

Antiproliferative effect of phenylethanoid glycosides from *Verbena officinalis* L. on colon cancer cell lines

LWT - Food Science and Technology (2015) 63, 1016-1022.

Manuel Alejandro Encalada^a, Sheyla Rehecho^a, Diana Ansorena^b, Iciar Astiasarán^b, Rita Yolanda Cavero^c, María Isabel Calvo^{a,*}.

^a Department of Pharmacy and Pharmaceutical Technology, Faculty of Pharmacy, University of Navarra, Irunlarrea s/n, 31008 Pamplona, Spain.

^b Department of Nutrition, Food Science, Physiology and Toxicology, Faculty of Pharmacy, University of Navarra, Irunlarrea s/n, 31008 Pamplona, Spain.

^c Department of Environmental Biology, Faculty of Sciences, University of Navarra, Irunlarrea s/n, 31008 Pamplona, Spain.

* Corresponding author: mcalvo@unav.es

Abstract

The cytotoxic effect of the aqueous extract from *Verbena officinalis*, was evaluated *in vitro* on DHD/K12/PROb rat colonic epithelial cell line and HCT-116 human colon adenocarcinoma cell line. In both cell lines, the IC₅₀ values were lower than 20 µg/mL after 72 h of treatment. Bioassay guided fractionation led to the isolation of 12 phenylethanoid glycosides with anti-proliferative activity, five of them are being reported for the first time. The new compounds were elucidated as 4''-acetyl-*O*-isoverbascoside, 2'',4''-diacetyl-*O*-verbascoside, 3'',4''-diacetyl-*O*-isoverbascoside, 4'',6''-diacetyl-*O*-betonyoside A and 3'',4''-diacetyl-*O*-betonyoside A. The IC₅₀ results suggest that antiproliferative activity is determined by not only the number of acetyl-groups but also their position in the aliphatic rings. Compounds exhibiting vicinal acetyl-groups in the sugar rings such as 3'',4''-diacetyl-*O*-isoverbascoside and 3'',4''-diacetyl-*O*-betonyoside A are particularly strong cytotoxic compounds against both cell lines. This investigation indicated that diacetyl-phenylethanoids might be valuable as cancer chemopreventive agents.