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**Review Article** 

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#### ROLE OF MUCUNA PRURIENS IN THE MANAGEMENT OF PARKINSON'S DISEASE

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#### **ABSTRACT**

Parkinson's disease (PD) is the second most common progressive neurodegenerative disease after Alzheimer's disease, affecting about 7 to 10 million patients worldwide. It is caused by degeneration of dopaminergic neurons in substantia nigra. The deficiency of dopamine in the basal ganglia leads to a movement disorder which is characterized by classical Parkinsonian motor symptoms which include bradykinesia, muscular rigidity and resting tremor as well as non-motor symptoms include olfactory dysfunction, cognitive impairment, psychiatric symptoms and autonomic dysfunction. *Mucuna pruriens* is an annual and perennial legume which belongs to the family Fabaceae having different types of therapeutic activity. It is widely utilized as a potent aphrodisiac. For this purpose, Google Scholar and PubMed were explored to obtain studies giving evidence for anti-parkinsonian activity and other pharmacological activities of *Mucuna pruriens*. *Mucuna pruriens* possesses different types of pharmacological activities. The neuroprotective activity and anti-Parkinsonian activity of *Mucuna pruriens* was explored by several researchers. Levodopa is the important constituents responsible for the anti-Parkinsonian activity of *Mucuna pruriens*. Apart from anti-parkinsonian activity, *Mucuna pruriens* possesses some most

common pharmacological activities including Anti-oxidant, anti-inflammatory, aphrodisiac, hepatoprotective and improving semen quality etc. This review emphasizes the anti-parkinsonian role of *Mucuna pruriens* in the management of Parkinson's disease systematically through literature review available to date. Some other activities of *Mucuna pruriens* are also summarized in this review.

**Key word:** Parkinson's disease, *Mucuna pruriens*, dopamine, levodopa.

#### INTRODUCTION

Parkinson's disease (PD) is one of the most common progressive neurodegenerative diseases caused by degeneration of dopaminergic neurons in substantia nigra.<sup>1,2</sup> It is the second most common neurodegenerative disease after Alzheimer's disease, affecting about seven to 10 million patients worldwide.<sup>3</sup> The deficiency of dopamine in the basal ganglia leads to a movement disorder which is characterized by classical Parkinsonian motor symptoms which include bradykinesia, muscular rigidity and resting tremor as well as non-motor symptoms include olfactory dysfunction, cognitive impairment, psychiatric symptoms and autonomic dysfunction.<sup>4</sup> The current therapy available for PD primarily relies on Levodopa that offers the potential of slowing down disease progression to some extent but includes lot of side effects. Any potential drug capable of treating or halting the disease still remains to be identified.<sup>5</sup> The progression of Parkinson's disease is characterized by a worsening of motor features; however, as the disease progresses, there is an emergence of complications related to long-term symptomatic treatment. The available therapies for Parkinson's disease only treat the symptoms of the disease.<sup>4</sup>

*Mucuna pruriens* is an annual and perennial legume which belongs to the family Fabaceae having different types of therapeutic activity.<sup>6</sup> Conventionally, *Mucuna pruriens* seeds are used for maintaining male infertility in traditional medicine and It is widely used as a potent aphrodisiac.<sup>6,7</sup> Reportedly, *Mucuna pruriens* is used as a rejuvenator drug having neuroprotective property.<sup>7</sup> The neuroprotective activity of *Mucuna pruriens* was shown by several researchers. The anti-Parkinsonian activity of *Mucuna pruriens* was explored since

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the nineteenth century. Mucuna is a well-known natural source of levodopa (L-dopa).<sup>8</sup> Levodopa is the important constituents responsible for the anti-Parkinsonian activity of *Mucuna pruriens*.<sup>6</sup>

Levodopa is largely unavailable and unaffordable in many low-income countries. Also dopaminergic anti-parkinsonian medications, such as levodopa cause drug-induced dyskinesias in majority of patients with Parkinson's disease. High dose *Mucuna pruriens* induces greater motor improvement than conventional levodopa. *Mucuna pruriens* could be used for symptomatic management of Parkinson's disease.

### **METHODOLOGY**

A systematic literature search was carried out to review articles and to gather the information available in the literature regarding 'Mucuna pruriens' in the view of description of the plant, chemical constituents, part used, therapeutic action and therapeutic uses, and recent scientific evidence of anti-parkinsonian activity. All the available information on 'Mucuna pruriens' was compiled from Unani, Ayurveda textbooks and electronic databases such as Google scholar and PubMed.

