

Gynura nepalensis DC: A Potential Wonder Medicinal Plant

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This is a review of published studies on *Gynura nepalensis* DC, a medicinal plant locally known as Philippine Ashitaba. Although it is utilized as an ethno-medicine to cure a wide array of human sickness such as indigestion, diabetes, cuts/wounds, cough, asthma, kidney stones, urinary tract bleeding, hepatitis, gall bladder stones, hemorrhoids, constipation, diarrhea, vomiting, blood poisoning, septicemia, fertility problems, skin allergy, rheumatism, high cholesterol levels, and high-low blood pressure, only its hypolipidaemic, hepatoprotective, antimutagenic, anti-oxidant and radioprotective potentials using either ethanolic or methanolic extracts have been directly investigated based on reported studies. There are no reported studies on the antibacterial, antifungal, and cytotoxic potentials of *G. nepalensis* DC extracts using other solvents like water, DCM, DMSO, chloroform, ether, and others. Thus, many of the pharmacological claims that are based on anecdotal report need scientific validation.

Preliminary phytochemical screening of the leaf extract of the plant indicated the presence of alkaloids, tannins, flavonoids, steroid (with 2-deoxysugars), flavonoid (leucoanthocyanin), and saponin. *G. nepalensis* DC is often taken as similar with other *Gynura* spp. No molecular studies to establish the DNA barcode or fingerprint of the plant have been reported. Therefore, since many cures have been attributed to *G. nepalensis* DC, making it a potential wonder medicinal plant, the scientific community needs to conduct more researches - bioassays, molecular studies, and chemical profiling - on this plant.

Keywords: *Gynura nepalensis*, ethno-medicine, medicinal plants, phytochemistry

1 Introduction

Grierson and Long (2001) (Afroz, Uddin, & Hassan, 2014) ? Grierson and Long (2001)

Medicinal plants are considered alternative medicines and their use in the rural communities is encouraged. As the price of synthetic medicines continues to rise, people resort to medicinal plants as cheaper alternatives hoping to get cured of their illnesses. Such decision, however, is usually based on local wisdom or folk knowledge. *Gynura nepalensis* DC is a medicinal plant with wide curative application based on ethno-medical knowledge. As such, many researchers, having recognized this wide potential, have conducted studies on *G. nepalensis* DC. This rapid review seeks to bring together the current state of research on *G. nepalensis* DC and highlight ethno-medicinal claims on *G. nepalensis* DC that have

scientific bases as reported in scientific journals. The review tried to consider available reports accessible or searchable in the internet. We searched reputable data bases such as Elsevier for published articles on *G. nepalensis* DC but there is a dearth of published studies. Thus, the review includes studies in refereed journals, two unpublished theses (i.e., Salas, 2016) and (San Andres, 2014) that the authors have personal knowledge of, a report presented in a scientific convention (i.e., Bustillos, Quijano, Tandigan, & Cabling, 2013), and two websites Plantlist.org and Tropicos.org for the taxonomic classification. Studies published in personal or public websites were excluded for being not peer-reviewed.

2 Distribution and Biology of *Gynura nepalensis* DC

Distribution of the Gynura species.

About 44 species of *Gynura* are distributed in Africa, South and East Asia and Australasia (Vanijajiva, 2009; ?) Aside from *G. nepalensis* DC, a study by Vanijajiva (2009) revealed 9 species in Thailand: *G. bicolor* Roxb.ex Willd. DC, *G. calciphila* var. *calciphila* Kerr, *G. calciphila* var. *dissecta* F.G. Davies, *G. cusimbua* D. Don S. Moore, *G. hmopaengensis* H. Koyama, *G. integrifolia* Gagnep, *G. procumbens* Lour. Merr., *G. pseudochina* L. DC, and *Gynura* sp. which is claimed to be endemic in Thailand. Furthermore, ?, in their revision of an earlier study on *Gynura* in Thailand, reported 3 additional new species, namely, *G. daviesiae* Vanijajiva & Kadereit, *G. villosa* Vanijajiva & Kadereit, and *G. siamensis* Vanijajiva & Kadereit and 3 new combinations, namely, *G. dissecta* F.G. Davies Vanijajiva & Kadereit, *G. annua* F.G. Davies Vanijajiva & Kadereit, and *G. aurantiaca* (Bl.) DC. subsp. *parviflora* F.G. Davies Vanijajiva & Kadereit.

