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A149 Extracts of rhizomes of Cyperus articulatus is an antagonist at the NMDA receptors and an agonist at the GABAB receptors expressed by Xenopus laevis oocytes

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Cyperus articulatus is a marshland plant commonly used in Africa and Latin America to treat many diseases in traditional medicine (1,2). Extracts of rhizomes of C. articulatus have been shown to possess pharmacological activities (3,4,5). In these studies, the effect of the decoction and the water extract of Cyperus articulatus rhizomes were tested in some receptors [NMDA (h NMDA R1A/2A), non-NMDA (r GluR3), GABA_B (R1b/R2) and potassium channels (Kir3)] expressed by Xenopus oocytes. Two electrodes (1-2 MΩ) voltage clamp experiments were performed with the membrane potential clamped to -70 mV. Water extract of the rhizomes of C. articulatus dosedependently decreased the ionic current induced by the EC80 of glutamate through NMDA receptors. Concentration 0.3 mg/ml of water extract inhibited 57% of this current. In contrast, the decoction of C. articu latus could not antagonize current through non-NMDA receptors. Even concentration of 0.3 mg/ml induced very little current (27% of total ionic current induced by the EC₁₀₀ of glutamate) through those receptors. These results are in accordance with the one obtain by Ngo Burn et al. (3) where the water extract of rhizomes of C. articula tus antagonized NMDA but not non-NMDA induced depolarisations in the rat cortical wedge preparation. In addition the water extract of rhizomes of C. articulatus dose-dependently induced an inward current in oocytes expressing heteromeric GABA_B (R1b/R2) together with heteromultimeric G protein activated inward rectifying potassium channels (Kir3). Concentration 3 mg/ml induced 57% of the current induced by a saturating concentration of GABA. In conclusion, rhizomes of C. articulatus could contain compounds that possess NMDA antagonising and GABAB agonizing properties. Further studies that are ongoing in our laboratory could allow the isolation of NMDA antagonist in C. articulatus.

References: 1. Burkill, H.M. (1985) The Usefull Plants of Tropical Africa. Royal Botanic Gardens Kew, London. 2. Schultes, R. E. and Raffauf, R. F. (1990) The healing forest: Medicinal plants of the Northwest Amazonia. Dioscorides Press, Portland. 3. Ngo Bum et al. (1996), J. Ethnopharmacol. 54: 103-111. 4. Rakotonirina et al. (2001), Fitoterapia 72: 22-29. 5. Ngo Bum et al. (2001), J. Ethnopharmacol. 76: 145-150.

A150 Coumarins having anti-amnestic activity from Angelica gigas

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Acetylcholinesterase (AChE) inhibitory activity—guided fractionation of Angelica gigas Nakai (Umbelliferae) led to the isolation of a new coumarin, peucedanone and eleven known coumarins. Among them, decursinol represented the highest inhibitory activity towards AChE in vitro. The study on their structure-activity relationships revealed that the cyclization of isoprenyl (IP) unit at C-6 of coumarin was essential for the AChE inhibitory activity (1). We evaluated the anti-amnestic activities of decursinol and its structural derivative, decursin, the major coumarin constituents of Angelica gigas in vivo using ICR mice with amnesia induced by scopolamine (1 mg/kg body weight, s.c.). Decursinol and decursin, when administered to mice at 1 and 5 mg/kg body weight i.p., significantly ameliorated scopolamine-induced amnesia as measured in the passive avoidance test (2). In contrast to in vitro study, decursin showed higher anti-amnestic activity than decursinol in vivo. Decursin (1 mg/kg body weight, i.p.) improved the spatial memory deficit of scopolamine-induced amnestic mice in the Morris water maze test (3). Moreover, decursin (1 mg/kg body weight, i.p.) significantly inhibited AChE activity by 34% (P < 0.05) of control in the hippocampus of treated mice. These results indicate that decursin may exert anti-amnestic activity in vivo through the inhibition of AChE activity in the hippocampus.

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References: 1. Kang et al. (2001) J. Nat. Prod. 64: 683-685. **2.** Christensen et al. (1992) Brain, 115: 1681-1699. **3.** Morris (1984) J. Neurosci. Methods. 11: 47-60.