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ASWAGANDHA - A CRITICAL REVIEW

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ABSTRACT

Objective: The objective of this paper is to compile the information on *Aswagandha* from various Ayurvedic textbooks, research papers and reports. Design: This review is in a descriptive format and consists of all relevant publications on *Aswagandha* that were identified through an organized search in major computerized medical databases and Ayurvedic texts. The therapeutic properties are compiled based on the useful part and important chemical constituents. Results: Studies indicate that *Aswagandha* possesses properties such as anti-inflammatory, anti-tumor, anti-stress, antioxidant, immunomodulatory, hepatoprotective, anti-parkinsonism, hemopoetic effects. It also exerts a positive influence on the endocrine, cardiopulmonary, and central nervous systems. Toxicity studies reveal that *Aswagandha* appears to be a safe compound. Conclusion: Preliminary studies have established *Aswagandha*, *Withania somnifera* exhibit a range of therapeutic effects with little or no associated toxicity. These results are quite inspiring and indicated the need of further extensive studies on this herb to confirm these properties and other potential therapeutic effects.

Keywords: Aswagandha, Withania somnifera, anti-inflammatory, antiparkinsonism, antioxidant, anti-stress.

INTRODUCTION

Nature acts as a major repository for new and novel therapeutics¹. According to the World Health Organization (WHO) report, about 4 billion people of the world presently use herbal medicines for their primary healthcare as an alternative system of medicine². Ayurvedic classics have a unique source of medicinal plants and immense traditional knowledge of herbal medicine for the treatment of various diseases³. Aswagandha is an important drug mentioned in the Ayurvedic literature with enormous medicinal value. The roots of the plant are described with rasayana karma, which are proved to promote health and longevity by enhancing the defense mechanism, slowing

the ageing, rejuvenating the body in debilitated conditions and by creating a sense of mental wellbeing⁴. Aswagandha is also mentioned in Samhitas such as Charaka Samhita, Susruta Samhita and Nigantus viz., Astanga Nigantu, Dhawanthari Nigantu, Sodhala Nigantu on their therapeutic applications. The objective of this paper is to review the available literature on Aswagandha (Withania somnifera) which have a proven therapeutic potential. Present study compiled the ayurvedic literatures, taxonomical features, phytochemistry, and pharmacological properties of this drug.

Aswagandha

Basonym

Aswagandha- aswagandhasyevagandho asyaadhavaaswasyevagandhotsaah: (kaamavega:) sevanenasa⁵|

Having smell like that of horse and promotes sexual potency like that of horse.

Main synonyms and Interpretation

Asakandha- aswasyamedramivakandamoolasya⁵ | - Root resembles penis of aswa; Aswaavarohaka:-aswagandhaavruksha:⁵ | - Having smell like that of horse and promotes sexual potency like that of horse; Kanchuka- kaci-pratibandhanayo:⁵ | - Retains semen; Kaamaroopini-

kaamammanonjamroopamasyatasya⁶|- Promotes complexion; Gandhapatri- gandhayuktapatramasya⁵|- Leaves having smell like that of horse; Putradaputramgarbhamdadatisevanenaiti⁶|- Increases the potency and quality of sukrathereby increasing the chance of progeny; Balada- balamdadati | poshtikebaladaasmruta⁵|- Promotes strength; Vrushaprajananasaamarthyam ⁵|- Promotes sexual potency; Hayahwaya- haya: hayagandhoasyataya aswa: | aswagandhaayam⁵|- It is known by the title of horsebecause of having the smell of horse.

Vargeekarana

Charaka Samhita-Balya, Brumhaneeya, *Maduraskanda*⁷; Susrutha Samhita-*Urdwabhagahara*⁸; Bhavaprakasa Nigantu-Gudoochyadi varga⁹; Kaiyadeva Nigantu- Oshadi varga¹⁰; varga¹¹; Raja Nigantu-Satahvadi varga¹²; Madanapala Nigantu-Abhayadi Gudoochyadi varga¹³; Dhanvanthari Nigantu-Saligrama Nigantu- Godoochyadi varga¹⁴; Sodala Nigantu- Gudoochyadi varga¹⁵; Niganthu Adarsh-Kantakaryadi varga¹⁶; Priya Nigantu- Satapushpadi varga¹⁷

Taxonomy¹⁹

Order: Tubiflorae; Family: Solanaceae; Genus: Withania; Species: somnifera

This plant is distributed throughout the dry areas of India, Balochisthan, Ceylon- Mediterranean regions, Canaries and Cape of Good Hope. It is a perennial branched erect under shrub of 0.3-1.5 m height. Stem

is erect and branches are terrete, herbaceous above, woody below.

