

REVIEW ON ISOLATION, IDENTIFICATION AND APPLICATIONS OF *Simarouba glauca* PLANT

D. Siva Selvi¹, N. Vijayakumar^{1,*}, R. Jayaprakash² and M. Reddi Nagesh¹

¹Department of Biochemistry and Biotechnology, Annamalai University,
Annamalainagar-608002, Tamil Nadu, India

²Department of Chemistry, School of Arts and Science, AVIT Campus, VMRF,
Chennai-603104, Tamil Nadu, India.

*E-mail: nvkbiochem@yahoo.co.in

ABSTRACT

Natural Plant species and traditional methods are valuable in the medical field to control various diseases. Recently natural plant-based medicines have received major attention in the medical research field and act as a vital part of healthcare. Also, these natural bioactive compounds are the basic molecules for the new drug synthesis. But the researchers are facing problems such as species identification, isolation of active compounds and their medicinal action in different therapeutic fields in research. This work investigated the completed research and its outcomes of the identification of new biologically active plants and their active parts. Based on the social problem, this review also investigated the *Simarouba Glauca* plant extraction, isolation of active compounds from the plant and phytochemical analysis of the reported research outcomes. Because, global health challenging illnesses like cancer, diabetes and AIDS are increasing due to the lacking of synthetic medicines. So the new researchers are concentrating on natural plants, algae and sea living organisms for drug development. Hence this work concentrated to identify the usefulness of *Simarouba Glauca* plant and different work carried on the plant products. These kinds of plant extracts are tested for various biotechnology methods to evaluate its drug suitability.

Keywords: *Simarouba Glauca*, Medicinal Plants, Identification, Isolation, Applications.

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INTRODUCTION

Nowadays plants are acting as an important role in the medical field. Traditional plants are used as medicines by the peoples still today. The main advantages of the natural plants are not showing any side effects between the peoples. These treatments are classified into different types based on the plant parts using methods such as Siddha, Rasashastra, Unani, Sa-Rigpa, Ayur vedha, yoga, and Chinese as per the Indigenous Medicines of India.¹ World Health Organization (WHO) exposed that 80% of the world people are depending on the plant-based medicines because of low cost and availability to fulfill their primary healthcare needs.² But due to the side effects of synthetic drugs, herbal drugs and their requirements are increasing day-by day.³ Developing countries people are using medicinal plants due to poverty and its availability in nearby areas.⁴ The habit of using medicinal plants has been increasing every year and most of the Americans are also started to use herbal products.⁵ So, natural drug usage is increasing to overcome the different ill effects.⁶ Hence this research reviewed the reports about the *Simarouba Glauca* (SBGA) tree parts and its usefulness in different therapeutic fields instead of selecting small herbs and small plants.⁷ Due to the availability of the trees than herbs or plants, this work selected the Shorgum Maram (Lakshmi Taru tree) which is growing in water-holding soil and sub-soil moisture. The commonly known Paradise or Bitter wood tree (Fig.-1) is in the order of Sapindales which is in the family of Simaroubaceae.

Alternative Names of *Simarouba Glauca*

The *Simarouba Glauca* can be called as bitter ash, Paradise tree, acajou blanc, daguillo gabilan, olivio, palo amargo, bwa fwenn, doliv fwenn, bois amer, quinquina, Mountain Damson, *Simarouba*, Dysentery bark and Acituno based on the available countries.

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Fig.-1. *Simarouba Glauca* (SBGA) Plant Parts

Global Distribution of Paradise Tree

The paradise tree is globally available in the following countries such as Southern Florida, West Indies, Brazil, Bahamas, Costa Rica, Cuba, El-Salvador, Guatemala, Haiti, Honduras, Jamaica, Mexico, Puerto Rico, United States of America, Sri Lanka, India, Philippines and Myanmar.⁸

Various Uses of Plant Parts

The seeds of *Simarouba glauca* contain 60-75% oil. It can be easily processed which is suitable for both edible and non-edible applications. Each complete grown tree yields 15 to 30 kg nuts, which is equivalent to 2.5-5 kg oil. The oil is isolated from the *Simarouba Glauca* seeds.⁹