Taxonomy of Gynura nepalensis DC.

According to Tropicos.org (2014), the plant of interest belongs to Class Equisetopsida, Subclass Magnoliidae, Superorder Asteranae, Order Asterales, Family Asteraceae and Genus *Gynura*. Its synonyms are *Gynura dielsii* H. Lév., *Gynura nudibasis* (H. Lév. & Vaniot) Lauener & D.K. Ferguson and *Senecio nudibasis* H. Lév. & Vaniot. *G. nepalensis* DC has several common names, to wit: *ni bo er ju san qi* in China (Tropicos.org, 2014), *Tong kribi* in Bhutan (Grierson & Long, 2001), and Philippine Ashitaba in the Philippines (Gracilla & Bagunu, 2014).

Morphological Description of Gynura nepalensis DC.

Based on various sources, Vanijajiva (2009) provides the following morphological description of *G. nepalensis* DC, which is quoted in full here: "Plants 2–5 m high or more, stems erect, densely white- or tawny-tomentose, roots fibrous. Blades elliptic, narrowly elliptic, rhomboid or lyrate, 2–20 x 1–6 cm, densely tomentose, base cuneate, apex acute, margins entire to denticulate. Petioles 0.5–5 cm, exauriculate, tomentose. Capitula 3–7 per corymb, peduncles stout, 2–10 cm long; calycular bracts 3–8, 3–7 mm long, densely tomentose; involucre 8–12 mm long, 7–12 mm in diam.; phyllaries 13–14, 8–10 mm long, 1–2 mm broad, densely tomentose or sometimes glandular. Florets 25–40, orange to yellow, 9–15 mm long, exserted part 2.5–4 mm long. Anthers 2.5 mm long, anther collars elongate. Style arms 3.5 mm long. Cypselas

4–6 mm long, brown, pilose; carpopodium cylindrical, yellowish; pappus 10–12 mm long, white or dirty-white.

Phenology, ecology and distribution of Gynura nepalensis DC

In terms of phenology, "the plant is flowering and fruiting all year round" according to Vanijajiva (2009) but Afroz et al. (2014) reported its phenology from March to August. Ecologically, it grows "at the edge and in open places of mountain forests, 700-4500 m alt" (Vanijajiva, 2009). It is native in Nepal and distributed in India, Bhutan, China, Myanmar, Thailand, and Bangladesh (Vanijajiva, 2009; Afroz, Uddin & Hassan, 2014). In the Philippines the plant is also distributed but Ursulom & Rialubin (2013) reported that the Ashitaba grown in the country is *G. nepalensis* DC, which they wrote is also called *G. procumbens* and *G. acutifolia*. These three names, however, actually represent three different *Gynura* species.

3 Ethnomedicinal Use

Gynura nepalensis DC is used to cure many human diseases.

Indigestion.

The leaf extract of *G. nepalensis* DC is one of the 40 ethnomedicinal plants used to treat indigestion by the members of the Apatani tribe, in the Ziro Valley of Arunachal Pradesh, one of the 28 states in Northeast India and also considered the 12th mega bio-diverse regions of the world (Kala, 2005).

Diabetes.

The use of *G. nepalensis* DC leaves to treat diabetes has been reported in Bangladesh (Afroz, Uddin, & Hassan, 2014) and in the Philippines (Ursulom & Rialubin, 2013).

Cuts and Wounds.

G. nepalensis DC's juice is used to heal cuts and wounds in Nepal (Manandhar, 2002, cited in Afroz, Uddin, & Hassan, 2014) and external wounds in the Philippines (Ursulom & Rialubin, 2013).

Array of Diseases.

