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**Review Article** 

# PHARMACOLOGICAL ACTIVITIES OF STEPHANIA GLABRA, WOODFORDIA FRUTICOSA AND CISSEMPELOS PAREIRA-A REVIEW

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#### ABSTRACT

Ayurveda is a profound and comprehensive system of health care that originated in India. This system endeavors to rationalize the all phenomena governing empirical experiences with natural products in medicine. Ayurvedic medicinal plants are validated with a view to bring their usage in the main stream. Ayurvedic medicare system has attained popularity at global level to replace the synthetic chemicals as they have shown less adverse reactions. One of the main branch of Ayurveda is 'Rasayana'. Rasayana therapy of Ayurveda is a dedicated stream of medication for immune promotive, antidegenerative and rejuvenative health care and is known for preventing the effects of ageing and improving the quality of life of healthy as well as diseased individuals. Numbers of plants have been mentioned in classical text of Ayurveda for the management of several diseases. Number of researchers have given different reports on the therapeutic activity of different kinds of Ayurvedic medicinal plants. Such studies should be brought in the knowledge of every concern man. The present study is therefore focused on the review of literature of pharmacological activities on three such Ayurvedic medicinal plants viz. Stephania glabra, Woodfordia fruticosa and Cissempelos pareira.

Keywords: Natural product, Stephania glabra, Woodfordia fruticosa, Cissempelos pareira

## INTRODUCTION

Plant-derived drug occupy an important place in both traditional and modern medicine. 1,2 Ethnopharmacological and ethnobotanical knowledge are percolating down to these days among the tribal population, but much of this information is empirical at best, and lacks preclinical scientific validations3,4. Chemical principles from natural sources have become much simpler and have contributed significantly to the development of new 'Rasayana' drugs from medicinal plants<sup>5</sup>. Biologically active compounds from natural sources have always been of great interest to scientists working on different diseases. Research to find out scientific evidence for claims of plants used for Indian Ayurvedic system of medicine has been intensified. Detailed research on the chemistry and pharmacology of therapeutic products of plant origin are much essential and this may eventually lead to the discovery of medicine that can be used in the treatment of several hazardous diseases<sup>5</sup>. Several plants like Oxalis corniculata as an antiinflammatory6, Dorstenia barteri as an anti arthritic7, Paeonia lactiflora as an antiarthritic8 and so many others has reported for their remarkable pharmacological activities. Numbers of pharmacological activities have been reported for plant Stephania glabra, Woodfordia fruticosa and Cissempelos pareira.

Stephania glabra is a large climbing shrub with greenish yellowish flowers and large tubers weighing as much as 30kg, it grows at

subtropical and temperate Himalayas from an altitude of 7000 ft from sea level from Sindh eastward and Khasia hills and Pegu<sup>9</sup>.

Woodfordia fruticosa (Lythraceae) is much branched, semi deciduous, undershrub or shrub up to 1-3 m high, rarely upto 3m and found in throught India, ascending to 1500m in Himalaya and also in the gangestic plains. Plant is known as Dhataki in Sanskrit, fire flame bush in English and dhai in Hindi. The Brhat gangadhara churna is an important formulation in the dose of 3-6g which has been specified in Ayurvedic pharmacopoeia<sup>10</sup>. Woodfordia fruticosa is an important ingredient of several Ayurvedic formulations. According to Ayurveda, 'Rasayana' from the plant, is used as an Atisera, Raktapitta, Trsna, varna, and visarpa<sup>10</sup>. Cissempelos pareira (Menispermaceae) is a climbing shrub, 2 - 5m high with a thickened root. Leaves have an orbicular shape 7. 14 cm in diameter. They are membranous or leathery, veined, glabrous to densely pilose. Flowers are green, male ones in spikes, 7 - 10cm long, with a little round leaflet at the base of every flower. Cissempelos pareira Linn. is a very variable, lofty, slender, dioecious, perennial, climber commonly distributed throughout topical and sub topical region of India, ascending up to an altitude of 2000 m and traditionally known as Laghupatha in Ayurveda, an Indian traditional system of medicine<sup>9,10</sup>. By considering the pharmacological importance of these plants the present study is focused on their review of literature.





Fig. 1: showing picture of Stephania glabra<sup>11</sup>

## PHARMACOLOGICAL ACTIVITY ON STEPHANIA GLABRA

## Folklore uses

In folklore the decoction of *Stephania glabra* roxb. (Menispermaceae) is used in the treatment of diabetes, fever, gastric problem, amoebic dysentery, as an anthelmintic in rheumatic bodyache, blood dysentery, in leprosy and even as anticancer drug<sup>12</sup>.