### **RESULTS & DISCUSSION**

## Scientific classification of Mucuna pruriens29

Kingdom: Plantae

Division: Magnoliophyta

Class Magnoliopsida

Order: Fabales

Family: Fabaceae/Leguminosae

Genus: Mucuna

Species: pruriens



Fig.1: Leaves, Flowers, Pods and Seeds of Mucuna pruriens

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Vernacular names

English: Cowhage, Velvet Bean

Tamil: *Poonaikali* 

Sinhala: Wanduru-me

Unani Tibbi name: Konch

Sanskrit name: *Kapikacchu* 

Description of the Mucuna pruriens plant<sup>30</sup>

A semi-woody twining climber with slender branches clothed with short white deflexed

hairs:

**Leaves:** alternate, stipulate, 3-foliate with stipels, large, rachis 7.5-12.5 cm long, sparingly

deflexed hairy, leaflets 7.5-10 cm long, 5-7.5 cm broad, on short, thick, hairy stalks, terminal

one smallest and rhomboid-oval, lateral ones very unequal with the lower half greatly

dilated, all acute, mucronate, pubescent above, densely covered with shining silvery

adpressed hairs beneath, stipules linear, setaceous, hairy.

**Flowers:** irregular, bisexual, dull dark purple with a yellowish green keel, numerous, 3.7-4.3

cm long, on short pubescent pedicels, usually 2 or 3 together at intervals on a slender,

pubescent raceme, 15-30 cm long, bracts 1.2 cm long, lanceolate, hairy, deciduous; sepals 5,

fused into a campanulate calyx, densely silky, two upper segments completely connate,

lowest much the longest; petals 5, exserted, very unequal, wings twice as long as the

standard, keel rather longer than wings, curved into a stiff beak at apex; stamens 10,

diadelphous; ovary superior, surrounded at base by a small crenulate disc, unilocular with

marginal ovules, style beardless, stigma capitate.

**Fruit:** legume 6.2-7.5 cm long, 1.2 cm broad, linear, blunt, falcately curved at both ends with

a longitudinal rib along the whole length of each valve but without wings, densely covered

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with close rather weak orange-brown irritant bristles pointing backwards and readily detached, 4-6 seeded with partitions between them.

**Seed:** ovoid, 0.6 cm long, compressed, brownish, mottled with black, hilum oblong.

Parts used: Seed, leaf and root.31

**Chemical Constituents:** The plant is reported to have L-3-4 dihydroxy – phenylalanine (L-DOPA) as a major constituent mainly in seeds. Alkaloidal constituents such as Mucunadine, Mucunine, Prurienidine, Prurienine are reported from seeds.<sup>31</sup>

## Properties of Mucuna pruriens according to the traditional systems of medicine

Table 01 shows the properties of the *Mucuna pruriens* according to Unani and Ayurveda systems of medicine.<sup>32,33</sup>

Table 01: Properties of *Mucuna pruriens* according to Unani and Ayurveda systems of medicine

Unani	Ayurveda
Taste: Sweet	Rasa (taste): Sweet and Bitter
<i>Mizaj</i> (Temperament): Hot 2 <sup>0</sup> Dry 1 <sup>0</sup>	Guna (attributes): Guru (Heavy) and
Naf 'e Khas (Actions):	Snigdha (Unctous)
Qabiz (Astringent)	Virya (potency): Usna (Hot)
Mulayyin (Laxative)	Vipaka (post digestive effect): Sweet
Muqawwi-e-Bah (Aphrodisiac)	Karma:
Muqawwi-e-Asab (Nervine Tonic)	It alleviates <i>Vata</i> , <i>Pitta</i> and <i>Kapha</i> , all three
Mudir-e-Baul (Diuretic)	dosas.
Mudir-e-Tams (Emmenagogue)	It helps in nourishing all the <i>Dhatu</i> (tissue elements) in the body, especially <i>Sukra</i>
Qatil-e-Deedan (Vermifuge)	(Semen) in the males.

