Contents lists available at ScienceDirect

## Phytochemistry

journal homepage: www.elsevier.com/locate/phytochem

# *trans*-Tiliroside: A potent $\alpha$ -glucosidase inhibitor from the leaves of *Elaeagnus angustifolia* L

Hafize Yuca<sup>a,b,\*</sup>, Hilal Özbek<sup>a,b</sup>, Lütfiye Ömür Demirezer<sup>c</sup>, Handan Gökben Kasil<sup>a</sup>, Zühal Güvenalp<sup>a,b</sup>

<sup>a</sup> Department of Pharmacognosy, Faculty of Pharmacy, Ataturk University, 25240, Erzurum, Turkey

<sup>b</sup> Medicinal and Aromatic Plant and Drug Research Center, Ataturk University, 25240, Erzurum, Turkey

<sup>c</sup> Department of Pharmacognosy, Faculty of Pharmacy, Hacettepe University, 06100, Ankara, Turkey

#### ARTICLE INFO

Keywords: Elaeagnus angustifolia Elaeagnaceae a-Glucosidase inhibition Radical scavenging Natural products Flavonoids trans-Tiliroside Igdoside A-D

### ABSTRACT

Elaeagnus angustifolia L. (Elaeagnaceae) is an important medicinal plant associated with numerous pharmacological activities. Its leaves are used as a therapeutic agent in traditional medicinal systems to treat diabetes. However, the active compounds responsible for the beneficial effects of E. angustifolia remain unclear. In this study, we determined the bioactive profile of E. angustifolia leaves using open column chromatography and semipreparative HPLC. Further, we sought to determine its *a*-glucosidase and *a*-amylase inhibitory activities, and its DPPH and ABTS radical-scavenging activities. Four undescribed flavonol glycosides, igdoside A-D, and four known glucosides were isolated from the ethyl acetate and n-butanol extracts of E. angustifolia leaves. Thereafter, the compound structures were identified using spectroscopic methods, including NMR and mass spectrometry. Of the compounds extracted, kaempferol-3-O-(6"-trans-p-coumaroyl)-β-D-glucopyranoside (trans-tiliroside), exhibited the highest a-glucosidase inhibitory activity with an IC<sub>50</sub> value of 2128  $\pm$  63  $\mu$ M compared to the positive control, acarbose (IC<sub>50</sub> = 6561  $\pm$  207  $\mu$ M). trans-Tiliroside was also found to exhibit potent scavenging activity against the ABTS radical, with an IC\_{50} value of 5  $\pm$  0  $\mu M,$  compared to the positive controls, trolox  $(31 \pm 1 \mu M)$  and  $\alpha$ -tocopherol (50  $\pm 1 \mu M$ ). In addition, isorhamnetin-3-O- $\beta$ -D-galactopyranoside  $(IC_{50} = 6 \pm 0 \mu M)$  and astragalin  $(IC_{50} = 6 \pm 0 \mu M)$  showed similar ABTS radical-scavenging activity as *trans*tiliroside. Based on HPLC, the content of trans-tiliroside was 9.69% in the ethyl acetate extract, 1.04% in decoction, 0.34% in 70% methanol extract, and 0.23% in infusion. None of the extracts and compounds showed  $\alpha$ -amylase inhibition or DPPH-scavenging activity.

#### 1. Introduction

The Elaeagnaceae family is represented by two floral species in Turkey. One of them, *Elaeagnus angustifolia*, is a thorny shrub that can reach a height of 7 m. *E. angustifolia* is deciduous; grows naturally in Europe, Turkey, Caucasus, Syria, Iran, Afghanistan, and Pakistan; and is widely cultivated throughout Turkey (Güner et al., 2012; McKean, 1965).

*E. angustifolia* has been identified as having numerous ethnobotanical uses by the local people of Turkey and is a potential herbal medicine for the treatment of diabetes mellitus (DM). The decoction of its leaves is consumed as a blood glucose-lowering agent (Arituluk and Ezer, 2012) while the infusions from its flowers and leaves are consumed as diuretics and antipyretics (Baytop, 1999). Previous studies have shown that *E. angustifolia* exerts antiarthritic, anti-inflammatory, antimicrobial, antimutagenic, antinociceptive, antioxidant, antitumor, cardioprotective, gastroprotective, hypolipidemic, insecticidal, tyrosinase inhibitory, and wound-healing effects (Carradori et al., 2020; Farzaei et al., 2015). The *in vitro*  $\alpha$ -glucosidase and  $\alpha$ -amylase inhibitory effects of the 70% methanol extract prepared from the leaves of *E. angustifolia* have been evaluated using different methods. However, to the best of our knowledge, further studies have not been conducted to determine the effective compounds in the extract (Saltan et al., 2017). By investigating the hypoglycemic effect of polysaccharides obtained from the fruit of this

https://doi.org/10.1016/j.phytochem.2021.112795

Received 25 November 2020; Received in revised form 18 April 2021; Accepted 23 April 2021 Available online 25 May 2021 0031-9422/© 2021 Elsevier Ltd. All rights reserved.





Abbreviations: ABTS, 2,2'-azinobis-3-ethylbenzothiazoline-6-sulfonic acid; DPPH, 1,1'-diphenyl-2,2'-picrylhydrazyl; HPLC, high performance liquid chromatography; NMR, nuclear magnetic resonance.

<sup>\*</sup> Corresponding author. Department of Pharmacognosy, Faculty of Pharmacy, Ataturk University, 25240, Erzurum, Turkey. *E-mail address:* hafize.yuca@atauni.edu.tr (H. Yuca).