RESULTS AND DISCUSSION

Various Reports on SBGA

Recently SBGA extracts and its parts are used for various applications. Hence at first, this work referred to the recent reports. Based on the reports, this review extended up to 2000. Jose *et al.* has testified the hexane, ethyl acetate, 70% ethanol-water, chloroform, leaf extracts of SBGA on HCT-116 and HCT-15 cells by well-known Sulforhodamine-B assay.¹⁰ The outcomes revealed that the chloroform extract exhibited maximum anticancer activity and they have isolated a potent molecule from the extract and shown in Fig-2(a). The reported anticancer activity of chloroform extract has displayed in Fig-2(b). Sridevi *et al.* have exposed the in vitro antioxidant and MTT assay on T-24 (Bladder cancer cell) cancer cell using methanol, chloroform, ethanol, hydro alcohol ethyl acetate and water extracts of the leaf.¹¹ Using Folin-Ciocalteu reagent, the Phenolic content of the extracts was measured and reported. The same extracts were carried for 2,2-Diphenyl-1-picrylhydrazyl (DPPH) radical scavenging assay, Ferric ion reducing antioxidant power (FRAP) assay, Phosphomolybdenum (PM) assay for their antioxidant property. Their result and conclusion stated that alcoholic extracts showed more antioxidant and cancer activity. K.S. Lakshmi *et al.* have reported the different solvent extraction of the leaves using chloroform, methanol and ethyl acetate solvents.¹² Further, these extracts were carried for the phytochemical analysis followed by an antioxidant, antimicrobial, thrombolytic and hemolytic activities. Their Phytochemical analysis confirmed the main chemical constituents such as alkaloids, a phenolic compound, terpenoids, flavonoids, glycosides, and cardenolides. The outcomes exposed that the increase in concentration

exposed to increasing activity. The extract exhibited scavenging activity $IC_{50} = 6.72 \mu\text{g/mL}$ for chloroform extract. Rajurkar B.M *et al.* have described the leaves of *Clerodendrum infortunatum* Linn, *Simarouba glauca* and *Psoraleacory lifolia* ethanolic extracts and their antimicrobial activity.¹³ The research outcomes showed the good inhibitory activity of the extracts when compared to tetracycline standard drugs. Patil *et al.* have reviewed the various uses of *Simarouba Glauca*.¹⁴ This review also exposed the bark of SBGA used for dysentery control. Also, this review exposed the medicinal properties of bark, leaf extracts pharmacological activities such as antiparasitic, antipyretic, anticancerous and antidysentric.

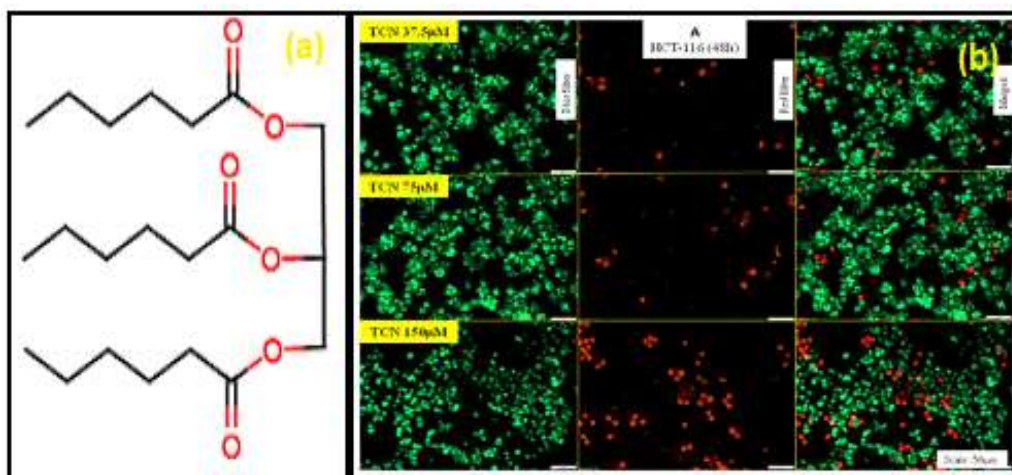


Fig.-2: (a)Tricaproin from Chloroform Extract, (b)Anticancer Activity of the SBGA Chloroform Extract¹⁰

In addition to this they have summarized the pharmacological, Phytochemical and Ethnobotanical characters of the SBGA. Some of the chemicals isolated from the seed also reported in the review, which is displayed in Fig.-3. Padovan *et al.* have reported the effect of the root of the *Tabebuia rosea* Bertol and evergreen *Simarouba Glauca* when they planted in a selected soil.¹⁵ In addition to this, the report also correlated the root effect with the rainfall. The research conclusion exposed that the *Simarouba Glauca* roots effect on coffee roots and enhance the soil nutrients.

The vegetable oils are the source for biodiesel research nowadays as per the reports.¹⁶ They have examined the *Jatropha*, *Pongamia* and *Simarouba* oils for biodiesel research. The outcomes exhibited the biodiesel property of the seed oil. The seed and oil quantity relation graph showed in Fig.-4. The conclusion confirmed that the *Simarouba Glauca* is the best source for the biodiesel preparation. The same kind of work already carried and reported by Chavan *et al.*¹⁷ Their outcomes also supported the biodiesel property of SBGA.

