



Analgesic, hypoglycemic, sedative, antidiarrheal, thrombolytic, anti-Inflammatory activity of *Syzygium fruticosum* DC

Md. Hasan Ali^{1,3}, Sauda Sultana Mimi^{*1}, Sadia Afrin Chhanda¹, Mohammad Mahmudul Hasan¹, Md. Omar Sha Rafi¹, Nusrat Tabassum Shristy¹, Mohammad A. Rashid², Tanvir Muslim¹

¹Department of Chemistry, University of Dhaka, Dhaka -1000, Bangladesh

² Phytochemical Research Laboratory, Department of Pharmaceutical Chemistry, Faculty of Pharmacy, University of Dhaka, Dhaka-1000, Bangladesh

³Bangladesh Council of Scientific and Industrial Research (BCSIR)

*Corresponding author: saudamimi1549@gmail.com
(accepted December 17, 2024)

ABSTRACT

Context: *Syzygium fruticosum* DC. is a medicinal plant of Bangladesh. **Objectives:** testing analgesic, hypoglycemic, sedative, antidiarrheal, thrombolytic, and anti-inflammatory activities *in vivo*. **Methods:** organic extracts were given to swiss albino. **Results:** extracts exhibited various degrees of activities in all tests performed. **Conclusion:** our findings provide scientific validation for the plant's extensive traditional use in the Chittagong Hill tracts.

Keywords: medicinal plants, Bangladesh

INTRODUCTION

Plants, often regarded as the "Almighty's gift," hold a unique and indispensable role in the natural world (Ozioma et al., 2019). They provide numerous health benefits, forming a cornerstone of human survival and well-being (Kumar et al., 2012; Hamburger et al., 1991). Life on Earth would be unsustainable without plants, as they supply oxygen, food, shelter, and raw materials (Mancuso et al., 2015; Modak et al., 2007). Since ancient times, plants have been central to health systems, valued in both traditional and modern medicine for their natural therapeutic properties (Dias et al., 2012). Medicinal plant extracts preserve active secondary metabolites with therapeutic effects on organisms (Ehrman et al., 2007). Bangladesh, especially Sylhet, Chittagong Hill tract, and Sundharban are endowed with a vast source of medicinal plants (Saifullah et al., 2021; Ahmed et al., 2021). Most of the medicinal plants in Bangladesh have not been studied (Haque et al., 2020). One of these is *Syzygium fruticosum* locally

known as khudijam (Elliot et al., 1990; Nigam et al., 2012; Chadni et al., 2015; Nasrin et al., 2018; Moni et al., 202; Uddin et al., 2022).

METHODS

Plant material

The mixture of ripe (blackish purple) and unripe (green) fruits of *S. fruticosum* was accumulated from the botanical garden of Jahangirnagar University, Savar. The plant and its fruit were authenticated (DACB-87367) by a scientific officer, Shaharina Hasin in Bangladesh National Herbarium (BNH), Mirpur, Dhaka, Bangladesh. Fruits were washed, sun-dried, ground into powder, and 480 g dried sample was dissolved in 1.7 liters of Methanol for 15 days. The resulting Methanol soluble Partition was filtered and dried using a rotary evaporator. 5 g dried methanolic extract was taken for partitioning into different solvents such as n-Hexane, Dichloromethane, Ethyl acetate and Aqueous by following modified Kupchan process (Emran et al., 2015). These Methanol soluble Partition (MESP), Hexane soluble partition (HSP), Dichloromethane soluble partition (DCMSP), Ethyl acetate soluble partition (EASP), and Aqueous soluble partition (AqSP) were investigated.

Animal study

Swiss albino mice, aged between four and five weeks, consisting of both males and females, were kept in the Pharmacy Department at the State University of Bangladesh for pharmacological research. Prior to the study, ethical approval was obtained from the Ethics Review Committee of the same department. The approved Animal Ethics Number is 2024-03-29/SUB/I-ERC/003. The mice were housed and maintained in plastic polypropylene cages under typical animal housing configurations (temperature $24\pm 2^{\circ}\text{C}$), humidity levels ranging from 60%-70% and dark-light cycles (12/12) h the mice had limitless access to water ad libitum and conventional laboratory feed from Hindustan Animal Feeds. Animal study design according to Ali et al., 2024.