Ursulom & Rialubin (2013) reported that *G. nepalensis* DC and *Angelica keiskei* Koidzumi, which they seem to suggest to be the same, can treat an array of diseases including cough, asthma, kidney stones, urinary tract bleeding, hepatitis, gall bladder stones, hemorrhoids, constipation, diarrhea, vomiting, blood poisoning, septicemia, fertility problems, skin allergy, rheumatism, high cholesterol levels, high-low blood pressure. In addition, it can also suppress further cancer cell growth, help bring about balanced blood cells and aid in bone injury recovery. The plant part used for the treatment of the enumerated diseases is the leaf, which can be boiled, dried, pounded and mixed with oil and applied as poultice. Alternatively, 1 or 2 leaves can be eaten raw for prevention and 2 or more leaves for treatment, after rinsing the same with water and salt. The citations or references for the above long list of claims, however, cannot be found in the literature cited section of the paper. In their interview of 50

users, Ursulum & Rialubin (2013) reported that the most dominant beneficial effect [of using *G. nepalensis* DC] is giving a lighter and healthier feeling, while the least are aiding indialysis, treating pharyngitis, goiter, and pain in nipples; thus, it is a health enhancer, medicine and good for health maintenance.

Ethnopharmacological studies on *Gynura nepalensis* DC.

Several studies on the pharmacological potentials of *G. nepalensis* DC have already been published. However, some studies tend to be indirect insofar as findings on other *Gynura* species are applied (or generalized) to *G. nepalensis* DC.

4 Studies on *Gynura nepalensis* DC

Direct Studies.

- **Hypolipidaemic and Hepatoprotective Activity.** Nigam, Paarekh, Singh, Goyanar, Upmanyu, & Banweer (2012) have established the hypolipidaemic and hepatoprotective activity of the aqueous flower extract of *G. nepalensis* DC on Streptozotocin-induced diabetic mice, at doses of 100 mg/kg and 200 mg/kg b.w., respectively. After 21 days of extract administration, they observed significant reduction in serum cholesterol and triglyceride levels, significant decrease in HDL, and significant increases in LDL and VLDL in the experimental animal group. Significant increases in hepatic enzyme levels, namely, Serum Glutamate Pyruvate Transaminase (SGPT), Serum Glutamate Oxaloacetate Transaminase (SGOT), and Alkaline Phosphatase (AP), were significantly reduced after 21 days of treatment with the extract.
- **Radioprotective Activity.** Gracilla and Bagunu (2014) reported the radioprotective potential of the methanolic leaf extract of *G. nepalensis* DC. At 100
- **Antimutagenic Activity.** Bustillos, Quijano, Tandigan and Cabling (2013) have determined the antimutagenic activity of *G. nepalensis* DC using micronucleus assay.
- **Phytochemicals in *Gynura nepalensis* DC.** Phytochemical screening on the Ethanolic leaf extract of *G. nepalensis* DC [conducted in Pampanga State Agricultural University] indicated the presence of alkaloids, tannins and flavonoids (Sales, 2014). Another phytochemical screening of the Ethanolic leaf extract of *G. nepalensis* DC revealed the presence of primary alkaloids, steroid (with 2-deoxysugars), flavonoid (leucoanthocyanin), saponin and tannin (San Andres, 2014). According to Tiwari, Kumar, Kaur, Kaur, & Kaur (2011) alkaloids and tannins have antimicrobial, anthelmintic and antidiarrheal activity; flavonoids have antimicrobial and antidiarrheal activity; and, saponins have antidiarrheal activity.

Indirect Studies.

- **Based on *Gynura procumbens* and *Gynura divaricata*.** Gracilla and Bagunu (2014) stated that the ethno-pharmacological and phytochemical properties of Philippine Ashitaba (*Gynura nepalensis* D.C.) have been established and that the extract contains alkaloids, catechic tannins, saponins and flavonoids.

These statements, however, are based on two studies which were conducted on *G. procumbens* (Hassan, Yam, Ahmad, & Yusof, 2010) and *G. divaricata* (Wan, Yu, Zhou, Liu, Tian, & Cao, 2011). These plants, however, belong to different species of *Gynura*.