## **Antimicrobial**

Ethanol extract of *Stephania glabra* tubers has been evaluated for their antibacterial and antifungal activities against five bacterial species namely *Staphylococcus aureus* (along with ten hospital strains), *Staphylococcus mutans, Staphylococcus epidermidis, Escherichia coli, Klebsiella pneumonia* and six fungal species *Aspergillus niger, Aspergillus fumigatus, Penicillum citranum, Microsporum gypseum, Microsporum canis, Trichophyton rubrum,* obtained from different culture media. The extracts has been found to be active against most of the tested microorganism with MIC range of 50-100μg/ml<sup>13,14</sup>.

#### **Chemical Constituent**

Novel morphine alkaloid named gindarudine (structure1) was isolated from the ethanol extract of stephania glabra tubers together with four alkaloid palmatine, palmatine, dehydrocorydalmine and stepharanine. Gindarudine was elucidated as 3,6-0,N-detrimethyl-10-hydroxy-1-methoxy-thebaine by means of spectroscopic data including 2D NMR studies<sup>15</sup>. Glabradine tetrahydropalmitine (structure-2) also has been isolated from *Stephania glabra*.

## Hypotensive activity

The isolated alkaloids of Stephania glabra were reported for their hypotensive activity<sup>16</sup>.

## Analgesic and antipyretic

Analgesic and antipyretic activities of Gindarudine (GN) has been reported. The reason for this activity was found to be because of the close resemblance of Gindarudine to the structure of thebaine  $^{17}$ .

## Antihyperglycemic effect

Antihyperglycemic effect of Stephania glabra tubers in alloxan induced diabetic mice was reported by using different doses of ethanolic extract <sup>18</sup>.

## Antihistaminic activity

The in vitro H¹-receptor antagonist activity of methanolic extract of tuber of Stephania glabra was reported by employing *in vitro* screening models of guinea pig ileum and goat tracheal chain preparation. Goat isolated trachea and guinea pig ileum contracted to histamine in a dose-dependent manner while chlorpheniramine blocked this effect. The methanolic extract produced significant dose-dependent H¹-receptor antagonist activity by blocking histamine-induced contraction  $^{19}. \\$ 

#### Anthelmintic activity

The *in vitro* Anthelmintic efficacy of *Stephania glabra* along with two more medicinal plants from Northeast India against Hookworms, *Ancylostoma ceylanicum* has been reported. The concoction of rhizome pulp of *Stephania glabra* with areal roots of *Trichosanthum multiloba* and root tubers peel of *Flemingia vestita* were used. For this, adult parasites from intestine of goldenhamsters (*Mesocricentus aurants*) were collected and then exposed to various concentrations of crude extracts. These extracts at a concentration of 100mg/ml took less time for onset of paralysis in *A.ceylanicum*. Concentration of 100mg/ml of the concoction of rhizome pulp of *Stephania glabra* with areal roots of *T.multiloba* in the concentration ratio of 1:1 took lesser time for onset of paralysis. Whereas, 100mg/ml extract of peeled roots of F.vesta took very less time than the paralysis effect of mebandazole at same concentration<sup>20</sup>.

## PHARMACOLOGICAL ACTIVITY ON WOODFORDIA FRUTICOSA





Fig. 2: Showing picture of Woodfordia fruticosa. (Whole plant leaves and flowers.)<sup>21</sup>

## Antimicrobial activity<sup>22, 23, 24</sup>

Different extracts of dried flowers of W.fruiticosa have been reported for their significant antibacterial activity against fourteen human pathogen. Dried flowers were extracted by using five different solvents like ethanol, methanol, chloroform, petroleum ether and water. Out of the five extracts tested, the petroleum ether extract showed significant activity when compared to Gentamicin as standard drug<sup>22</sup>. The in vitro antibacterial activity of the crude methanolic extract of W.fruticosa flower has been reported by comparing it with standard drug ciprofloxacin using the agar well diffusion method. The methanolic extract has been reported to be most active against *Pseudomonas pseudoalcaligenes*. The methanolic extract was reported more effective against Gram negative bacteria as compare to Gram positive bacteria<sup>23</sup>.The petroleum ether, chloroform, diethylether and acetone extract of Woodfordia fruticosa leaves were evaluated against four bacterial species strains by using disc diffusion method. The extracts of petroleum ether, chloroform, diethyl ether and acetone were found to be effective against all the srains<sup>24</sup>.