Central analgesic activity

Tail immersion method (Jahan et al., 2024).

Peripheral analgesic activity

Chemical nociception model (Ajaib et al., 2024).

Hypoglycemic activity

Glucose tolerance test (Ming et al., 2024).

Sedative activity

Thiopental Sodium-induced sleeping assay (Jahani et al., 2022).

Anti-diarrheal activity

Castor-oil induced method (Rahman et al., 2020).

Thrombolytic activity

Thrombolytic method (Ramjan et al., 2014).

Hemolysis activity

Hypotonicity and heat-induced hemolysis methods (Yesmin et al., 2020; Aidoo et al., 2021).

Statistical analysis

Paired t-test analysis utilizing the GraphPad software, prism-10 edition (prism-10 for Windows, Version 10.0, GSL Biotech, California, USA). The data are presented as the mean \pm SD. Standard Error Mean (SEM) was also recorded. The mean variation between the positive and negative control was deemed substantial at P values < 0.01 and 0.005 .

RESULTS

Central analgesic activity

MESP at the dose of 600 mg/kg evoked after 90 mins a prolongation of time of reaction to pain of 620% (Figure 1)

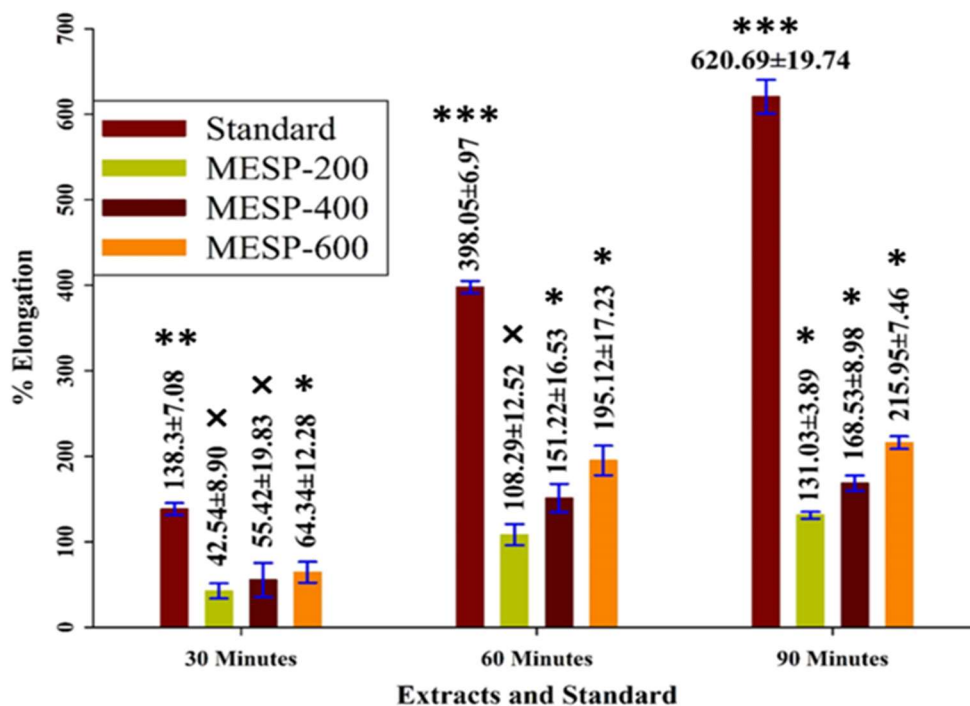


Figure 1: Central analgesic activity

% Time of standard and extracts at different time intervals. Values are expressed as mean \pm SD (n = 4); ***p < 0.001 are very statistically significant, *p < 0.05 statistically significant and Xp > 0.05 compared to control followed by Dunnet test (GraphPad Prism 10). MESP = Methanol Soluble Partition

Peripheral analgesic activity

Methanol extract at the dose of 600 mg/kg reduced writhing by 70.4% (Figure 2)