Part Used- Root, Leaves, seeds²⁰.

Dosage- 3-6 g of the drug in powder form¹⁸.

Important phytochemical constituents

Root- The roots are reported to contain amino acids, alkaloids, volatile oil, steroids, starch, reducing sugars, hentriacontane, dulcitol, glycosides, withaniol, an acid, and a neutral compound²¹; Steroidal alkaloids-Roots contain steroidal alkaloids with variety of therapeutic efficacy such as isopelletierine, anaferine, cuscohygrine, anahygrine, tropine, pseudotropine, anaferine, isopelletierine, withananine, withananinine, pseudo-withanine, somnine, somniferine, somniferinine (Basic alkaloids), 3-tropyltigloate (Neutral alkaloids), withanine, withasomnine, visamine²¹; Steroidal lactones (withanolides)withanolides, withaferins²²; Free amino acids- aspartic acid, glycine, tyrosine, alanine, proline, tryptophan, glutamic acid, and cystine²³; Others: resin, fat, Iron, coloring matters, a reducing sugar, phytosterol, Ipuranol and a mixture of saturated and unsaturated acids.

Leaves- Contains 12 Withanolides, 5 unidentified alkaloids, many free amino acids, chlorogenic acid, glycosides, glucose, condensed tannins, and flavonoids²³. Withanolides—a group of C28 steroids characterized by a 6-membered lactone ring in the 9-carbon atom side chain. Withanolides— Withaferin $A[(4\beta,5\beta,6\beta,22R)-4,27-Dihydroxy-$

5,6:22,26diepoxyergosta-2,24-diene-1,26-dione]²³;

Fruits- The green berries contain amino acids, a proteolytic enzyme, condensed tannins, and flavonoids. They have a high fraction of free amino acids which include proline, valine, tyrosine, alanine, glycine, hydroxyproline, aspartic acid, glutamic acid, cystine and cysteine²³; **Shoots-** The tender shoots are abundant in crude protein, calcium and phosphorous, and are not fibrous. They are reported to contain scopoletin²³.

Rasa panchaka^{9, 10, 11,12,13, 16, 17}

The Ayurvedic pharmacological properties of root were explained in the different Ayurvedic texts is described in the table 1

Nigantu	Bp.N	R.N	P. N	es of root explained in Ni.R	D.N	Mp.N	Kai. Ni
Rasa	Tikta, Kashaya	Katu, tikta	Tikta	Kinchitkatu, tuvara	Kashaya, tikta	Tikta, kashaya	Tikta, kashaya
Guna				Laghu			
Veerya	Usna	Usna	Usna	Usna	Usna	Usna	Usna
Vipaka							
Karma	Vatakaph ahara, balya, rasayani, atisukrala , sopha, switra, kshayahar a	Vatahara, balya, kasa, swasa, kshayavrana hara	Vataha ra, balya, vrushy, rasayan i	Jaravyadhinasaka, dhatuvrudhikara,ba lada, kantiprada, sareerapushtikari, vrushya, vatakaphahara, swasa, kasa, vrana, swetakuhta, visha, krumi, sopha, kshata, kshaya, kantuhara	Vatakaphah ara, visha, vrana, kshayahara, Kanti, veerya, balaprada	Vatakaphah ara, balya, rasayani, atisukrala, sopha, switra, kshayahara	VatakaphaharaVru shya, rasayani, Balapushtiprada, kasa, vrana, sopha, kandu, visha, switra, krumi, swasa,kshata, kshaya

Specific action (mentioned in Sarngadhara and Charaka samhita)

Sarngadhara- Sukrala²⁴

Charaka- Balya, Brumhaneeya, Madura skanda⁷

Amayigaprayoga with details of administration

- In *Granthi, Ganda, Apaci*, the external application of paste of *aswagandha* is mentioned¹⁵.
- In *Swasa*, *Kshara* is advised to be taken with honey and ghee²⁵.
- In *Granthivisarpa*, the part is advised to be applied with warm paste of *aswagandha* ²⁵.
- In Vatavyadhi, ghruta prepared with kashaya, kalka of Aswagandha along with ksheera is advised for vatavyadhi, mamsavardhana (Cakradatta-Vatavyadhiadhikara)²⁶.
- In *Vandyatwa*, *Aswagandha kashaya* added with *milk* and *ghruta* is advised to be taken internally²⁷.
- In *Karsya*, *Aswagadhachoorna* is advised to be given with milk or *ghruta*or*taila* or *sukhambu* for one month²⁸.
- In, *Nidranasa* choorna of Aswagandha is advised to be given with sita and sarpi²⁹.
- In *Kshaya*, *Aswagandhachoorna*, *pippali*, *sita* in equal quantities is advised to be taken internally with *ghruta* and *madhu*³⁰.