## Chemical constituents

Different chemical constituents have been isolated from different parts of the plant. From the leaves and flowers, tannins, flavanoids, anthraquinon glycosides and polyphenols has been reported<sup>25</sup>.The presence of three dimeric hydrlyzable tannins viz. Woodfordin A, B and C along with seven known hydrolysable tannins including oenothein B has been reported in the plant26. An amount of isoschimawalin A is present in the tannin fraction of the plant. Essential oil of Woodfordia fruticosa was extracted by hydrodistillation. The main components present in the essential oil of leaves are sesquiterpenoids (β-caryophyllene, γ-curcumen, gerrmacrene-D, β-selinene, elemol) and monoterpenoids (αpinene 2, 6 dimethyl-1, 3, 5, 7 octatetraene) 27. Besides previously reported hydrolyzable tannin oligomers (woodfordins A-D), a new hydrolyzable tannin monomer (isoschimawalin A) and five oligomers (woodfordins E-I) have been isolated from the dried flowers of Woodfordia fruticosa (an Indonesian crude drug, Sidowaya), and their structures were elucidated on the basis of spectra and chemical evidence. Woodfordins G and H were characterized as dimers with structures related to woodfordin B. Woodfordins I, E and F were a macrocyclic dimer, trimer and tetramer, respectively28. Woodfruticosin (woodfordin C), a new cyclic dimeric hydrolyzable tannin having an inhibitory activity toward deoxyribonucleic acid (DNA) topoisomerase II, has been isolated from the leaves of Woodfordia fruticosa Kurz (Lythraceae) along with three known flavonol glycosides and three known flavonol glycoside gallates. The structure of woodfruticosin (woodfordin C) was determined by the use of two-dimensional nuclear magnetic resonance (2-D NMR) spectroscopy including heteronuclear multiple quantum coherence (HMQC) and heteronuclear multiple bond connectivity (HMBC) techniques<sup>29</sup>. The structure of Woodfordin 'C' has been depicted below (structure 3)

## Hepatoprotective activity

Woodfordia fruticosa posses significant hepatoprotective activity <sup>30</sup>, <sup>31,32,33,34</sup>. Hepatoprotective activity of petroleum ether, chloroform, ethyl alcohol and aqueous extract of the flower of Woodfordia fruticosa has been reported against carbon tetrachloride induced hepatotoxicity<sup>30</sup> and phenytoin induced liver damage in rats<sup>31</sup>. The methanolic extract of the flowers of Woodfordia fruticosa has been reported for hepatoprotective activity against acetaminophen induced hepatic injury in rats<sup>32</sup> and declofenac sodium induced hepatic damage in rats<sup>33</sup>.

## Cardioprotective activity

*Arjunarishta* (Parthadyarishta) is an important Ayurvedic formulation used for cardiovascular disorders and is prepared by fermenting the decoction of specified plant materials using flowers of  $Woodfordia\ fruticosa^{34}$ .

## Antioxidant activity

Antioxidant potential of *Woodfordia fruticosa* was confirmed by using DPPH and ABT free redial scavenging models<sup>35, 36, 37</sup>. For this solvent Petroleum ether, chloroform, methanol and water depending upon the increasing order of polarity were used as solvents for extraction. The extracts were tested for phytoconstituents. Solvents were subjected to preparatory HPLC. From IR LC-MS and NMR spectra gallic acid and ascorbic acids were found to be main constituents involved in free radicals scavenging activity of the plant. Among ascorbic acid and gallic acid the latter one is known to play pivotal role in this activity <sup>37</sup>.

## Role in Leukaemia

The effects and action mechanisms of woodfordin I, a macrocyclic ellagitannin dimer was investigated, on human chronic myelogenous leukemia (CML) cells. The results showed that woodfordin I was able to suppress the proliferation and induce apoptosis in CML cells. Apoptosis was evaluated by cytomorphology, internucleosomal DNA fragmentation, and externalization of phosphatidylserine. Woodfordin I treatment caused a rapid and sustained loss of mitochondrial transmembrane potential (MMP), transient generation of reactive oxygen species (ROS), transient elevation of intracellular Ca2+ concentration, and cytosolic accumulation of cytochrome c. The activation of caspase-9 and 3, but not caspase-8, was also demonstrated, indicating that the apoptotic signaling triggered by woodfordin I was mediated through the intrinsic mitochondria-dependent pathway<sup>38</sup>.