- Based on *Angelica keiskei* Koidzumi. Ursulom and Rialubin (2013) also reported that some studies have shown that *G. nepalensis* DC and *Angelica keiskei* Koidzumi, which they suggest to be the same, are "anti-oxidant, anti-cancer, anti-aging, anti-inflammatory, antihypertensive, and antidiabetic". The authors [Ursulom and Rialubin (2013)] cited three studies on *Angelica keiskei* Koidzumi, which is also called Ashitaba in Japanese. These three Japanese studies reported the potential of *A. keiskei* Koidzumi extract "to improve insulin resistance and hypertriglyceridemia in rats fed with high-fructose drink" (Ohnogi, Kudo, Tahara, Sugiyama, Enoki, Hayami, Sagawa, et al., 2012), "to produce[s] elevation of serum HDL levels and a reduction of liver triglyceride levels in stroke-prone spontaneously hypertensive rats (SHRSP)" (Ogawa, Nakashima, & Baba, 2003), and "to promote anti-tumor activity" (Okuyama, Takata, Takayasu, Hasegawa, Tokuda, Nishino, Nishino, & Iwashima, 1991). These studies, however, were conducted specifically on *A. keiskei* Koidzumi and not on *G. nepalensis* DC. Thus, there seems to be an attempt to generalize the findings on *A. keiskei* Koidzumi to *G. nepalensis* DC. *A. keiskei* Koidzumi belongs to the carrot Family Apiaceae (carrot) (<http://www.theplantlist.org>) while *G. nepalensis* DC belongs to Family Asteraceae / Compositae (<http://www.tropicos.org>; <http://www.theplantlist.org>). These two plants are separate species.
- Extraction Solvents used. Several solvents are used to extract the bioactive components of medicinal plants. The ability of these solvents depends on their polarity. Tiwari, Kumar, Kaur, Kaur, & Kaur (2011) reported the phytochemicals that specific solvents can extract, to wit: water "anthocyanin, starches, tannins, saponins, terpenoids, polypeptides, and lectins; ethanol "tannins, polyphenols, polyacetelenes, flavonols, terpenoids, sterols, and alkaloids; methanol "anthocyanins, terpenoids, saponins, tannins, xanthoxylines, totarol, quassinoids, lactones, flavones, phenones, and polyphenols; chloroform "terpenoids and flavonoids; ether "alkaloids, terpenoids, coumarins and fatty acids; and, acetone "phenol and flavonols. So far, published studies on *G. nepalensis* DC have used either ethanol or methanol as extraction solvents. Both ethanol and methanol are polar protic solvents. There is no report on the use of other solvents such as water, DCM, DMSO, acetone and others.

5 Conclusion and Recommendation

The review presented here has yielded several observations for the following conclusions. As a medicinal plant, *Gynura nepalensis* DC appears to have great potentials as it is claimed to be able to cure a lot of diseases. However, the scientific bases for many of these "cures" are not yet well established. *G. nepalensis* DC is confused with other species particularly *G. procumbens*. But insofar as *G. nepalensis* DC is a different species, applying the findings of other species on *G. nepalensis* DC should be approached with

caution as this needs further validation. So far, there is no published study on the antibacterial, antifungal, and cytotoxic potentials of the plant. Furthermore, the usual extraction solvent used in published studies reviewed here is either methanol or ethanol. Thus, other solvents need to be used to validate the presence of other phytochemicals in *G. nepalensis* DC. No molecular studies on any of the *Gynura* species have been published or can be found in the literature available or searchable in the internet. It is safe to conclude that no DNA barcoding or fingerprinting studies have been undertaken.

The scientific community needs to address the gaps identified in this review. More bioassays need to be conducted on *G. nepalensis* DC involving extraction solvents other than ethanol and methanol to scientifically validate the pharmacological claims on this plant. Alternatively, bioassays already conducted on *G. nepalensis* DC should be reported to the scientific community through publication in national or international journals. Molecular studies for DNA barcoding or fingerprinting of the various *Gynura* species and the chemical profiling of *G. nepalensis* DC are hereby recommended.

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