

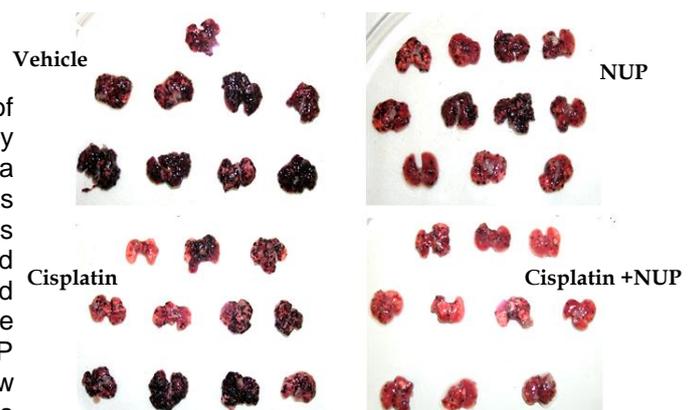
# Anti-Inflammatory and Anti-Cancer Effects of *Nuphar lutea* (yellow Water Lilly)



Natural products of *Nuphar lutea* (Nymphaeaceae) have been widely used for treating inflammatory conditions in ethnic medicine. Scientific investigations on the medicinal properties of Nuphar extracts have indicated several potential applications, such as anti-leishmanial, anti-bacterial and anti-cancer activities. Over the years we have screened a large number of Mediterranean plants used in ethnic pharmacopeia for biological activities including *Nuphar lutea* in order to identify the active compounds of these plants. As a result of our screening, we have identified Nuclear Factor  $\kappa$ B (NF $\kappa$ B) inhibitory activity in extracts of various plant parts of *Nuphar lutea*, in which the major components were sesquiterpene thioalkaloids. The NF $\kappa$ B family of transcription factors plays a pivotal role in inflammation and immune responses, proliferation, apoptosis and expression of certain viral genes. Therefore, the NF $\kappa$ B signaling pathway has also provided a focus for pharmacological intervention, primarily in chronic inflammation or in cancer, where the pathway is often constitutively active and plays a role in the disease. The therapeutic and preventive effects of many natural products may, at least in part, be due to their ability to inhibit NF $\kappa$ B. Translocation of NF $\kappa$ B to the nucleus stimulates the transcription of a wide variety of genes. NF $\kappa$ B induces interleukin-1 (IL-1)  $\alpha$  and  $\beta$ , Tumor Necrosis Factor  $\alpha$  (TNF- $\alpha$ ) and other molecules. Given the role of NF $\kappa$ B in cell proliferation and survival, it is not surprising that constitutive NF $\kappa$ B signaling has been implicated in oncogenesis and tumor progression.

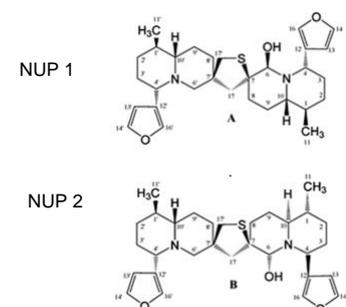
## The Technology

We have shown that that a partially purified mixture of thioalkaloids from of *Nuphar lutea* (NUP) as well two highly purified NUP fractions inhibited NF $\kappa$ B activation, leading to a powerful effect in inhibiting pro-inflammatory cytokines as well as promoting anti-inflammatory cytokines. These results were observed both in sera from mice injected with LPS and in-vitro, on peritoneal mouse macrophages also activated with LPS. Moreover, mice pretreated with NUP, were protected from lethality induced by LPS. In addition, NUP partially protected mice with peritonitis, providing a window for more efficient treatment with antibiotics during this condition. Animals treated with a combination of the cytotoxic anti-cancer drug cisplatin and NUP acted synergistically and more importantly, NUP showed a very significant reduction of B16 melanoma experimental lung metastasis.



## Patent Status

Granted



## Research Team

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