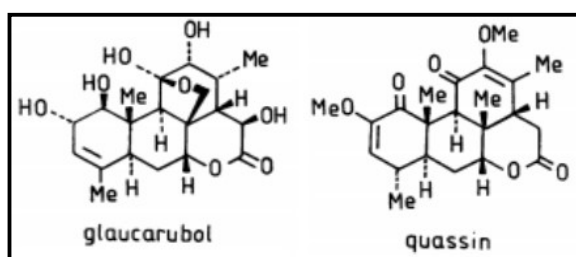


Fig.-3: Isolated Compounds from SBGA¹⁵

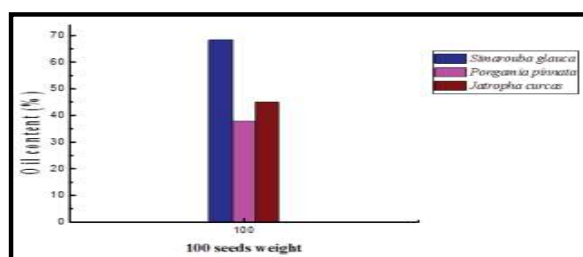
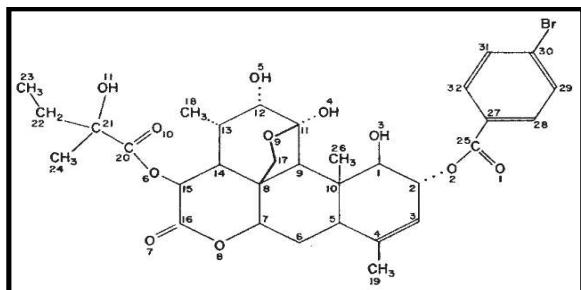
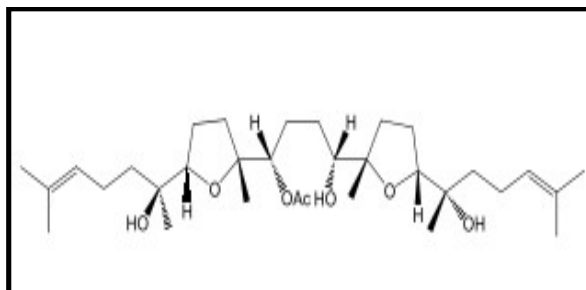


Fig.-4: Graphical Representation of weight of Seeds Vs. Oil Content¹⁶

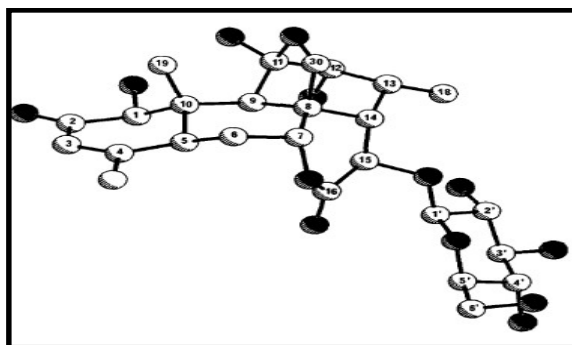
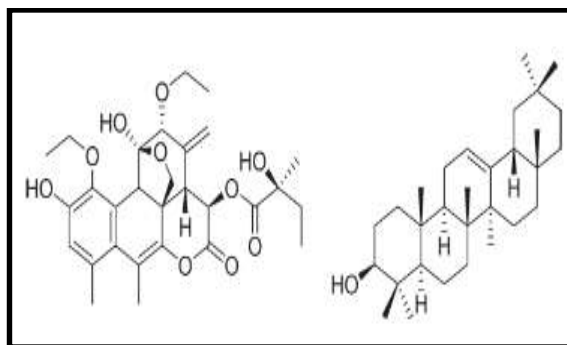
Phytochemical Analysis

Sharanya *et al.* have reported the SBGA various parts of the plants for diseases like amoebiasis, malaria, ulcer, antibacterial, cancer, antifungal, snake bite, etc. this pharmacological review revealed the *Simarouba glauca* medicinal potential.¹⁸ Assendelft *et al.* have isolated the crystalline glycoside for the

treatment of amoebiasis.¹⁹ Similarly Kartha *et al.* has reported the structure of characterized crystal structure.²⁰ Both works have completed the glycoside structure which is displayed in **Fig.-5**. SBGA plants exhibited both antimalarial and anticancer activity.²¹ Twelve compounds were isolated from the chloroform extract of southern Florida SBGA twig.²² They were tested against the epidermoid tumor cell line. Out of twelve, 14-Deacetylerylene (Fig.-6) was active against the Lu1 cancer cell line and inactive for in-vivo hollow fiber assay.

Fig.-5: Glycoside Structure from SBGA²⁰Fig.-6: Structure of 14- Deacetylerylene²²

Two toxic quassinoid glucosides were isolated from aqueous extract Simarouba glauca seeds for new biological compounds preparation. Their structures were deduced by spectral and single-crystal (Fig.-7) X-ray analysis.²³ SBGA plant can be effectively identified by Random Amplified Polymorphic DNA (RAPD) to eliminate the non-profitable male seedlings.²⁴ SBGA various parts (bark, flowers and leaves) were extracted using ethanol, chloroform, methanol, and water solvents for their antioxidant property. Different antioxidant assays were carried and the results revealed that the aqueous extract of the bark is an effective antioxidant when compared to other parts.²⁵ Different varieties of 32 genera and 170 species, including Simaroubaceae and their chemical constituents were reviewed and the research gaps regarding patents were initiated.²⁶ Some of the triterpenes have presented in Fig.-8.

