

14th World Congress on

TOXICOLOGY AND PHARMACOLOGY

March 12-14, 2018 Singapore

Celastrol induces apoptosis-mediated cell death in multi-drug resistance human nasopharyngeal cancer cells**Ming Ju Hsieh and Mu-Kuan Chen**
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Nasopharyngeal carcinoma (NPC) belongs to squamous cell carcinoma that occurs in the epithelial lining of the nasopharynx. Because of the anatomical position close to the cervical lymph node, some patients have a distant metastasis at the time of diagnosis that leads to treatment failure. Although early stages have a high curability and excellent prognosis, advanced NPC urgently requires new drugs developed to reinforce the effectiveness of therapy without noticeable side effects. Celastrol, a chemical compound isolated from the root extracts of *Tripterygium wilfordii* (thunder god vine) and *Celastrus regelii* has been reported to possess anticancer potential. The aim of the present study was to determine the anticancer activity of celastrol and further elucidate the underlying molecular mechanisms. In this study, we first demonstrated that celastrol potently suppressed cell viability in MDR-NPC cell lines. Treatment of cells with celastrol induced G2/M arrest and apoptosis. Further studies showed that celastrol increased the expression of cleaved caspase-3, -8, -9 and subsequently activated apoptosis. Moreover, we found that celastrol-induced activation of bax, bim and t-Bid involved in the apoptosis. The expression of anti-apoptotic proteins Bcl-2 was significantly reduced, but expression of Bcl-XL was no significantly change after treatment of celastrol. Celastrol treatment also increased the expression of Fas, DcR2, DR5, RIP and TRADD. The cytotoxic effect of celastrol on NPC cells is mainly due to apoptosis, mediated by Fas-Fas ligand and mitochondrial pathway. These results suggested that celastrol could be a potential anticancer agent for NPC.

Recent Publications

1. Lin H F, Hsieh M J, Hsi Y T, Lo Y S, Chuang Y C, Chen M K, Chien S Y (2017) Celastrol-induced apoptosis in human nasopharyngeal carcinoma is associated with the activation of the death receptor and the mitochondrial pathway. *Oncol Lett*; 14(2): 1683-1690.
2. Chen J C, Hsieh M J, Chen C J, Lin J T, Lo Y S, Chuang Y C, Chien S Y, Chen M K (2016) Polyphyllin G induce apoptosis and autophagy in human nasopharyngeal cancer cells by modulation of AKT and mitogen-activated protein kinase pathways *in vitro* and *in vivo*. *Oncotarget*; 7(43): 70276-70289.

Biography

Ming Ju Hsieh is a Biochemistry Doctor, specializing in biochemistry, biotechnology, tumor metastasis, apoptosis and other research areas. He has considerable experience in research results in the hospital and research units.

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