, laxative, cardioprotective, antipyretic, and hepatoprotective properties. EO also reported as a hair tonic and digestive medicine, also effective against peptic ulcer [1].

Chemical constituents of *Emblica officinalis*

It's challenging for drug industries to recognize the exact genotype of medical plants. Precincts of morphological and techniques to authentication have to need to create the latest protocols in botanical quality control. A DNA-based marker for EO and the Random Polymorphic Amplified DNA (RAPD) technique has

been developed to identify putative markers (1.1 kb) unique to EO recognition. Furthermore, in commercial samples of EO SCAR (sequence characterized amplified region) marker is considered to be valuable for the identification, and the SCAR marker is generated through using the RAPD application [2]. Phenolic compounds, amino acids, tannins, alkaloids, and carbohydrates are the major components of the EO. The highest level of vitamin C in fruit juice is (478.56 mg/100 ml) and when mixed with other fruit, the fruit improved its nutritional profile with vitamin C content [3]. Many types of beneficial compounds such as quercetin, ellagic acid, gallic acid, corilagin, chebulinic acid, 1,6-di-O-galloyl beta-D glucose, 3,6-di-O-galloyl-glucose, 3 ethylgallic acid (3 ethoxy 4,5 dihydroxy benzoic acid), 1-O-galloyl-beta-D-glucose, chebulagic acid, and isostrictiniin, isolated from it. In addition to, also possess some flavonoids including, kaempferol 3 O alpha L (6" ethyl) kaempferol and rhamnopyranoside 3 O alpha L (6 "methyl) rhamnopyranoside [4]. Similarly, a new acylated apigenin glucoside (apigenin 7 O (6 "butyryl beta glucopyranoside), along with the known compounds gallic acid, 1,2,3,4,6-penta-O-galloylglucose, methyl gallate, and luteolin-4'-Oneohesperidoside were also identified after the isolation from the methanol extract of the *Phyllanthus emblica* leaves [5]. A number of chemical components found in amla are listed in table 1.

Fruit pulp (Kumar, et al. 2012)	Leaves (Singh, et al. 2011)	Seed (Khan, 2009)
Moisture	Gallic Acid	Fixed Oil
Mineral	Chebulic Acid	Phosphatides
Crude Cellulose	Ellagic Acid	Essential oil
Albumin	Chebulinic Acid	
Gum	Amlic Acid	
Tannin	Alkaloids Phyllantine	
Gallic Acid	Phyllantidine	

Table 1: Chemical Constituent from Different Plant Parts of Amla.

Phytochemical

In recent years, many scientific kinds of literature have recorded different Phyto-constituents of *E. officinalis*. The amla fruit includes numerous bioactive components including isostrictiniin, ellagic acid, apigenin, chebulinic acid, quercetin, gallic acid, chebulagic acid. The tannins also found in the fruit extract of amla arepedun-

culagin, emblicanin A, phyllaemblicin B, emblicanin B and punigluconine. 100 g of edible fruit have been reported to be 470–680 mg of Vit. C. In ethanolic extracts of amla [9], recently identified Quercetin and β -sitosterol. Another study confirmed the existence of 5 major plants, including 5-hydroxymethylfurfural, 1, 2, 3-benzenetriol (synonym: Pyragallol), 2-acetyl-5-methylfuran. Recently two new chalconoid analogues, emblirol B [3] and emblirol A [2], have been isolated from the roots of emblica with a molecular formula C₁₉H₂₄O₆ [3]. The discovery of various phytochemicals in different sections of amla is very significant and understand its therapeutic role, along with mechanistic action in the attack against different disorders. Also, known phytochemicals can be docked for the understanding of their goals and related therapeutic activities.

Applications of *Emblica officinalis*

Role in cancer treatment

Earlier Triphala was considered to reveal some prospective for chemoprevention. The benzo-a-pyrene (B-a-P) responsible for papillomagenesis incidence in the fore-stomach of mice could be reduced through the addition of Triphala in the diet. It has become more active in decreasing tumor incidences relative to its components. The antioxidant level of animals which would have led to chemoprevention also increased substantially with Triphala [10]. Breast cancer considered one of the most communal cancers among females. Major factors associated with breast cancer were lipid metabolism enzymes, fats, and lipoproteins. Kalpaamruthaa (KA) is a improved preparation of Siddha which contains EO, honey and *Semecarpus Anacardium*. As treated by KA and SA, the raised levels of phospholipids, free fatty acids triglycerides, total cholesterol, free cholesterol, and the replacement of ester cholesterol in blood, liver, and kidney in animals with cancer have reversed to almost normal rates [11].

