

## 優秀論文 II

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題目：Inhibitory effect of *Nelumbo nucifera* leaf extract on 2-acetylaminofluorene-induced hepatocarcinogenesis through enhancing antioxidative potential and alleviating inflammation in rats

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摘要：本篇文章主要探討荷葉萃取物(NLE)預防肝癌發生的作用及其機制。首先我們發現以0.03% 2-acetylaminofluorene(AAF)餵飼老鼠經12週後可誘導老鼠產生肝纖維化，而同時餵飼NLE(0.5% - 2%)可抑制肝纖維化的發生。另外經六

個月的實驗後觀察到單獨餵飼AAF的老鼠其血液肝損傷指標如ALT, AST,  $\gamma$ -GT, AFP, TG, total cholesterol等及發炎指標IL-6, TNF- $\alpha$ 皆大大提升，而同時餵飼NLE的各組老鼠則明顯降低這些指標，而在肝臟組織生化及切片分析中也觀察到NLE可減少AAF所誘導的肝脂質過氧化、8-OHdG、GST-Pi及肝癌細胞。進一步研究發現NLE乃透過減緩肝發炎及增加抗氧化轉錄因子Nrf2及其下游酵素如catalase, SOD-1, GPx的表現來抑制AAF所誘發的肝癌。

**Abstract:** Leaf extract of *Nelumbo nucifera* (NLE) has been demonstrated to possess anti-atherosclerosis, improve alcohol-induced steatohepatitis, prevent high-fat diet-induced obesity, and inhibit the proliferation and metastasis of human breast cancer cells. This study determines the chemopreventive role of NLE against 2-acetylaminofluorene (AAF)-induced hepatocellular carcinoma (HCC) in rats. AAF was used to induce hepatocarcinogenesis in rats through genetic and nongenetic effects. After administration for 12 weeks, NLE (0.5% - 2%) supplementation orally inhibited AAF (0.03%)-induced hepatic fibrosis which appears during the development of premalignant lesions in rats. After the 6-month experiment, NLE supplementation resulted in decreasing AAF-induced serum parameters of hepatic injury, including the level of triglycerides, total cholesterol, alpha-fetoprotein (AFP), and inflammatory mediator IL-6 and TNF- $\alpha$  as well as the activities of alanine aminotransferase (ALT), aspartate aminotransferase (AST), and gamma-glutamyl transferase ( $\gamma$  GT). NLE supplementation also reduced AAF-induced lipid peroxidation and 8-hydroxy-2'-deoxyguanosine (8-OHdG) formation in the rat liver. Hepatic histopathological investigation revealed that NLE supplementation attenuated the AAF-induced HCC and GST-Pi expression. Furthermore, NLE supplementation increased the expression of transcription factor Nrf2 and its downstream targets, including catalase, glutathion peroxidase (GPx), and superoxide dismutase 1 (SOD-1) in the rat liver. Our findings indicate that NLE supplementation inhibited AAF-induced hepatocarcinogenesis by enhancing antioxidative potential and alleviating inflammation in rats.