Fig.-7: Single Crystal Structure of Glucosides²³Fig.-8: Some of the Quassinoid reported Triterpenes²⁶

Antimicrobial and Anticancer Activity

In addition to these the SBGA parts and their biological activities were reviewed between 1953 and 2014 merely 105 journals.²⁷ The isolated greenish-yellow, odorless fat exhibited properties such as melting point - 26.4 °C, Iodine value - 52.6, saponification value - 190.5 and oil content of 60-100g. Latter on the review stated that the aqueous extract showed the equal contents of cinnamon bark. The SBGA crude extract was tested against cancer cell lines KG-1, K-562 and MOLT-3 using different concentrations leaf extract by MTT assay.²⁸ The polar methanolic solvent extract SBGA leaves exposed noticeable anticancer activity against MOLT-3 (IC₅₀-69.69 µg/ml) and K-562 (IC₅₀-74.21 µg/ml) when compared to KG -1 (IC₅₀-131.1 µg/ml) cell lines. The SBGA plant parts can be used for leukemic cancers. Three active quassinoids of Simarouba Glauca were inhibited against chloroquine-resistant Plasmodium falciparum strain by both in-vitro and in-vivo studies.²⁹ Plants of SBGA are used as drugs mainly to cure cancer which contains the anthraquinones, phenylpropanoids, Polyphenols, flavonoids, coumarins, limonoids,

lignans, quinines, vitamins and fatty acids which were identified by phytochemical analysis.³⁰ SBGA pharmacological study against entophytic fungus *Penicillium pimateouiense* was reported and the activity (Fig.-9) is due to the presence of the above reported different organic compounds (Fig.-10) by phytochemical analysis.³¹

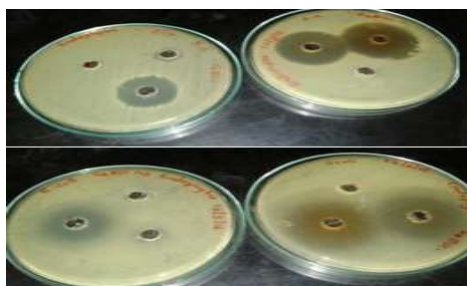


Fig.-9: Antimicrobial Activity of SBGA against the Fungal Extract Bacteria³¹

Phytochemical	Test	Result	Inference
Flavanoids	Shinoda's test	Reddish brown colour	Positive
Phenols	FeCl ₃ test	No dark green/blue colour	Negative
Alkaloids	Mayer's test	Pale creamy precipitate	Positive
Cardiac glycosides	Keller -Kiliani test	No greenish blue colour	Negative
Triterpenoids	Salkowski's test	Reddish brown colour	Positive
Carbohydrates	Molish's test	Reddish violet ring	Positive
Saponins	Frothing test	No stable froth	Negative
Tannin	FeCl ₃ test	No blue green or blue black colour	Negative

Fig.-10: Phytochemical Analysis of SBGA³¹

An anticancer ability of quassinoids compounds of SBGA was carried by docking against PI3K (Phosphoinositide 3- kinases. Fig.-11) and adme analysis. The resulting outcomes showed the binding ability and supported anticancer activity.³² Nanoparticles that can be prepared by using plant extracts were investigated using silver nitrate and silver nanoparticles (AgNPs) formed with leaf extract SBGA.^{33,34} It can act as a very good antimicrobial agent as shown in Fig.-12.

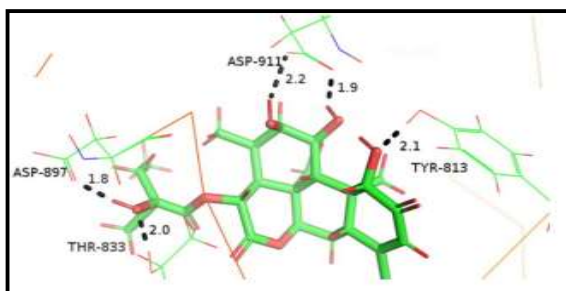


Fig.-11: Docking Image of Glaucarubinone against PI3K³²

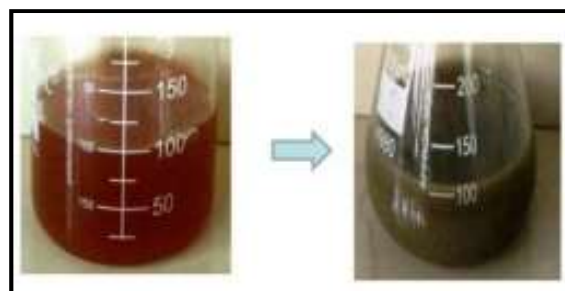


Fig.-12: Silver Nanoparticles from SBGA Leaf Extract³³

Medium polar ethyl acetate and non-polar petroleum ether solvent extracts of the leaves were carried for antibacterial, antioxidant by different methods.³⁵ Ethyl acetate extract exposed good antioxidant nature when compared to non-polar solvent extract. Using SBGA leaf extract, pure ZnO nanoparticles were prepared for the antioxidant and antimicrobial study.³⁶ The research revealed that the best biological