## Therapeutic uses of *Mucuna pruriens* according to the traditional systems of medicine

Table 02 shows the Therapeutic uses of the *Mucuna pruriens* according to Unani and Ayurveda systems.<sup>32,33</sup>

Table 02: Therapeutic uses of *Mucuna pruriens* according to Unani and Ayurveda systems of medicine

Unani	Ayurveda
1. Jiryan	1. The seeds are the most commonly used ingredient in many tonics
(Spermatorrhoea)	for impotency and for enhancing sexual vitality.
2. Sailanur Reham (Leucorrhoea)	It also works well as a restorative for conditions of debility and weakness.
3. Isterkha	2. The seeds powder of <i>Kapikacchu</i> and <i>Kokilaaksa</i> impart very potent effect, when taken with sugar and followed by milk. It
(Paralysis)	augments the seminal fluid, vitality and vigour.
4. Kirm-e-Shikam	3. The hot infusion of the seeds is an excellent panacea for premature ejaculation in men.
(Worm in the	4. The seeds are useful galactogouge in lactating mothers.
abdomen)	5. The roots of this plant are diuretic.
	6. The decoction of roots is beneficial in renal problems and dysuria.
	7. The roots also help regulating the menstrual cycle and enhance sexual vigour in women.
	8. In <i>Vata</i> diseases like Facial palsy, Cervical spondylosis, Parkinson's disease, Paralysis etc, the decoction of <i>Kapikacchu</i> is rewarding.

# Compound formulations of *Mucuna pruriens* according to the traditional systems of medicine

Table 03 shows the Compound formulations of the *Mucuna pruriens* according to Unani and Ayurveda systems<sup>32,33</sup>

Table 03: Compound formulations of *Mucuna pruriens* according to Unani and Ayurveda systems of medicine

Unani	Ayurveda
<ol> <li>Laboob-e-Kabeer</li> <li>Laboob-e-Sagheer</li> <li>Majoon-e-Konch</li> <li>Safoof-e-Mundi</li> </ol>	<ol> <li>Vanari Gutika</li> <li>Masabaladi Pacana</li> <li>Vajikarana ghrta</li> <li>Brmhana Gutika</li> </ol>

## Recent Scientific Evidence of Anti-Parkinson Activity of Mucuna Pruriens

Cilia R, et al. (2017) carried out a study to investigate whether *Mucuna pruriens* (MP may be used as alternative source of levodopa for indigent individuals with Parkinson disease (PD) who cannot afford long-term therapy with marketed levodopa preparations. They investigated efficacy and safety of single-dose intake of MP powder from roasted seeds obtained without any pharmacologic processing. Single-dose MP intake met all noninferiority efficacy and safety outcome measures in comparison to dispersible levodopa/benserazide. Clinical effects of high-dose MP were similar to levodopa alone at the same dose, with a more favorable tolerability profile.<sup>11</sup>

Adi YK, et al. (2018) evaluated the neuroprotective effect of n-Propanol extracts from fresh, boiled, and fermented seeds of *Mucuna pruriens* in a Parkinson's disease (PD) rat model, based on the total number of dopaminergic (DA) neurons in the substantia nigra pars compacta (SNpc). n-Propanol extract of boiled and fermented seeds could produce a higher neuroprotective effect against DA neuron than fresh seeds in a PD rat model.<sup>12</sup>

Rai SN, et al. (2017) investigated the effects of aqueous extract of Mp (100 mg/kgbwt) on neuroinflammation, orally administered to mice intoxicated with 1-methyl-4-phenyl-1,2,3,6-tetrahydropyridine (MPTP) as well as the molecular mechanism involved in the progression of PD. The findings observed thereby indicate that Mp extract have suggestively ameliorated MPTP induced neuroinflammation, restored the biochemical and behavioral abnormalities in PD mouse and thus provided a scientific basis for its traditional claim.<sup>13</sup>

Yadav SK, et al. (2014) carried out a study to evaluate the neuroprotective effect of an ethanolic extract of Mp seed in the 1-methyl-4-phenyl-1,2,3,6-tetrahydropyridine (MPTP) model of PD and compared to estrogen, a well reported neuroprotective agent used for treating PD.. Notably, the effect of Mp was greater than that elicited by estrogen. Mp down regulates Nitric Oxide production, neuroinflammation and microglial activation and all of these actions contribute to Mp's neuroprotective activity. These results suggest that Mp can be an effective treatment for neurodegenerative diseases, especially PD by decreasing oxidative stress and possibly by implementing neuronal and glial cell crosstalk.<sup>14</sup>