Important formulations in each Kalpana

Important formulations in the form of kashaya, choorna, gulika etc of Aswagandha are described below

Kashaya-MaharasnadiKashaya,

AswagandadiKashayam³¹, **Amrutadi** Kashayam (Vatikaootrakruchra³²: Choorna-Aswagandhadichoorna, Rasnadichoorna³¹; **Gulika-**Yogarajaguggulu³¹; **Lehyam-** Jeerakadilehyam (1 Rajayakshma), adanakameswarilehyam niska. Suklakara). (Chaturjatarasayana, Madhusnuhirasayana (1 karsa, prabhatakalasevitam, deepanam, balapradam, sarvadhatuvivardhanam)³¹; Arista- Aswagandharishtam (50 pala- Aswagandha, 1/2 panam, Moorcha. apasmruti, $sosha)^{31}$ pala-(100 Balarisahtam pala, vataroga), (1 Saraswatharishtam karsha, $dhanwatharinirmitham)^{31}$; sishyanamupakarartham Lavangasava pala, prameha, $dhatukshaya)^{31}$; Tailam-Astadasasatikaprasarnitailam (3 pala), Ekadasasatikaprasarinitaila (3 pala), Trisatiprasarinitaila (100 pala, Vataroga, pumsatwa), Saptasatikaprasarinitaila (100 pala, vividhavataroga) Chandanaditaila. Triphaladitaila, Dhanwantaramtaila (Sarvavatavikara), Naravanataila (10 pala, vataroga),

Mahanarayanataila (1/2)pala, vataroga), Puvankuruntalatadiavanakenna (2 kazhanju, two times, swasa, jwara), Prabhanjanavimardhanataila (2 pala, Aseetivatarogansudarunan), Balaaswagandhalakshaditaila (Sarvajwara, unmada), Mashaditaila (1 karsa, vataroga, pakshaghata, ardita). Mahamashaditaila (Pakshaghata, hanustambha). Lakshadi (Sarvajwara, kshaya, unmada), Vilwampachotyadienna (Sarvaamayapaha).. adm. urustambha)³¹: Kushtaditaila (internal Ghrutam-Aswagandhadighrutam, Mahakushmandaghruta, Vastyaamayanthakaghruta, Sasavasadighruta, varahyadighruta, Sarvaamayanthakaghruta, Sukumaramghruta³¹; **Taila-**Narayanataila (Aswagandha- 10 pala, Vataroga), Lakshaditailam (Aswagandhakarsha). 1 Satavaritailam (1/2 palam, Vandyatwa)³¹

Substitutes and Adulterants

Classical substitute for Kakoli and Ksheerakakoli³³.

Research works

Toxicity studies

In acute toxicity studies, oral LD50 of *W.somnifera* was found to be >2000mg/kg body weight. The 14th day observation and weekly bodyweight assessment revealed no toxic effects in rats. There was no abnormal behaviour shown by the rats during these 14 days of observations³⁴. In the sub-acute toxicity study, all the animals survived 28 days. Physical and behavioural examination did not reveal any treatment related adverse effects at the dose of 500mg/kg/day, 1000mg/kg/day and 2000 mg/kg/day³⁴.

In- vitro studies

Root

Antibiotic activity

W. somnifera revealed the antibacterial activity against Bacillus subtilis, E. Coli, P. Flourescens, E. Aerogens, K.pneumoniae, R.planticola A. Tumefaciens³⁵

Anti-tumour Activity

The ethanolic extract of root showed invitro cytotoxicity against five human cancer cell lines such as PC-3, DU-145 (Prostrate), HCT-15 (colon), A-549 (lung), IMR-32 (neuroblastoma)³⁶.

Antioxidant and Free Radical Scavenging Activity

Antioxidant activity was evaluated by DPPH radical scavenging assay, ferrous reducing power, nitric oxide radical scavenging activity, Fe²⁺ chelating activity assay, superoxide anion and hydrogen peroxide radical scavenging activity. The result of the study suggested that indigenous root showed more activity than the exported root³⁷.