compatibility of the leaf extracted ZnO nanoparticles. SBGA DC effect on the growing plant induced additives and regulators through organogenesis which is shown in Fig.-13.³⁷

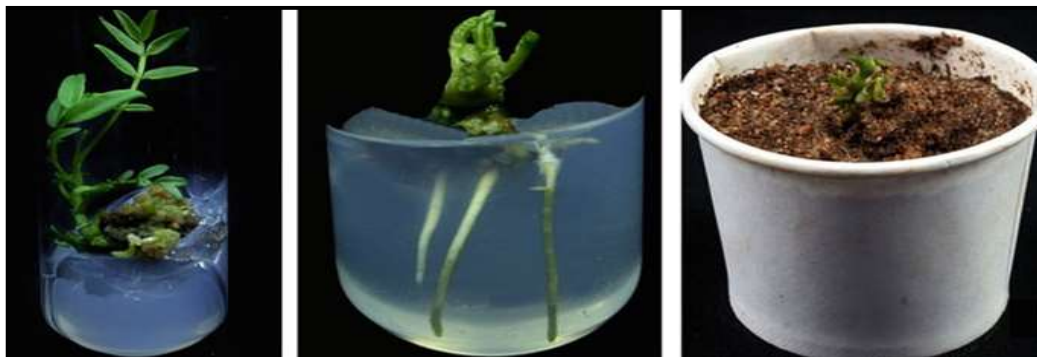


Fig.-13: Various Steps involved in SBGA Organogenesis³⁷

Screening of the ethanolic and methanolic crude extracts for the antifungal activity on *Fusarium oxysporum*, *Aspergillus parasiticus* respectively. SBGA extract showed good inhibition against *Aspergillus parasiticus*.³⁸ Leaves and bark of SBGA were collected for antibacterial activity against two pathogens such as *S. sobrinus* and *S. mutan*.³⁹ Alcoholic leaf extracts of Anticancer SBGA were phytochemically analysed and using Thin Layer Chromatography compositions were compared.⁴⁰ SBGA Bark, Flower and Leaves extracts were tested for their Anticancer Activity on Hct116 cancer Cells.⁴¹ The results revealed that the cell viability of chloroform extracts showed poor cell viability when compared to ethanol extracts- 35.8% at 50µg/ml and methanol extract -66.8% at 50 µg/ml. The extract of SBGA showed anti-inflammatory activity and the resultant outcomes exposed the dropping of inflammation.⁴² Antifungal activity of SBGA ethanolic extract against *Aspergillus parasiticus*.⁴³ was tested and showed good results. The silver nanoparticles were prepared by green synthesis using SBGA leaf which showed good resistivity against common bacteria and antioxidant character.⁴⁴ The chemical structure and the functional groups of SBGA component glaucarubin have been reported and ten oxygen atoms of the molecule exposed lactone, six hydroxyl and ester.⁴⁵ Using ethanol, aqueous and methanol SBGA bark extracts were carried for in-vitro antioxidant activity and exhibited promising activity.⁴⁶ Similarly, more stable with antimicrobial active gold nanoparticles were prepared from SBGA leaf extract and characterized.⁴⁷ SBGA callus (Fig.-14) and three places of the leaf (basal, middle, tip) random amplified polymorphic DNA was studied. The DNA sequence change was observed in SBGA due to media.⁴⁸ More quantity of biodiesel (methyl ester) was prepared from SBGA seed oil using alkaline catalyzed transesterification.⁴⁹ Both *A. Muricata* and SBGA leaf extracts were tested for *E. Faecalis* and the outcomes, not favored SBGA.⁵⁰ The various parts extracts of SBGA insecticidal activity (Fig.-15) were tested against *Plutella xylostella*, *Helicoverpa armigera* respectively. The report confirmed the highest mortality of the bark methanolic extract.⁵¹