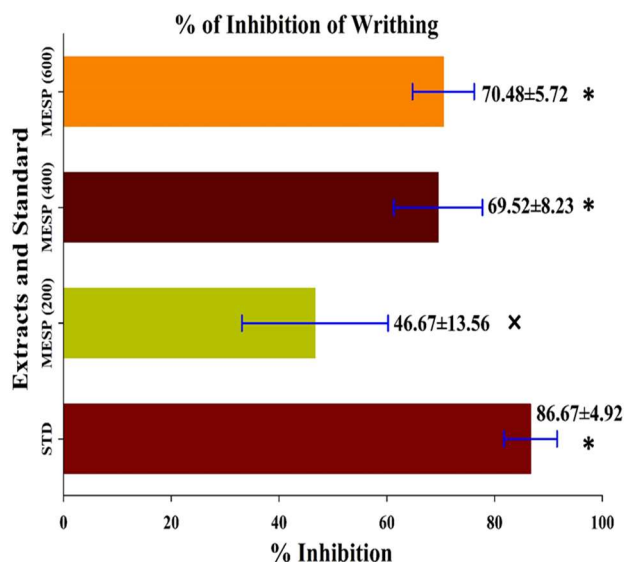


Figure 2: % Inhibition of writhing

Values are expressed as mean ± SD (n = 4); *p < 0.05 statistically significant and x p > 0.05 compared to control followed by Dunnet test (GraphPad Prism 10). MESP = Methanol Soluble Partition

Hypoglycemic activity

MESP at the dose of 400 mg/kg decreased postprandial glycemia from 12.6 (control) to 7.7 mmol/L at 30 min (Table 1, Figure 3).

Table 1. Hypoglycemic activity

Code No.	Plasma level of glucose (Mean ± SD)/ (mmol/L)			
	00 min	30 min	60 min	120 min
CTL	6.77±0.98	12.69±5.55	9.04±3.69	6.81±0.58
STD	3.55±0.31	7.12±2.03	5.18±1.63	3.38±0.71
MESP (200)	5.05±0.34	9.625±1.68	7.35±1.60	6.05±0.71
MESP (400)	4.89±0.53	7.78±1.04	6.92±0.85	5.36±0.49
MESP (600)	5.25±0.64	7.975±0.62	6.45±1.45	4.75±0.53

CTL= Negative control, STD= Standard Drug

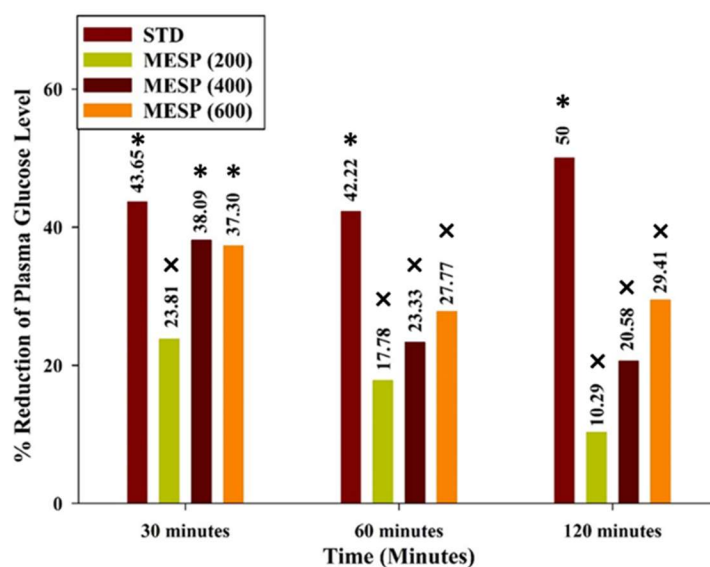


Figure 3: Hypoglycemic activity

Values are expressed as mean \pm SD (n = 4); *p < 0.05 statistically significant and x p > 0.05 compared to control followed by Dunnet test (GraphPad Prism 10). MESP = Methanol Soluble Partition

Sedative activity

MESP at the dose of 600 mg/kg was the most active (Figure 4).