Antimycobacterial

When tested by MIC method, aqueous extracts of different concentrations of *W. somnifera* revealed consistent reduction in activity against M.TB. It was observed that the increase in anti-mycobacterial activity by *W. somnifera* was dose dependent and the increase in anti-mycobacterial activity was significant in 3rd day and in which extends up to the 7th day. The highest antibacterial activity of M.TB was found in *W.somnifera* (1.0 mg/mL) which is 64.47% inhibition and least activity inhibits in lower dose of 0.01 mg/mL which observed 17.88%³⁸.

In- Vivo studies

Root

Anti-inflammatory activity

The effect of *W. somnifera* root powder with a dosage of 1000 mg/ kg, orally daily for 15 days revealed a reduction in the paw swelling and bony degenerative changes in Freund's adjuvant-induced arthritis in rats by radiological examination. The reductions were better than those produced by the reference drug, hydrocortisone (15 mg/kg)⁴⁰.

Antitumour activity

Withaferin A, withanolide D & E exhibited significant antitumour activity in vivo against Ehrlich ascites carcinoma, Sarcoma 180, Sarcoma Black (SBL), and E 0771 mammary adenocarcinoma in mice with doses 10, 12, 15 mg/kg body-weight. Growth of Ehrlich ascites carcinoma was completely arrested in more than half the mice which survived for 100 days without the evidence of growth of the tumour⁴¹.

Immunomodulatory Activity

Aswagandha showed a significant modulation of immune reactivity in animal models. Root extract of *W.somnifera* was found to prevent myelo-suppression in mice treated with three immunosuppressive drugs *viz.* cyclophosphamide, azathioprin, and prednisolone.

Treatment with Aswagandha was found to significantly increase Hb concentration, RBC count, platelet count, and body weight in mice. Administration of Aswagandha extract was found to significantly reduce leucopenia induced by cyclophosphamide (CTX) treatment⁴².

Anti-stress/Adaptogenic Activity

Anti-stress effect of *W.somnifera* was studied in rats using cold water swimming stress test. Root powder was given in the dosage of 100 mg/kg orally as an aqueous suspension every day for consecutive seven days before the swimming test in water at 10°C. The swimming test also showed increase in total swimming time, indicating better stress tolerance in rats. The alcohol extract of *W. somnifera* (100 mg/ kg, two times daily orally on day 1, 4 or 7) lessened stress-induced increases in blood urea nitrogen levels, blood lactic acid, and adrenal hypertrophy, but did not affect changes in thymus weight and hyperglycemia in rats⁴³.

Anticonvulsant Activity

Administration of *Aswagandha* root extract was found to lessen jerks and clonus in 70% and 10% animals respectively with a dosage of 100mg/kg and reduction was evident from EEG wave pattern. The protective effect of Aswagandha extract in convulsions has been stated to involve GABAergic mediation⁴⁴.

Hepatoprotective Activity

Withaferin A at the dose of 10mg/kg exhibited significant hepatoprotective effect against CCl4-induced hepatotoxicity in rats⁴⁵.

Neuropharmacological Activity

The root extract of *W. Somnifera* showed prolonged hypotensive, bradycardiac and respiratory stimulant activities in dogs. The total alkaloids produced a calming and a gentle depressant effect (tranquillizer-sedative type) on the CNS in several experimental animals⁴⁶. The drug elevated the level of acetylcholine receptor capability in the cortical muscarinic indicates the cognition-enhancing and memory improving effects of extract from *W. Somnifera*⁴⁷.

Hypolipidemic effect

W. somnifera root powder decreased total lipids, cholesterol and triglycerides in hypercholesteremic animals. It also significantly increased plasma HDL-cholesterol levels, HMG-CoA reductase activity and

bile acid content of liver. A comparable trend also stated in bile acid, cholesterol and neutral sterol excretion in the hypercholesteremic animals with WS administration⁴⁸.

Antiparkinsonism effect

Rats were previously treated with the *W. Somnifera* extract orally for 3 weeks. On day 21, 6OHDA(6-Hydroxy dopamine) was infused into the right striatum. Three weeks after 6-OHDA injections, rats were evaluated for neurobehavioral activity. The animals were killed after 5 weeks lessoning for the estimation of lipidperoxidation, reduced glutathione content, activities of glutathione-S-transferase, glutathione reductase, GPX, SOD and CAT, catecholamine content, dopaminergic D2 receptor binding and tyrosine hydroxylase expression. *W. Somnifera* extract inverted all the parameters significantly in a dose-dependent manner⁴⁹.