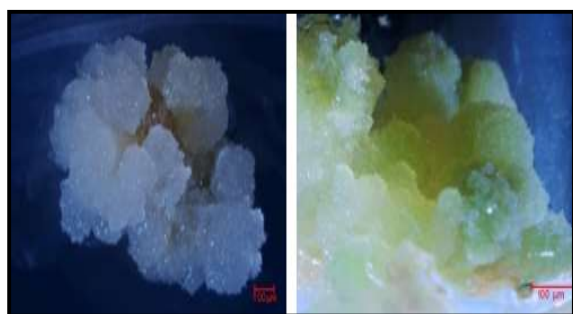


Fig.-14: Callus from 30 and 120 Days Culture⁴⁸

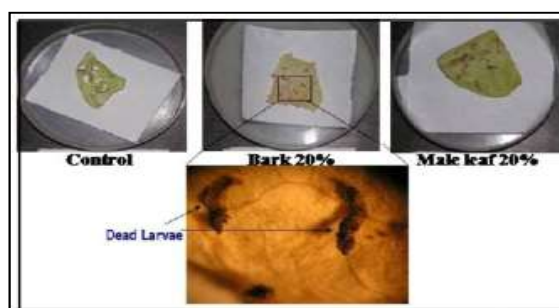


Fig.-15: Insecticidal Bioassay SBGA against *Plutella xylostella*⁵¹

Due to the various biological efficacy of SBGA cropping development carried in nurseries. The raised bed method and polybag (25 x 18 cm) are suitable for good cropping.⁵² In-vivo nephroprotective activity

of SBGA in albino Rats was tested which is treated with gentamycin induction. The methanolic extract exposed a good effect in nephrotoxicity in test species.⁵³ An effect of fungus pathogens *G. mosseae* plus, *B. coagulans* on SBGA nutrition was studied and they enhanced the nutrients, growth respectively.⁵⁴ The low concentration acetone extract of SBGA leaf controlled the *Allium cepa* L (mitotic) cells.⁵⁵ Similar way, green synthesis of copper nanoparticles was reported from *Uncaria gambir* ROXB.⁵⁶ Nowadays various plant extracts were used for medicinal uses including leaf extract of *Coriandrum Sativum*.⁵⁷

CONCLUSION

From the various reports and reviews, this work observed and noticed the importance of the selected plant SBGA in the medical field. But, some of the extracts and their usefulness are not clearly explained still now. Most of the research papers reported the phytochemical analysis with biological activity of the SBGA plant. Only a few research reports have characterized the isolated extracts using different analytical techniques. The limited reports of SBGA are observed on therapeutic effectiveness and usefulness of synthesized nanoparticles by the pollution-free method. Extraction methods, solvent effect, chemical constituents, nanoparticle preparations and the applications in a wide range of anti-microbes are lacking. Hence this will be useful for future researchers.

REFERENCES

1. M. Pandey, S. Rastogi, S., and A.K.S. Rawat, *Evidence-Based Complementary and Alternative Medicine*, 1(2013), DOI:10.1155/2013/376327
2. S. Parasuraman, *Current Pharmacogenomics and Personalized Medicine*, **16**, 63(2018), DOI: 10.2174/1875692116666180419153313
3. J. Rout, A.L. Sajem and M. Nath, *Indian Journal of Traditional Knowledge*, **11**(3), 520(2012)
4. M. Ekor, *Front Pharmacology*, **177**, 1(2014)
5. A.H. Hussin, *Malaysian Journal of Pharmacy*, 39(2001)
6. B. Petrovska, *Pharmacognosy Reviews*, **6**, 1(2012), DOI:10.4103/0973-7847.95849
7. J. Antony, A.Thomas, D.Gnanasekaran and H.Elizabeth, *International Journal of New Technology and Research*, **2**(10), 59(2016)
8. S.Joshi and S.Joshi, *University of Agricultural sciences, Bangalore and Indian council of Agricultural Research*, New Delhi, India, 86(2002)
9. I. Arivu, M. Muthulingam, and R. Palaniappan, *Journal of Advanced Scientific and Technical Research*, **1**(7), 86(2017)
10. A. Jose, A.Chaitanya, E. Kannan, and S.V. Madhunapantula, *Frontiers in Pharmacology*, 9(2018), DOI:10.3389/fphar.2018.00127
11. I.P. Sridevi, S.C. Ghagane, R.B. Nerli, S.S. Jalalpure, and M.B. Hiremath, *Pharmacognacy Journal*, **9**, 906(2017), DOI:10.5530/pj.2017.6.142
12. K.Santhanalakshmi, D. Sangeetha, S.Sivamani, M.Tamilarasan, T.P Rajesh, and B.Anandraj. *International Journal of Pharmaceutical Sciences and Research*. **5**(2), 432(2014)
13. B.M. Rajurkar, *International Journal of Research and Review in Pharmacy and applied science*, **1**(4), 278(2011)
14. M.S. Patil and D.Gaikwad, *Journal of Pharmaceutical Science and Research*, **3**, 1195(2011)
15. M.P. Padovan, V.J. Cortez, L.F. Navarrete, E.D. Navarrete, A.C. Deffner, L.G. Centeno, and B. Rapidel, *Agroforestry Systems*, **89**(5), 857(2015), DOI:10.1007/s10457-015-9820-z
16. G. Vaidya, G. R. Naik, *International Journal of Current Research Reviews*, **10**(5), 46(2018), DOI: 10.7324/IJCR R. 2018.1058
17. R. L. Chavan and B. V. Tembhurne, *Karnataka Journal of Agricultural Science*, **28**(2), 235(2015)
18. V.K.Sharanya, K. Gayathiri, M. Sangeetha and G.Shyam Prakash, G.S.Kumar, J. V. Vathini and R.S. Kavimani, *International Journal of Pharma Research & Review*, **5**(6), 32(2016)
19. V.F. Assendelft, D.T. Mintz, J.A.Schack, P. Ottolenghi, and H.Most, *American Journal of Tropical Medicine and Hygiene*, **5**(3), 501(1956)
20. G. Kartha, D. J. Haas, H.M. Schaffer, and K.K. Kaistha, *Nature*, **202** (4930), 389(1964), DOI: 10.1038/202389b0