Chemoprevention with food phytochemicals is generally recognized to be one of the most vital cancer control methods. EO is known for its special flavonoids and tannins, which reveal very strong antioxidants properties. In Swiss albino mice, the inhibition of tumor incidence rate from this plant's fruit extract was tested on a two-stage cycle of skin carcinogenesis. Chemopreventive potential of the EO fruit extract was found on skin tumorigenesis induced by 7,12-dimethylbenz(a)anthracene (DMBA) in Swiss albino mice [12]. A significant polyphenol, gallic acid, observed in Triphala responsible for the reduction of cancer cell growth [13]. Experimentally,

EO ethanoic extract was probed for protection against DMBA induced genotoxicity. Before single intraperitoneal DMBA injection different concentrations (100, 250, 500 mg/kg) of EO fruit administered orally to Swiss albino mice for seven consecutive days which significantly reduced the frequency of bone marrow micronuclei. The anti-oxidant ability and modulative impact on detoxification of enzymes and hepatic activation are the main factors in EO's defense system [14].

Diabetic uses

Blood sugar levels fell significantly in healthy rats and alloxan (120 mg/kg) after oral administration (100 mg/kg body weight) within 4 hours. Production of diabetic cataract substantially decreased by EO and a supplemented fraction of its tannins in rats [15]. Aldose reductase (AR) engages in the growth of complications of secondary diabetes comprising cataracts. EO as a verified inhibitor of Aldose reductase. A significant approach to managing diabetic problems would be to examine the medicinal advantages of natural ingredients which would incorporate people into their everyday lives [16].

Impacts on the liver

Amla fruits reportedly have been used in Ayurveda [17], for hepatoprotection, and *Phyllanthus emblica* extract has been tested for hepatic rat damage due to ethanol [18]. The extent of hepatic fibrosis caused by thioacetamide and carbon tetrachloride was lessened by a hydroalcoholic (50%) extract of EO fruit (EO-50). Because of its antioxidant function, EO-50 effectively reverting profibrogenic events. EO-50 probed hepatoprotective role in antituberculosis drug-induced hepatic injury. Due to the perceived character of antioxidant, membrane stabilizer, and inhibitory CYP2E1, the EO-50 has hepatic protection activity [19]. Liver toxicity in Wistar rats also inhibited by EO [20]. EO and Chyayanaprash hepatoprotective functions were reviewed in rats with CCL causing liver damage. The extracts revealing reduced liver and serum peroxide, alkaline phosphatase (ALP), and glutamate-pyruvate transaminase (GPT) have been reported to suppress hepatotoxicity triggered by acute as well as chronic CCl₄ administration extract. The use of CCl₄ also induces hepatic fibrosis as revealed with elevated levels of pathological and collagen-hydroxyproline testing. The two extracts have been reported to significantly hinder these increased levels, suggesting that extracts can decrease the induction of fibrosis in rats [21].

The cardio-protective activity

The chronic effects of Amla's homogeneous and fresh fruit as oral administration on myocardial system antioxidant and ischemic-reperfusion injury (IRI) to the oxidative stress was tested in mice. Prolonged administration stimulates myocardial adaptation through swelling endogenous antioxidants as well as secures the heart of rats against IRI-associated oxidative stress [22].

Anti-ulcers characters

A herbomineral Ayurvedic formulation called Pepticare has been analyzed for its anti-ulcer and antioxidants in rats consisting of EO, *Tinospora cordifolia*, and *Glycyrrhiza glabra*. Peptic has anti-ulcer attitudes, due to its antioxidant properties [22]. EO extract (EOE) was tested for the treatment of ulcers. EOE has powerful curative and therapeutic impacts on the ulcer, which can affect both defense and offensive mucosal factors [23].