## Antiulcer activity 39, 40

The antiulcer potential of W.fruticosa has been reported in ethanol,  $HCL^{40}$  and NSAIDS (Diclofenac sodium)  $^{40}$  induced ulcer in stomach of female wistar albino rats. The roots were extracted with chloroform and methanol. Both the extracts have found to have significant antiulcer activity.

#### Immunomodulatory activity

The in vitro and in vivo immunomodulatoy activity of ethanolic extract of the flowers of Woodfordia fruticosa has been reported. For this the effect of non specific immune responses in mice was examined. *In vitro* immunomodulatory activity of the extract was examined on murine peritoneal macrophage phagocytosis (using nitroblue tetrazoleum dye lysosomal enzyme activity, nitric oxide reduction. myeloperoxidase) and on proliferation of bone marrow cells by salforhodamine 'B' (SRB) assay. The in vivo activity has shown on macrophages and bone marrow cells by using carbon clearance test and cyclophosphamide induced myelosupression respectively. The significant increase in the release of myeloperoxidase, nitric oxide, lysosomal enzyme and superoxide from macrophages along with significant increase in phagocytic index in carbon clearance test indicates stimulatory activity of the extract in macrophages. The extract was found to show 60% increased bone marrow cells proliferation and offer protection towards cyclophosphamide induced myelosuppression which represent the stimulation of bone marrow 41.

## **Antifirtility activity**

The antifirtility activity of various extract of dried flowers of *Woodfordia fruticosa* has been reported on female albino rats. The ethanolic extract of the powder of the dried flowers was prepared by extracting successively with petroleum ether, benzene, chloroform and ethanol and also extracted individually with 50% aqueous alcohol and water. Antifertility activity of successive alcoholic, individual aqueous and individual hydroalcoholic extracts was studied in female albino rats. The results revealed that the alcoholic extract posses significant abortifacient activity, whereas aqueous and hydroalcoholic extracts hold moderate activity as compared to the control. Thus, the successive alcoholic extract showed promising abortifacient activity at 100 mg/kg body weight 42.

## **Antitumor activity**

Woodfordin C, a macro-ring hydrolyzable tannin dimer from dried flower was reported to posses antitumor activity  $^{43}$ .

## PHARMACOLOGICAL ACTIVITY ON CISSEMPELOS PAREIRA





Fig. 3: Show the pictures of Cissempelos pareira.  $^{44}$ 

# Anti-inflammatory activity- $^{46,47,48,49,50}$

1) 50% ethanolic extract of *Cissempelos preira* roots was evaluated in acute, sub acute and chronic models of inflammation. Doses of 200 and 400 mg/kg (p.o.) have shown significant anti-inflammatory activity. The percentage inhibition of inflammation in rat by using different models to induce inflammation has been depicted in table  $1^{46}$ 

Table 1: Show the percentage inhibition of inflammation by using different models<sup>46</sup>

Percentage of Treatment in Acute Inflammation					
1.	Carrageenin induced inflammation	59.55 and 64.04%			
2.	Histamine induced	15.38 and 30.77%			
3.	5-HTP induced inflammation	17.78 and 31.11%			
4.	PGE induced inflammation	19.23 and 30.77%			
Percentage of Treatment in Sub-acute Inflammation					
1.	Formaldehyde induced inflammation	38.6 and 47.95%			
3. 5-HTP induced inflammation 17.78 and 31.11% 4. PGE induced inflammation 19.23 and 30.77%  Percentage of Treatment in Sub-acute Inflammation 1. Formaldehyde induced inflammation 38.6 and 47.95%  Percentage of Treatment in chronic Inflammation					
1.	Cotton pallet granuloma	15.02 and 19.19%			

- 2) 50% ethanolic extract of the aerial part of *Cissampelos pareira* (Menispermaceae) was tested for analgesic activity and anti-inflammatory by abdominal writhes and hot plate model in rats and mice, respectively. Oral administration of extract exhibited significant and dose dependent anti-inflammatory activity in the carrageenin test which was based on interference with prostaglandin synthesis as confirmed by the arachidonic acid test. In the abdominal writhing test induced by acetic acid higher dose of the plant extract had the highest analgesic activity. In the hot-plate test the best dose was reported as 100 mg/kg <sup>47,48</sup>.
- 3) Anti-inflammatory activity of the ethanolic extract of the Cissampelos pariera Linn leaves was studied in albino wistar rats

using the carrageenin induced rat paw edema model. The ethanolic extract of *Cissampelos pariera* (400 mg/kg p.o.) inhibited carrageenin induced rat paw edema. In the result significant anti-inflammatory activity when compared with the standard drug indomethacin (10 mg/kg p. o.) and untreated control was obtained<sup>49</sup>.

