산딸나무 열매로부터 분리한 2차대사산물의 생리활성 검증

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Objectives

Cornus kousa Burg. (Cornaceae), a deciduous tree distributed in the mountains of Korea, China, and Japan, is in flower from May to June. Its fruits are red or pink, 2–3 cm in diameter and very delicious, and the seeds ripen from July to August. In Korea the fruits of this plant have traditionally been used as a hemostatic agent as well as for the treatment of diarrhea. Recently, the immuno-regulatory properties of *C. kousa* fruits extract have been reported. Our phytochemical study on the fruits of this plant led to isolation and structure determination of a new lignan glycoside together with several known compounds, such as six lignans, seven triterpenoids, four flavonoids, five sterols, and some lipids

Materials and Methods

 1 H-NMR (400 MHz), 13 C-NMR (100 MHz) and 2D-NMR spectra were recorded on a Varian Unity Inova AS-400 FT-NMR spectrometer. Methanol- d_4 with TMS as an internal standard was purchased from Sigma. RPMI Medium 1640, Dulbecco's Modified Eagle Medium and Penicillin-Streptomycin were purchased from GIBCO. FBS was from Hyclone. MTT and DMSO were purchased from Sigma. The methanol extract was fractionated into an EtOAc layer, an n-BuOH layer and a H₂O layer through solvent fractionation. The repeated SiO₂, ODS and Sephadex LH-20 c. c. of EtOAc fractions yielded twenty-two compounds (1-22).

Results

From the results of spectroscopic data including EIMS, FABMS, UV, IR, 1 H and 13 C-NMR, DEPT and 2D-NMR (COSY, HSQC, HMBC), the chemical structure for the new lignan glycoside was determined as (7'S, 8'R)-dihydrodehydrodiconiferyl alcohol-4'-O- β -D-xylopyranoside (1), namedcornuskoside A; for the lignans as (+)-pinoresinol (2), (-)-balanophonin (3), (+)-laricresinol (4), erythro/threo-guaiacylglycerol- β -coniferyl aldehyde ether

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(5), (7'S, 8'R)-dihydrodehydrodiconiferyl alcohol (6); for triterpenoids as ursolic acid (7), corosolic acid (8), taraxasterol (9), betulinic aldehyde (10), tormentic acid (11), ajunolic acid (12), nigaichgoside-F1 (13); for flavonoids as kaempferol (14), astragalin (15), hyperin (16), isoquercitrin (17); and for sterols as β -sitosterol (18), stigmasterol (19), 6a-hydroxystigmast-4-en-3-one (20), 6β -hydroxystigmast-4-en-3-one (21) and daucosterol (22). They were the first to be isolated from C. kousa. Their cytotoxic activities were assessed using MTT assay. Lots of lignans and triterpenoids showed the cytotoxicity against HCT-116, HeLa, MCF-7, SK-MEL-5, SK-OV-3 tumor cell lines. Also, compounds 7, 8, 10 and 11 revealed relatively high hACAT-linhibitory activity with the value of 46.2±1.1%, 46.7±0.9%, 41.5±1.3% and 60.8±1.1% at the concentration of 100 µg/mL, respectively. Our results suggest that the fruits of C. kousa might good source for anti-cancer, anti-hypercholesterolemia, andanti-atherosclerosis agents.

Figure. Chemical structure of isolated compounds from the fruits of Cornus Kousa.