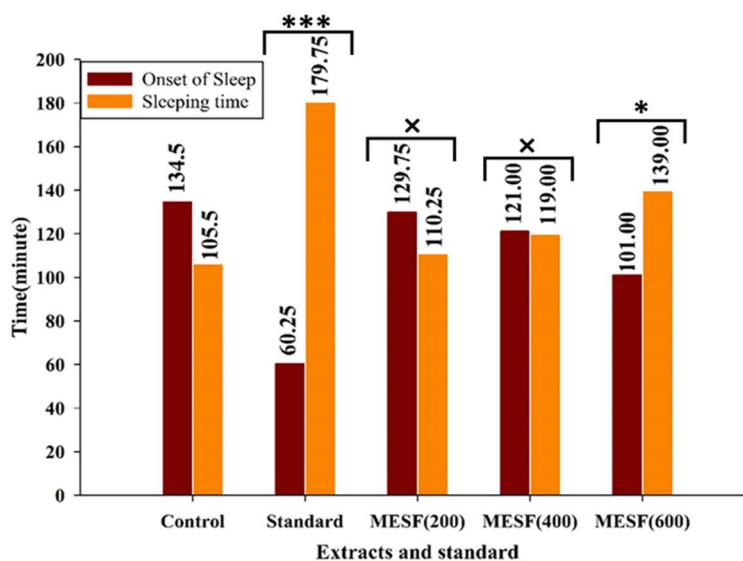


Figure 4: Sedative activity

Values are expressed as mean \pm SD (n = 4); ***p < 0.001 are very statistically significant, *p < 0.05 statistically significant and x p > 0.05 compared to control followed by Dunnet test (GraphPad Prism 10). MESP = Methanol Soluble Partition

Anti-diarrheal activity

MESP at the dose of 600 mg/kg reduced diarrheal discharges by 61.1% (Table 2)

Table 2: Anti-diarrheal activity

Code No.	No of diarrheal feces (Mean)	% Reduction of diarrhea	Standard deviation SD	Standard Error of mean	t-test value	P value
STD	1.5	83.33	1	0.50	5.4232	0.0016
MESP (200)	4.25	52.77	2.5	1.25	2.6444	0.0006
MESP (400)	3.5	61.11	1	0.5	3.9754	0.0073
MESP (600)	3.0	66.67	2.16	1.08	3.5663	0.0118

Thrombolytic activity

EASP evoked a 19.4% of cloth lysis (Table 3)

Table 3. Thrombolytic activity

Fractions	% Of clot lysis (w5/w4) x 100
HSP	9.03
DCMSP	7.20
EASP	19.42
AQSP	19.10
MESP	9.31
Blank	5.22
streptokinase	65.67

Hemolysis activity

EASP was the most active (66.4 %) in the in hypotonic solution-induced hemolysis test (Figure 5) while MESP was the most active extract in the induced hemolysis test (74.5 %) (Figure 5).

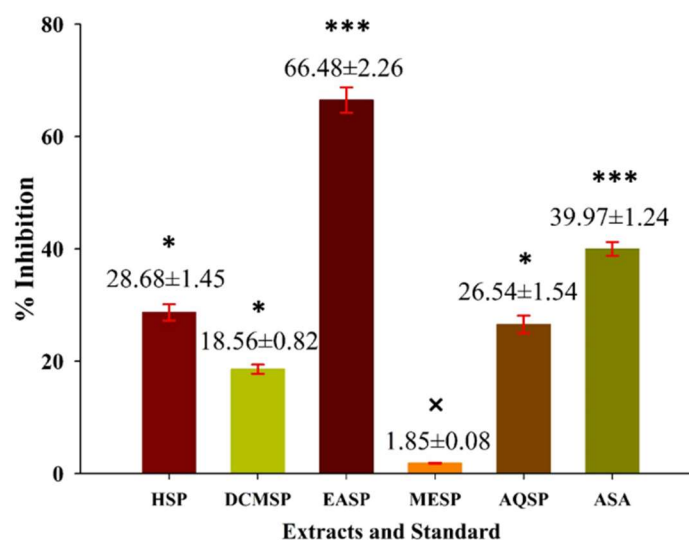


Figure 5: Membrane Stabilization activity of in hypotonic solution
Values are expressed as mean ± SD (n = 4); ***p < 0.001 are very statistically significant, *p < 0.05 statistically significant and x p > 0.05 compared to control followed by Dunnet test (GraphPad Prism 10).