21. R. Duffy, C. Wade, and Chang, R, *Drug Discovery Today*, **17**, 942(2012), DOI: [10.1016/j.drudis.2012.03.013](https://doi.org/10.1016/j.drudis.2012.03.013)
22. J. F. Rivero-Cruz, R. Lezutekong, T. Lobo-Echeverri, A. Ito, Q. Mi, H. B. Chai, D. D. Soejarto, G. A. Cordell, J. M. Pezzuto, S. M. Swanson, I. Morelli and A. D.Kinghorn, *Phytotherapy Research*, **19(2)**, 136(2005), DOI:[10.1002/ptr.1642](https://doi.org/10.1002/ptr.1642)
23. S. Bhatnagar, J. Polonsky, T. Prange and C. Pascard, *Tetrahedron Letters*, **25(3)**, 299(1984), DOI: [10.1016/s0040-4039\(00\)99867-8](https://doi.org/10.1016/s0040-4039(00)99867-8)
24. L. Simon, P. Narayanaswamy, and S. Joshi, S. *The Journal of Horticultural Science and Biotechnology*, **84(5)**, 510(2009), DOI:[10.1080/14620316.2009.11512557](https://doi.org/10.1080/14620316.2009.11512557)
25. N.Sajeeda, K.Rr, S. Shivakumara, Y. Shivaraj, and C.S. Karigar, *Asian Journal of Pharmaceutical and Clinical Research*, **12(9)**, 56(2019)
26. Iasmine A.B.S. Alvesa, Henrique M. Mirandab, Luiz A.L. Soaresa and K.P. Randau, *Revista Brasileira de Farmacognosia*, **24**, 481(2014)
27. A.Kumar, G. Tyagi, Sunayana Sharma, Vikas Kumar and Reena Pundir, *Indian journal of Pharmacology*, **1(12)**, 735(2014)
28. C.K. Prajapati, M.N. Reddy and M.H. Bhatt, *International Journal of Botany Studies*, **3(2)**, 52(2018)
29. F.C. Valdés, A, M. Martinez, J. S. Lizama, R. Vermeersch, M. Cos, P.L. Maes, *Memorias Do Instituto Oswaldo Cruz*, **103(6)**, 615(2008), DOI:[10.1590/s0074-02762008000600019](https://doi.org/10.1590/s0074-02762008000600019)
30. B. Vikas, B.S. Akhil, S.R. Suja, and K. Sujathan, *Asian Pacific Journal of Cancer Prevention*, **18(7)**, 1765(2017), DOI:[10.22034/APJCP.2017.18.7.1765](https://doi.org/10.22034/APJCP.2017.18.7.1765)
31. S.Dinesh, D.S.N. Sasikumar, B. Girija, V.Lakshmipriya, Panicker, P.V.Kumar, S.Preetha and S.S.Sarma, *Journal of Applied Pharmaceutical Science*, **7(09)**, 142(2017), DOI:[10.7324/JAPS.2017.70919](https://doi.org/10.7324/JAPS.2017.70919)
32. K. S. Ramya, S. Iqbal, K.Gunasekaran, A.Radha, *Research Journal of Life Sciences, Bioinformatics, Pharmaceutical and Chemical Sciences*, 218(2018)
33. C. P. Chandrappa, N. Chandrasekar, M. Govindappa, C. Shanbhag, U. K. Singh, J. Masarghal, *International Journal of Current Pharmaceutical Research*, **9(4)**, 19(2017)
34. Ramesh, L. Cathrine, S. Gurupriya P. Pratheema, *International Journal for Research in Applied Science & Engineering Technology*, **6(III)**, 933(2018)
35. S. Kochath, S.A.Venugopal, M.C.Radhakrishnan, *South Indian Journal of Biological Sciences*, **2(1)**, 119(2016)
36. H. Kumar, N.K. Murali, M. Satish, A.B. Singh, S. Gowtham, H. G. Mahesh and H. M, Jagannath, *Journal of Cluster Science*, **31(2)**, 523(2020), DOI: [10.1007/s10876-019-01669-7](https://doi.org/10.1007/s10876-019-01669-7)
37. A.R. Lavanya, M. Muthukumar, S. Muthukrishnan, V. Kumaresan, T. S. Kumar, M. V. Venkatesh, and M. V. Rao, *Conservation and Crop Improvement*, **71**(2016), DOI:[10.1007/978-981-10-1917-3_4](https://doi.org/10.1007/978-981-10-1917-3_4)
38. K. Mikawlawng, S. Kaushik, A. K. Pushker, S. Kumar, M. K. Singh and G. S. Sharma, *Journal of Medicinal Plants Studies*, **2(3)**, 1(2014)
39. H.A. Salman, R.Senthilkumar and M. Vasundhara, *Biosciences Biotechnology Research Asia*, **15(2)**, 311(2018)
40. A. Kumar, V. Rawat, Amardeep and V. Kumar, *International Journal of Current Microbiology and Applied Sciences*, **5(6)**, 679 (2016)
41. S. Niketh, SL. Shivakumar, R.R. Kolgi, and C.S. Karigar, *International Journal of Scientific & Technology Research*, **8(10)**, 69(2019)
42. N. T.Abdullah, N. H. Khair and R. Koneri, *World Journal of Pharmaceutical Research*, **7(15)**, 550 (2018), DOI: [0.20959/wjpr201815-12963](https://doi.org/0.20959/wjpr201815-12963)
43. S. Kaushik, K. Mikawlawng, N.G. Sonone, S. Subramaniam and A. K. Choudhary, *International Journal of Current Research*, **6 (03)**, 5677(2014)
44. B. Hemashekhar, C. P. Chandrappa, M. Govindappa, N. Chandrasekhar, N. Ganganagappa and Y.L. Ramachandra, *Int. Journal of Engineering Research and Application*, **7 (8)**, 17(2017)
45. E.A. Ham, H. M. Schafer, R.G. Denkwalter, and N.G. Brink, *Journal of the American Chemical Society*, **76(23)**, 6066(1954), DOI: [10.1021/JA01652A060](https://doi.org/10.1021/JA01652A060)