4) 50% aqueous extract of roots of *Cissempelos pareira* along with leaves of *Vitex nigundo* and *Pongamia pinnata* were evaluated for analgesic and anti-inflammatory activity on swiss albino mice and wistar rats of either sex. Activity was tested by using hot plate method, acetic acid induced writhing in mice, and carrageenin induced paw edema <sup>50</sup>.

## Antiarthritic activity

50% aqueous ethanolic extract of roots of Cissempelos pareira was reported to posses antiarthritic activity.  $^{50,51}$ 

## Antioxidant activity

The alkaloidal fraction of roots of *Cissampelos pareira was* screened for *in-vitro* antioxidant activity and immunomodulatoy activity in mice. The HPTLC finger print profile was also established for the identification of extract which was found to contain 0.176 % of berberine. The extract possess strong antioxidant activity which was revealed by its ability to scavenge the stable free radical DPPH, superoxide ion and to inhibit lipid peroxidation in rat liver homogenate induced by iron/ADP/Ascorbate complex. The alkaloid fraction was found to have significant immunosuppressive activity at lower doses (25 and 50 mg/kg) while no activity was observed at higher doses (75 and 100 mg/kg).

The extract also found to reduce dose dependently the gastrointestinal transit from 46.4 and 38.7%, equivalent to 53.6 and 61.3%. However, *C. pareira* significantly reduced the lipid peroxidation and inhibited the decrease in antioxidant enzyme

levels (superoxide dismutase and catalase) on prior administration to castor oil-induced fluid accumulation. The extract of  $\it C. pareira$  had no effect on normal defecation at 25 mg kg $^{-1}$  in mice. However, 50 and 100 mg kg $^{-1}$  inhibited defecation by 100% in the initial 2 h and the activity was reduced to 40.0 and 73.0%, respectively, in the third hour  $^{52}$ .

## Immunomodulatory activity

Different extracts of *Cissempelos pareira* has been evaluated by different researchers for its immunomodulatoy activity<sup>52, 53</sup>. Methanol extract of roots of *C. pareira* was screened for its immunomodulatoy activity in mice at five different dose levels. Stimulatory activity on DTH response was found at 200-800 mg/kg dose. Phagocytic index was also increased significantly as compared to control animals. Methanol extract was not reported to alter the humoral response to SRBCs and significantly indicated no activity on humoral immunity. Higher doses of extract also offered protection against cyclophosphamide induced myelosuppression by increasing total WBC count significantly<sup>53</sup>.

## Antiparasite activity

Cissampeloflavone was reported to possess significant activity against *Trypanosoma cruzei* and *T. bruceirhodiense* but against human KB cell line it showed less low toxicity<sup>53</sup>.

#### Chemical constituents

From the aerial parts of *C.pareira*, chalcone a flavone dimer was isolated and was named cissampeloflavone. Its structure was identified by NMR and MS<sup>54</sup>. In earlier investigations roots from Kashmere contained 0.33 % of alkaloids, mainly hyatin and bebeerine, 0.2 % essential oils, 3.4 % fixed oils and a sterol <sup>54</sup>. Furthermore from the roots of *C.pareira* the bisbenzoylisochinoline alkaloids hyatin, haytinin and haytidin were found. Bebeerines has been reported as main alkaloids . In the roots of *C.pareira* from Peru toxic bisbenzoylisochinoline alkaloids were investigated. Cissampareine was isolated as one of bisbenzoylisochinoline, and its chemical structure was reported as  $C_{37}$   $H_{38}$   $N_2$   $O_6$ <sup>54</sup>. *Cissempelos pareira* was found to contain a group of plant chemicals called isoquinoline alkaloids. Different types of benzylisoquinolne isolated from *C. pareira* have been depicted in Table No.2<sup>55-63</sup>.