Antioxidant activities

Because of behavioral and biochemical abnormality changes due to cold stress, a study on EO has been carried out. The oral dispensation of Triphala by approximately one g/kg of the animal body until the 48-day period prevented cool stress-induced behavioral and biochemical abnormalities of albino rats. Hence the deemed shielding drug against stress is Triphala supplement [24]. In the aging rats, accelerated urea-nitrogen status is also suppressed by feeding Amla or Sun Amla ethyl acetate extract (EtoAc). Conspicuous lessening of thiobibetric acid-reactive elements volumes of serum, renal homogenate and mitochondria in old age mice are due to the oral intake of this extract. Thus urging that oxidative stress undergoing would be ameliorated by Amla. Moreover, EtoAc extracts from Amla or sun Amla extracts have substantially prevented up-grades in the COX-2 aorta of aged mice of nitric oxide synthase (iNOS) and cyclooxygenase. Restricting the NF-KappaB activation in the aged rats, the expression levels of COX-2 and iNOS are leveled down by the EtoAc extracts of Amla or Sun Amla. Ultimately for the prevention of age-associated renal ailments, Amla will come to be a very fruitful antioxidant [25]. Chashan the increase in renal gamma-glutamyl transpeptidase (GGT) activity triggered via hexachlorocyclohexane (HCH) was declined by pre-feeding Amla. Amla feeding was recognized to be an excellent part of a hepatic antioxidant mechanism and to deplete cytotoxic products which could be caressed by feeding HCH [26]. The rat's gut revealed for the entire body gamma radiation (WBI) was reported to have strengthened

the activity of xanthine oxidoreductase and decreased superoxide dismutase activity. However, the amount of sham irradiation was retrieved for 7 days when animals were treated with Triphala. Administration with Triphala recommended as the prevention from oxidative detriment caused by the exposure of the whole body to the radiation. The whole body of irradiated mice was shielded by Triphala. Protection was conciliated through repression of oxidative damage in cells and organs. The potential to thrive into a novel herbal radioprotector for practical application is signaled towards this drug [27]. Sedimentary extracts of the *Terminalia chebula*, EO, and *Terminalia bellerica* supplies and their combination of Triphala equi-portioned were tested for their in vitro antioxidant activities. The gamma-radiation promoting a strand break in plasmid DNA (pBR322) was considerably dwindled by the Triphala and its constituents. *Terminalia chebula*'s increased scavenging activities while its higher capability in lipid peroxidation and plasmid DNA testing is demonstrated by the EO, and their mixture Triphala is reckoned as more effective because of the incorporate activity of every single constituent [28].

Antipyretic and analgesic behavior

The extracts of EO are strengthened by strong antipyretic and analgesic action. There has been a substantial decrease in brewer's yeast hyperthermia in rats with just a single dose of ethanol extracts and aqueous extract (500mg/kg). Prominent inhibitory effects on acetic acid-induced writhing retort in mice were reduced by both of these extracts was revealed in analgesic test [29]. Such symptoms may have been caused by the involvement of alkaloids, tannins, amino acids, phenolic compounds, or carbohydrates.

Cytoprotective, gastroprotective and antitussive properties

Amla has been held responsible for its cytoprotective, chromium-induced immuno-modulatory effects of oxidative vandalism. Using the process of macrophage and phagocytosis gamma-IFN production was reinstated and chromium induced immunosuppression was obstructed by the EO. For the antitussive activity of EO in conscious rats, EO has been reviewed by mechanical stimulus of the air tracts tracheobronchial and laryngopharyngeal mucous region. Non-narcotic agent dropropizine was less effective in antitussive activity than that of EO but the classical narcotic antitussive drug, codeine is still at the highest efficiency for its antitussive activity. The antitussive activity shown by EO is presumed to be not only due to the antiphlogistic, anti spasmolytic and antioxidant

potency influences but also to its effects upon the mucous secretion in the airways [30]. EO is screened for antisecretory and counter-ulcer behaviors, using numerous experimental models in mice, such as pylorus ligation shay rats, indometacin hypothermic stress restriction triggered stomach ulcers, and necroticants. The exhibition of antisecretory, cytoprotective, and antiulcer properties by Amla was discovered then.

Memory enhancing effects

A dose-dependent increment of Amla fine powder is observed in the memory of juvenile and old rats. It was the transposition of amnesia caused by scopolamine and diazepam. Amla Churna is displayed to be gratifying healing for the treatment of Alzheimer's disease because of its multiple advantages including memory enhancement and memory arrears [31].

Management of ophthalmic conditions

Curcuma longa, *Terminalia bellerica*, *OE*, *cinnamomum*, *ocimum sanctum*, *acamphora*, *Medesumpapum* and *rosa damascena* have been integrated into the basic concepts of various herbs; *ophthacare* is an herbal eye-catcher. This herbal preparation was aimed at a clinical examination of those patients with various ophthalmic afflictions such as conjunctival xerosis, conjunctivitis, acute acryocytitis, and postoperative cataracts. The herbal eye drops primarily supplied the improved treatment for eye conditions. Patients well tolerated the eye drops and no side effects were ascertained for the time of the course of study. Ophthacare has shown its beneficial function in a variety of infectious, infective, as well as degenerative ophthalmic disorders [32].