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Lieu CA, et al. (2012) evaluated the effects of *Mucuna pruriens* to determine if its underlying mechanistic actions are exclusively due to LD. We first compared MPEP with and without carbidopa (CD), and LD+CD in hemiparkinsonian (HP) monkeys. Each treatment ameliorated parkinsonism. The distinctive neurophysiological findings in the basal ganglia and the ability to ameliorate parkinsonism without causing dyskinesias strongly suggest that *Mucuna pruriens* acts through a novel mechanism that is different from that of LD.<sup>15</sup>

Lieu CA, et al. (2010) conducted a study to compare the behavioral effects of chronic parenteral administration of a water extract of M. pruriens seed powder (MPE) alone without any additives, MPE combined with the peripheral dopa-decarboxylase inhibitor (DDCI) benserazide (MPE+BZ), LD+BZ and LD alone without BZ in the hemiparkinsonian rat model of PD. This study demonstrates that a simple water extract of *Mucuna pruriens* endocarp powder with no additives has a superior effect to the combination of MPE+BZ on parkinsonism and that MPE alone is superior to LD alone or LD+BZ combinational therapy in terms of efficacy of ameliorating parkinsonism with dramatically reduced risk for DID.<sup>16</sup>

Katzenschlager R, et al. (2004) assessed the clinical effects and levodopa (L-dopa) pharmacokinetics following two different doses of mucuna preparation and compared them with standard L-dopa/carbidopa (LD/CD). The rapid onset of action and longer on time without concomitant increase in dyskinesias on mucuna seed powder formulation suggest that this natural source of L-dopa might possess advantages over conventional L-dopa preparations in the long term management of PD.<sup>17</sup>

Improved L-DOPA concentrations in a soluble phenolic and antioxidant-rich M. pruriens background on day 1 sprouts have potential for Parkinson's disease management.<sup>18</sup>

Manyam BV, et al. (2004) evaluated the neurorestorative effect of *Mucuna pruriens* cotyledon powder on the nigrostriatal tract of 6-OHDA lesioned rats. Unlike synthetic levodopa treatment, *Mucuna pruriens* cotyledon powder treatment significantly restored the endogenous levodopa, dopamine, norepinephrine and serotonin content in the substantia nigra. Nicotine adenine dinucleotide (NADH) and coenzyme Q-10, that are shown to have a therapeutic benefit in Parkinson's disease, were present in the *Mucuna pruriens* cotyledon

powder. This additional finding of a neurorestorative benefit by *Mucuna pruriens* cotyledon powder on the degenerating dopaminergic neurons in the substantia nigra may be due to increased complex-I activity and the presence of NADH and coenzyme Q-10.<sup>19</sup>

## Other Pharmacological Activities of Mucuna pruriens

Apart from Anti-Parkinson's activity, *Mucuna pruriens* also shows following pharmacological activity as mentioned in Table 4.

Table 4: Other Pharmacological Activities of Mucuna pruriens

Pharmacological activities	References
Anti-inflammatory <sup>20,21</sup>	Martínez Leo EE, et al. (2018), Rachsee A, et al. (2020)
Anti-oxidant <sup>22,23,24</sup>	Jimoh MA, et al. (2020) Martínez-Leo EE, et al. (2019) Obogwu MB, et al. (2014)
Aphrodisiac <sup>24</sup>	Ahmad MK, et al. (2008)
Hepatoprotective <sup>25</sup>	Obogwu MB, et al. (2014)
Improving semen quality <sup>26,27,28</sup>	Gupta A, et al. (2011), Shukla KK, (2010), Shukla KK, et al. (2009)

#### **CONCLUSION**

Parkinson's disease is a progressive, disabling, neurodegenerative disease that requires long pharmaceutical treatment. Levodopa remains the gold standard treatment for Parkinson's disease globally, although it is largely unavailable and unaffordable for the majority of patients in low-income countries. Also dopaminergic anti-parkinsonian medications, such as levodopa cause drug-induced dyskinesias in majority of patients with Parkinson's disease. *Mucuna pruriens* has been shown to induce a great improvement of motor symptoms with

few adverse events in recent studies. We suggest the potential for *Mucuna pruriens* to replace or supplement levodopa-based medicines in countries where levodopa is unaffordable and inaccessible. However, caution is important until data on long-term safety of *Mucuna pruriens* are available. If long-term *Mucuna pruriens* proves to be safe and effective in clinical trials, it may be a sustainable alternative therapy for Parkinson's disease.

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