CONCLUSION

Our study provide preliminary evidence that *S. fruticosum* has analgesic, hypoglycemic, sedative, antidiarrheal, thrombolytic, and anti-inflammatory activities *in vivo*. These findings provide scientific validation for the plant's extensive traditional use in the Chittagong Hill Tracts.

DECLARATION OF CONFLICT OF INTEREST

No conflict of interest to declare.

DECLARATION OF HONOUR

We declare in our honor that our results are not fake and made up.

ACKNOWLEDGMENT

Md. Hasan Ali is grateful to both Bose Centre for Advanced Study and Research in Natural Sciences at University of Dhaka, Dhaka-1000 for Bose Fellowship and Ministry Science and Technology for National Science and Technology (NST) Fellowship. Sauda Sultana Mimi and all Co-authors are thankful to the Department of Chemistry, University of Dhaka, Dhaka-1000 for using Lab Facilities.

REFERENCES

- Ahmed, S.R., Rabbee, M.F., Roy, A., Chowdhury, R., Banik, A., Kubra, K., Baek, K.H. 2021. Therapeutic promises of medicinal plants in Bangladesh and their bioactive compounds against ulcers and inflammatory diseases. *Plants*, 10(7), 1348.
- Aidoo, D. B., Konja, D., Henneh, I. T., & Ekor, M. 2021. Protective effect of bergapten against human erythrocyte hemolysis and protein denaturation in vitro. *International Journal of Inflammation*, 2021(1), 1279359.
- Ajaib, M., Kamran, S. H., Siddiqui, M. F., Qasim, M., Azeem, M., Abideen, Z., El-Keblawy, A. 2024. Exploring the phytochemical, antioxidant, antimicrobial and analgesic potentials of *Solanum elaeagnifolium* as an alternative biological feedstock for producing sustainable biochemicals. *Biocatalysis and Agricultural Biotechnology*, 58, 103183.
- Ali, M. H., Islam, S., Hasan, M. M., Rahman, M. H., Rahman, A., Taher, M. A., Muslim, T. 2024. Studies of phytochemical analysis, in vitro and in vivo evaluation of the local *Pseudoelephantopus scipatus* plant. *African Journal of Pharmacy and Pharmacology* 18(7),114-126.
- Chy, M.N.U., Adnan, M., Chowdhury, M.R., Pagano, E., Kamal, A.T.M.M., Oh, K.K., Cho, D.H., Capasso, R. 2021. Central and peripheral pain intervention by *Ophiorrhizarugosa* leaves: Potential underlying mechanisms and insight into the role of pain modulators. *J Ethnopharmacol*, 276;114182.
- Dias, D.A., Urban, S., Roessner, U. 2012. A historical overview of natural products in drug discovery. *Metabolites*, 2(2), 303–336.
- Ehrman, T.M., Barlow, D.J., Hylands, P.J. 2007. Phytochemical databases of Chinese herbal constituents and bioactive plant compounds with known target specificities. *J Chem Inf Model*. 47, 254–63.
- Elliot, W.R., Jones, D.L. 1990. Encyclopaedia of Australian Plants Suitable for Cultivation; *Lothian Publishing Company Pty Ltd.: Port Melbourne, Australia*. 5, 512.
- Emran, T.B., Rahman, M.A., Uddin, M.M., Rahman, M.M., Uddin, M.Z., Dash, R., Layzu, C. 2015. Effects of organic extracts and their different fractions of five Bangladeshi plants on in vitro thrombolysis. *BMC Complement Altern Med*, 15:128.
- Hamburger, M., Hostettmann, K. 1991. Bioactivity in plants: the link between phytochemistry and medicine. *Phytochemistry*, 30(12), 3864-3874.
- Haque, M.R., Islam, M., Kuddus, M.R. 2020. In vitro and in vivo evaluation of pharmacological potential of *Begonia barbata* Wall. *Futur J Pharm Sci* 6, 112.
- Ichihara, Y.K., Shiraishi, Y., Kohsaka, S., Nakano, S., Nagatomo, Y., Ono, T., Takei, M., Sakamoto, M., Mizuno, A., Kitamura, M., Niimi, N. 2023. Association of pre-hospital precipitating factors with short-and long-term outcomes of acute heart failure patients: A report from the WET-HF2 registry. *International Journal of Cardiology*, 389, p.131161.
- Jahan, I., Ali, M. H., Shristy, N. T., Rafi, M. O. S., Mimi, S. S., Siddik, M. N. A., Chowdhury, T. A. 2024. Assessing Therapeutic Potentials of *Catharanthus roseus* (L.) G. Don Focusing on Anti-Diabetic, Analgesic and Anti-Diarrheal Activities. *Bangladesh Pharmaceutical Journal*, 27(2), 215-222.
- Jahani, R., Behzad, S., Saffariha, M., Tabrizi, N.T., Faizi, M. 2022. Sedative-hypnotic, anxiolytic and possible side effects of *Salvia limbata* C. A. Mey. Extracts and the effects of phenological stage and altitude on the rosmarinic acid content, *Journal of Ethnopharmacology*, 282;114630.
- Kumar, K.S., Umadevi, M., Bhowmik, D., Singh, D.M., Dutta, A.S. 2012. Recent trends in medicinal uses and health benefits of Indian traditional herbs *Aegle marmelos*. *The Pharma Innovation*, 1(4).
- Mancuso, S., Viola, A. 2015. Brilliant green: the surprising history and science of plant intelligence. Island Press.
- Modak, M., Dixit, P., Londhe, J., Ghaskadbi, S., Devasagayam, T.P.A. 2007. Indian herbs and herbal drugs used for the treatment of diabetes. *Journal of clinical biochemistry and nutrition*, 40(3), 163-173.
- Moni, J.N.R., Adnan, M., Tareq, A.M., Kabir, M.I., Reza, A.S.M.A., Nasrin, M.S., Chowdhury, K.H., Sayem, S.A.J., Rahman, M.A., Alam, A.K. 2021. Therapeutic Potentials of *Syzygium fruticosum* Fruit (Seed) Reflected into an Array of Pharmacological Assays and Prospective Receptors-Mediated Pathways. *Life*. 11(2),155.

