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Variation in the metabolites and alpha-glucosidase inhibitory activity of *Cosmos caudatus* at different growth stages

By: [Wan-Nadilah, WA](#) (Wan-Nadilah, Wan Ahmad)^[1,2]; [Akhtar, MT](#) (Akhtar, Muhammad Tayyab)^[2,6]; [Shaari, K](#) (Shaari, Khozirah)^[2]; [Khatib, A](#) (Khatib, Alfi)^[3]; [Hamid, AA](#) (Hamid, Azizah Abdul)^[4]; [Hamid, M](#) (Hamid, Muhajir)^[5]

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Abstract

Background *Cosmos caudatus* is an annual plant known for its medicinal value in treating several health conditions, such as high blood pressure, arthritis, and diabetes mellitus. The alpha-glucosidase inhibitory activity and total phenolic content of the leaf aqueous ethanolic extracts of the plant at different growth stages (6, 8, 10, 12 and 14 weeks) were determined in an effort to ascertain the best time to harvest the plant for maximum medicinal quality with respect to its glucose-lowering effects. Methods The aqueous ethanolic leaf extracts of *C. caudatus* were characterized by NMR and LC-MS/MS. The total phenolic content and alpha-glucosidase inhibitory activity were evaluated by the Folin-Ciocalteu method and alpha-glucosidase inhibitory assay, respectively. The statistical significance of the results was evaluated using one-way ANOVA with Duncan's post hoc test, and correlation among the different activities was performed by Pearson's correlation test. NMR spectroscopy along with multivariate data analysis was used to identify the metabolites correlated with total phenolic content and alpha-glucosidase inhibitory activity of the *C. caudatus* leaf extracts. Results It was found that the alpha-glucosidase inhibitory activity and total phenolic content of the optimized ethanol:water (80:20) leaf extract of the plant increased significantly as the plant matured, reaching a maximum at the 10th week. The IC₅₀ value for alpha-glucosidase inhibitory activity (39.18 μg mL⁻¹) at the 10th week showed greater potency than the positive standard, quercetin (110.50 μg mL⁻¹). Through an H-1 NMR-based metabolomics approach, the 10-week-old samples were shown to be correlated with a high total phenolic content and alpha-glucosidase inhibitory activity. From the partial least squares biplot, rutin and flavonoid glycosides, consisting of quercetin 3-O-arabinofuranoside, quercetin 3-O-rhamnoside, quercetin 3-O-glucoside, and quercetin 3-O-xyloside, were identified as the major bioactive metabolites. The metabolites were identified by NMR spectroscopy (J-resolve, HSQC and HMBC experiments) and further supported by dereplication via LC-MS/MS. Conclusion For high phytomedicinal quality, the 10th week is recommended as the best time to harvest *C. caudatus* leaves with respect to its glucose lowering potential.

Keywords

Author Keywords: *Cosmos caudatus*; alpha-Glucosidase inhibition; Total phenolic content; Harvesting age; Quercetin derivatives; NMR-based metabolomics

KeyWords Plus: ANTIOXIDANT ACTIVITY; PLANT-EXTRACTS; L. FRUIT; IN-VITRO; ANTIHYPERGLYCEMIC ACTIVITY; ANTIDIABETIC AGENTS; METHANOLIC EXTRACT; AQUEOUS EXTRACTS; PHENOLIC CONTENT; METABOLOMICS

Author Information

Reprint Address: Shaari, K (reprint author)

+ Univ Putra Malaysia, Inst Biosci, Lab Nat Prod, Serdang, Selangor, Malaysia.

Addresses:

+ [1] Univ Sultan Zainal Abidin, Fac Bioresources & Food Ind, Besut, Terengganu, Malaysia

+ [2] Univ Putra Malaysia, Inst Biosci, Lab Nat Prod, Serdang, Selangor, Malaysia

+ [3] Int Islamic Univ Malaysia, Kulliyah Pharm, Kuantan, Pahang, Malaysia

+ [4] Univ Putra Malaysia, Fac Food Sci & Technol, Serdang, Selangor, Malaysia

+ [5] Univ Putra Malaysia, Fac Biotechnol & Biomol Sci, Serdang, Selangor, Malaysia

+ [6] Govt Coll Univ, Inst Ind Biotechnol, Lahore, Pakistan

E-mail Addresses: khodzrah@yahoo.com.my

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