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## DRUG DISCOVERY &amp; DESIGNING

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***Moringa oleifera* aqueous leaf extract induces cell cycle arrest and apoptosis in human liver hepatocellular carcinoma cells**Charlette Tiloke<sup>1,2</sup>, Alisa Phulukdaree<sup>3</sup>, Robert M Gengan<sup>2</sup> and Anil A Chuturgoon<sup>1</sup><sup>1</sup>University of KwaZulu-Natal, South Africa<sup>2</sup>Durban University of Technology, South Africa<sup>3</sup>University of Pretoria, South Africa

**Background & Aim:** Hepatocellular carcinoma is one of the leading global epidemics with highest incidence in developing countries such as South Africa. Traditional herbal medicines have been utilized for generations and a medicinal tree, *Moringa oleifera* (MO), have been part of a variety of treatments including cancer. We investigated the anti-proliferative and apoptosis inducing effects of MO crude aqueous leaf extract (MOE) in human liver hepatocellular carcinoma (HepG<sub>2</sub>) cells.

**Methods:** HepG<sub>2</sub> cell viability was evaluated using the MTT assay. Oxidative stress and DNA damage was determined using the TBARS and comet assays respectively. Apoptosis was assessed by caspase-9, -3/7 activities and ATP levels (luminometry). Cell cycle,  $\gamma$ H2AX and cleaved PARP-1 were determined by flow cytometry. Protein expression of c-myc, Bax, p-Bcl2, Smac/DIABLO, Hsp70, SRp30a and cleaved PARP-1 was assessed using western blotting.

**Results:** HepG<sub>2</sub> cells were exposed to various concentrations of MOE for 24 h and an IC<sub>50</sub> of 4.479 mg/ml was determined. MOE caused a significant increase in lipid peroxidation, DNA fragmentation and  $\gamma$ H2AX levels. A significant decrease in G<sub>1</sub>, S and G<sub>2</sub>-M phase was seen with a concomitant increase in apoptosis. SRp30a protein expression was significantly increased which led to the alternate splicing and subsequent activation of caspase-9. Caspase-9 and -3/7 was significantly increased with a significant decrease in ATP levels. Apoptosis was further confirmed with the significant decrease in c-myc, p-Bcl2 and Hsp70 protein expression and a significant increase in Bax, Smac/DIABLO and cleavage of PARP-1.

**Conclusion:** MOE induces cell cycle arrest, alternate splicing and apoptosis in cancerous HepG<sub>2</sub> cells.

**Biography**

Charlette Tiloke has completed her PhD at the Department of Medical Biochemistry at University of KwaZulu-Natal and is currently a Post-doctoral Research Fellow. Her research interests include anticancer and antimicrobial activity of medicinal plants and their synthesized nanoparticles.

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