Dyslipidemia and cholesterol reduction role

Amla was investigated on low-density lipids (LDL) oxidation as well as cholesterol-induced in Cu²⁺ by cholesterol and LDL-fed rats, and its impact was examined in vitro and in vivo. For the hypercholesterolemia and prevention of atherosclerosis. Amla was concluded as an effective specimen [33]. With the flavonoids that can curtail the volumes of lipids in serum and tissues of mice induced hyperlipidemia, both EO and *Mangifera* are embraced, each causing breakdown and vanishing of cholesterol [34].

Snake venom neutralizer

It was the first time scrutinized that EO and vitex negundo have the anti-snake venom activity. The plant extracts were antagonized

with both in vitro and in vivo models, *Naja kaouthia* and *Vipera Rossellini* venom. The V. Rossellini venom-induced coagulants, defibrinogenating hemorrhages, and inflammatory activities by these plant extracts were distinctly nullified. The strong neutralizing capacities of snake venom are seen in extracts of the plants and need even greater consideration in speculating no precipitation bands formation between snakelike venom and plant extract [35].

Antimicrobial and antimutagenicity activities

Due to its antimicrobial activities, EO has been divulged [36]. Against *Escherichia coli*, *Kozaena*, *Klebsiella pneumonia*, *Proteus mirabilis*, *Pseudomonas aeruginosa*, *S. paratyphi B*, and *Serratia marcescens*, the EO plant is endowed with puissant antibacterial activities [37]. For an antimutagenic effect with a TA 98 and TA 100 Salmonella Typhimurium Test strain, using a histidine reversal assay to test a phenobarbitone-induced hepatic s9 rat, Triphala's chloroform, water, and acetone extracts were sussed out against the direct-active mutagens, 4 nitrophenylenediamine (NPD) and sodium azide and indirect-active promoutagens, 2 Aminofluorene (2AF). The mutagenicity reticence of both direct and s9 – based mutagens in the chloroform and acetone extracts in a succession was evident [38].

Applications for the treatment of certain other diseases

Triphala, which contains one of the constituents in the EO cycle, is used to medicate the ailments including such as fever, constipation, chronic ulcers, anemia, asthma and jaundice. Secluded from Triphala, polyphenolic fractions possess the anti-mutagenic effects [39]. Furthermore, the active principles of Triphala were reviewed and considered as a peerless clinical intervention for infected wounds [40]. Aqueous plant extracts have been trialed on Swiss albino mice for their radio-protective effects against sub-lethal gamma radiation (9Gy). The 100mg/kg body weight against radiation discovered to be the most effective dose of fruit pulp extracts. Conspicuously, the survival time was exalted and the mortality rate of mice was lessened with this dose. Moreover, it was noticed that the loss of body weight in extract administrated irradiated animals was noticeably low as compared to animals who were given radiations only [41]. By prompting highly puissant hypolipidaemic and hypoglycemic activities maximum beneficial action is exhibited by the flavonoids derived from EO [42]. Furthermore, for elevating the hemoglobin level in rats, flavonoids are found to be very effective. Its being as anti-tumor in actions is also reported [43]. Against

thioacetamide (TTA) and CCL4 [44], induced changes in rat liver EO fruit has been examined. Abnormal histopathology indicative of pre-fibrogenic events was caused by the treatment with TTA and CCL4 [45]. Such alteration was transposed by EO showing its preventive capability in pre-fibrogenesis of the liver [46]. The extracts of *Withania* are the result of a repeatable dose-dependent, colony-forming inhibition in CHO cells [47]. Hypercholesteremia is one of the key factors in the case of coronary artery disease. Besides, it has been documented that Triphala formulations show hypolipidemic properties on the experimentally induced hypercholesteremic mice [48].

Conclusion

Since ancient times Amla or Indian gooseberry played an important role in conventional medicine, tribal, and Ayurveda medicines. Several biological and biopharmaceutical studies have been performed over the last few decades in a wide group of phytochemicals such as tannins, flavonoids, terpenoids, tannins, and others, derived from Amla. Amla phytochemicals, including ellagic acid, emblicanin A, emblicanin B, gallic acid, phyllanthin, quercetin, phyllanthidine, etc., were found to have various biological activities such as antioxidants, antimicrobials, anti-inflammatory, antidiabetic, anti-radiation protection, chemopreventive, healing of wounds, etc. The current research showed that some *Emblca officinalis* bioactive compounds are also common in other pharmaceutical plant species. Further assessment of unexplored bioactive Amla compounds is therefore required, which will expose the ever new bioactivities of this strong medicinal plant.

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Interest of Conflict

The authors declare no conflict of interest.

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