- Nasrin, M.S., Mostofa, M.G., Harun-Or-Rashid, M., Islam, M.S., Khurshid, A.H.M.** 2018. Antioxidant, free radical scavenging, antibacterial and cytotoxic compound from the leaves of *Syzygium fruticosum*. *Int. J. Pharma Sci. Sci. Res.* 4, 69–73.
- Nigam, V., Nigam, R., Singh, A.** 2012. Distribution and medicinal properties of *Syzygium* species. *Current Research in Pharmaceutical Sciences*, 2(2), 73-80.
- Ozioma, E.O.J. and Chinwe, O.A.N.** 2019. Herbal medicines in African traditional medicine. *Herbal medicine*. 10, 191-214.
- Rahman, A., Hasan, M.M., Taher, M.A., Muslim, T.** 2020. Analgesic, Antidiarrheal and CNS-depressant Activities of *Flemingia macrophylla* (Willd.). *Bangladesh Pharmaceutical Journal*, 23(2), 141–145.
- Ramjan, A., Hossain, M., Runa, J.F., Md, H., Mahmodul, I.** 2014. Evaluation of thrombolytic potential of three medicinal plants available in Bangladesh, as a potent source of thrombolytic compounds. *Avicenna J Phytomed*, 4(6):430-6.
- Saifullah, M. and Jewel, K.N.A.** 2021. Sustainable Management of Non-wood Forest Products for Rural Livelihoods in Bangladesh. *Nonwood and Livelihood: Sustainable Management of Non-wood Forest Products for Rural Livelihoods in South Asia. SAARC Agriculture Centre, SAARC, Dhaka, Bangladesh*, 166(32).
- Uddin, A.B.M.N., Hossain, F., Reza, A.S.M.A., Nasrin, M.S., Alam. A.H.M.K.** 2022. Traditional uses, pharmacological activities, and phytochemical constituents of the genus *Syzygium*: A review. *Food Sci Nutr*.10(6),1789-1819.
- Yesmin, S., Paul, A., Naz, T.** 2020. Membrane stabilization as a mechanism of the anti-inflammatory activity of ethanolic root extract of Choi (*Piper chaba*). *Clin Phytosci* 6, 59.