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Topic of Research: Anti-viral Efficacy of Glycyrrhizin, Chrysin on the Life Cycle of Hepatitis B virus

Finding: Our study reveals that Glycyrrhizin and Chrysin have promising anti-viral efficacy against the Hepatitis B virus in cell line of hepatic origin. Further study needs evaluating their efficacy in other hepatic cell lines, non-hepatic cell lines and animal models of chronic HBV infection.

Background: Currently, **interferon (IFNs) and nucleoside analogues (NAs)** are the existing antivirals. Its long-term administration causes dose-dependent side effects, drug-resistance. Hence, it is of paramount importance to develop novel plant based anti-HBV candidates.

Methodology: HepG2 were treated with different concentrations of Glycyrrhizin and Chrysin for 72 h and non-toxic doses were determined by MTT assay. 1 μ g (pHBV 1.3X) wild type construct was transiently transfected in HepG2 cells. We performed ELISA by monitoring the concentrations of HBV surface antigen (HBsAg) and Hepatitis B e antigen (HBeAg). Extracellular HBV DNA and intracellular cccDNA were quantified by SYBR green real-time PCR assay. Molecular docking method was used where Glycyrrhizin, Chrysin and positive control lamivudine were docked with a HMGB1. The Absorption, Distribution, Metabolism, Excretion, and Toxicity (ADMET) properties were calculated by SwissADME and AdmetSAR web tool.

Results: Glycyrrhizin, Chrysin and lamivudine showed very good binding affinity and developed very stable complex with HMGB1 ($\Delta G = -7.0$ kcal/mol), ($\Delta G = -5.7$ kcal/mol) and ($\Delta G = -4.3$ kcal/mol) respectively. *In vitro* studies demonstrated that they decrease the expressions of HBsAg, HBeAg, supernatant HBV DNA, and cccDNA in a dose-dependent manner.

Conclusions: Glycyrrhizin and Chrysin have encouraging anti-HBV potential.

Keywords: Hepatitis B virus, Glycyrrhizin, Chrysin, HMGB1, cccDNA