46. S.D.E. Osagie-Eweka, *African Journal of Plant Science*, **12(1)**, 7(2018), DOI: [10.5897/AJPS2017.1547](https://doi.org/10.5897/AJPS2017.1547)
47. N. Thangamani and N. Bhuvaneshwari, N, *Chemical Physics Letters*, **732**, 2614(2019), DOI: [10.1016/J.CPLETT.2019.07.015](https://doi.org/10.1016/J.CPLETT.2019.07.015)
48. J Kakuturu, PC Josekutty, S Potlakayala, M Reitzel, K Salim, S Charyulu, R Adeyiga, S Menon, SL Goldman, P Patel, MJ Chorney and S Rudrabhatla, *African Journal of Biotechnology*, **13(53)**, 4766 (2014)
49. S.R. Mishra, M.K. Mohanty, S.P. Das and A.K. Pattanaik, *Research Journal of Chemical Sciences*, **2(5)**, 66(2012)
50. J. Mathew, R. George, R. Theruvil, T.C. Padavil, L. Tomy and A. Kurian, *The Journal of Contemporary Dental Practice*, **17(8)**, 650(2016)
51. S.S. Bangar, M.S. Dudhare, A.G. Deshmukh and H.A. Wagh, *Agriculture Update*, **12**, 436 (2017)
52. R. L. Chavan and V. Tembhurne, *Karnataka Journal of Agriculture Science* **28(2)**, 235 (2015)
53. N.T.Abdullah, R. Koneri and D. K. Jha, *Journal Pharaceutical Sciences Review and Research*, **51(1)**, 182(2018)
54. G.L. Sailo, and D.J. Bagyaraj, *Biological Agriculture & Horticulture*, **20(4)**, 339(2003), DOI: [10.1080/01448765.2003.9754977](https://doi.org/10.1080/01448765.2003.9754977)
55. S.V. Ajith, Chandralekha and C.T, BinuThomas, *World Wide Journal Of Multidicipinary Research And Development*, **3(7)**, 35(2017)
56. N. Elisma , A. Labanni , Emriadi , Y. Rilda , M. Asrofi and S. Arief, *Rasayan Journal of Chemistry*, **12 (4)**, 1752(2019), DOI: [10.31788/RJC.2019.1245347](https://doi.org/10.31788/RJC.2019.1245347)
57. S.M. Sinaga, G. Haro, S. Sudarmi and Iksen, *Rasayan Journal of Chemistry*, **12(4)**, 1992(2019), DOI: [10.31788/RJC.2019.1245451](https://doi.org/10.31788/RJC.2019.1245451)

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