Table 2: Show names and molecular formula of different constituents of C. pareira

S. No.	Name	Molecular Weight	Structure	Molecular formula
1.	Curine(Bebeerines)	594	5	$C_{36}H_{38}N_2O_6$
2.	Hayatine	594	6	$C_{36}H_{38}N_2O_6$
3.	Isochondodendrine	594	-	$C_{36}H_{38}N_2O_6$
4.	Cissamampareine	606	7	$C_{37}H_{38}N_2O_6$
5.	Hyatidinine	608	8	$C_{37}H_{40}N_2O_6$
6.	Hyatinine	608	9	$C_{37}H_{40}N_2O_6$
7.	4"-o-Methylcurine	608	10	$C_{37}H_{40}N_2O_6$
8.	Insularine	620	11	$C_{37}H_{40}N_2O_6$
9.	Cycleanine	662	12	$C_{38}H_{42}N_2O_6$

An amorphous white alkaloid 'pelosine' was studied in association with an indifferent body, deyamittin<sup>64, 65, 66</sup>. Cissempelosine was reported from *C.pareira* which was later on shortened as 'Pelosine' (structure 4).

Structure-6

Ayurvedic uses of Cissempelos pareira<sup>67</sup>has been depicted in table 3.

Table 3: shows Ayurvedic uses of C.pareira.67

Main Actions	Other Actions	Standard Dosage	
Stop bleeding	Kills bacteria		
Balance menstruation	Prevent convulsions	Decoction: 1 cup 2-3 times daily	
Relieves pain	Fights free radicals	Tincture: 2-3 ml 2-3 times daily	
Reduces spasm	Prevents ulcers	Capsules: 1-2 g 2-3 times daily	
Relaxes muscles	Reduces mucus		
Stop inflammation	Reduce fever		
Increase urination	Protects liver		
Lowers blood pressure	Balances hormones		

# Diuretic activity

The methanolic root extract of *Cissampelos pareira* was tested for diuretic activity and its moderate diuretic activity was reported<sup>68</sup>.

# Anti-hemorrhagic effects

To establish the antihemorrhagic activity of aqueous extract from leaves of *C. pareira*, the skin of mice were injected with a mixture of extract and venom, and extract was found to produced a total inhibition of this activity. Anti-proteolytic activity was also

conducted by observing the effect on casein in a test tube or on biotinylated casein in a microplate. Both procedures were unable to show any inhibitory activity  $^{69}$ .

## **Antifertility activity**

C. pareira leaf extract, when administered orally, altered the estrous cycle pattern in female mice. The extract was found to prolong the length of estrous cycle with significant increase in the duration of diestrus stage and reduced significantly the number of litters in albino mice.

The analysis of the principal hormones involved in estrous cycle regulation showed that the plant extract altered gonadotropin release (LH, FSH and prolactin) and estradiol secretion. The oral LD50 of the extract was found to be  $7.3~\rm g/kg$  in mice $^{70}$ .

## Chemo preventive effects

The protective effect of *C. pareira* extract was studied against benzopyrene induced gastric cancer in mice, and the tumor incidence was found to be reduced and the mean number of tumor and the tumor multiplicity were also reported to be reduced significantly. The modulatory effect of *C.pareira* extract was also examined on carcinogen metabolizing phase I and phase II enzymes, antioxidant enzymes, glutathione content, lactate dehydrogenase, and lipid peroxidation in liver. Significant increases in the level of acid-soluble sulfhydryl (-SH) and cytochrome P450 contents and in enzyme activities of cytochrome P450 reductase, cytochrome 5 reductase, GST, DTD, SOD, catalase, glutathione (GSH) peroxidase, and GSH reductase but decreased malondialdehyde (MDA) were observed<sup>71</sup>.

## **Toxicity**

In a acute toxicity test, oral administration of 2 g/kg of *C. pareira* was reported to produce no mortality, no changes in behavior or any other physiological activities in mice. In subacute toxicity studies, no mortality was observed when two doses of 1 or 2 g/kg day of 50% aqueous ethanolic extract of *C. pareira* were administered p.o. for a period of 28 days in rats. No significant change was observed in the blood chemistry analysis in both sexes of animals. Hematological analysis showed no marked differences in any of the parameters examined in the control or treated group of both sexes. Pathologically, no gross abnormalities and histopathological changes were observed<sup>72</sup>.

## CONCLUSION

The present study is focused on the review of literature on *Stephania glabra*, *Woodfordia fruticosa* and *Cissempelos pareira*. Literature from various journals has been reported here. From the above literature it can be concluded that these plants are rich in medicinally important components which are responsible for various pharmacological activities. Furthermore the literature highlights their importance in the field of Ayurvedic medicine.

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