Clinical trial

Anti-ageing Effect

Double-blind clinical trial was carried out to study the effect of *W. Somnifera* on the prevention of ageing in 101 normal healthy males in 50-59 years age group. Root powder at a dose of 0.5 g was given orally three times a day for 1 year. Results showed statistically significant increase in Hb, RBC, hair melanin, and seated stature in treated group in comparison to placebo group. Decrease in serum cholesterol was more in treated group than in placebo group.

Ashwagandha (W. somnifera) root powder supplementation on the vo2 max. and hemoglobin in hockey players

Thirty two male hockey players, with a mean age of 17.4 ± 1.7 years and BMI 20.9 ± 2.9 kg/m2 were undertaken for the study. Subjects were randomly allocated into two groups Group I (n=16):

W. somnifera group (experimental group) were given 500 mg capsules of aqueous roots of Aswagandha two times daily for eight weeks and Group II (n=16): Placebo (control) group., received starch capsules. VO2 max. with 12 minute run test and hemoglobin (Hb) of both experimental and control groups were measured before and after the administration of W. somnifera and placebo respectively. A significant progress in the VO2max and hemoglobin was observed in experi-

mental group who received 500 mg capsules of aqueous roots of *Aswagandha* twice daily for eight weeks⁵⁰.

Anti-stress activity

RCT was conducted with 130 participants, out of which 98 completed the study. Participants were randomly allocated into extract group and placebo group. Stress levels were assessed on 0, 30 and 60 day using modified Hamilton anxiety scale. Biochemical and clinical variables were noted on 0 and 60 day. The extract dose at 125mg QD showed significantly decreased Hamilton score, serum cortisol, serum CRP, pulse rate and blood pressure⁵¹.

Management of Sarcopenia

The participants of the RCT within the age group of 55 and 74 years were given 500 mg *W. somnifera* (in capsule) twice daily for 3 months. It has shown a significant role in recovering muscle functioning. The changes in blood creatine kinase levels with the use of this herb imply a possible increase in muscle metabolism or a possible decline in muscle catabolism⁵².

DISCUSSION

Aswagandha is an ayurvedic medicine with potential therapeutic efficacy. It is the important ingredient of a variety of formulations prescribed for musculoskeletal disorders and for promoting the longevity and wellness of the individuals. All the synonyms of Aswagandha emphasize its importance in the usage for rasayana karma. Ayurvedic pharmacological properties are mentioned as tikta, kashaya rasa, laghu guna, usna veerya, madhura vipaka. Aswagandha is also known to possess balya, brumhana, rasayana and vajeekarana karma¹⁸. Aswagandha is prescribed in the treatment of various diseases such as vatavyadhi²⁶, swasa²⁵, nidranasa²⁹, granthi, apachi¹⁵. It is the ingredient of hundreds of formulations including kashaya, choorna, arista, asava, taila³¹. Modern pharmacological studies reveals that aswagandha possesses properties such as anti-inflammatory, anti-tumor, anti-stress. anti-oxidant, immunomodulatory, hepatoprotective, anti-parkinsonism, hemopoetic effects.

In this review, an attempt has been made to systematically explain the drug *Aswagandha* starting from its etymology and synonyms. Unlike other reviews on

this drug, we have compiled the information on the classification (vargheekarana) of the drug in various samhitas and nigantus, opinions of their ayurvedic pharmacological properties (Rasa panchaka), therapeutic effects (Amayikaprayoga), different formulations such as kashaya, ghruta, taila, choorna containing Aswagandha as its ingredient. In addition to these, chemical constituents of different parts, and pharmacological review including the toxicity as well as clinical studies and in-vitro and in-vivo studies were also compiled.

CONCLUSION

From the review it was found that, the drug Aswagandha is found to possess immense therapeutic efficacy which can be utilized in the management of various disease conditions. This review will help the researchers to study the traditional claims and utilizing the different formulations of this drug in treating various disorders. Aswagandha has been explained as best the drugs explained for brumhana, among balavardhana and sukrajanana. The studies relating the assessment of pharmacodynamics of ayurvedic pharmacological actions of Aswagandha such as brumhana, sukrala, balya has to be conducted for explaining its mode of action in various clinical conditions.

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