

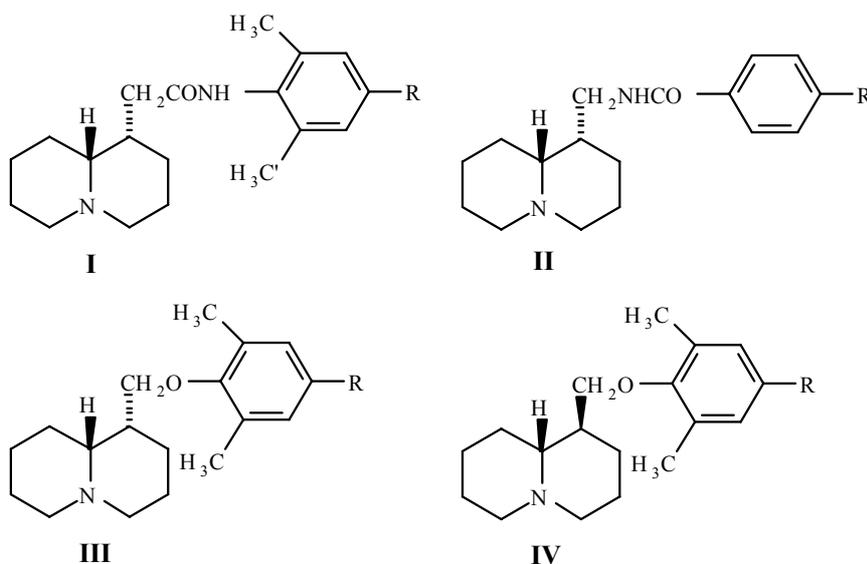
NOVEL QUINOLIZIDINYL DERIVATIVES AS ANTIARRHYTHMIC AGENTS

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On the pattern of well known N-(dialkylaminoalkyl)anilides, particularly of lidocaine and pilsicainide, a set of N-homolupinanoyl anilides was prepared in the past [1] and found endowed with strong antiarrhythmic activity *in vitro* and *in vivo* tests. These compounds exhibited an unusual pharmacological profile, being devoid of local anaesthetic activity, Ca²⁺ channel and β -adrenergic blocking activities. Since these peculiarities might be connected to the presence of the quinolizidine nucleus, new sets of compounds bearing this nucleus linked to different aromatic moieties (structures **I-IV**) have been prepared.



R= H; NO₂; NH₂; NHCOCH₃; NHSO₂CH₃; CO-Ar

Compounds of structure **III** and **IV** are somehow related to the 2,6-dimethyl-4-R-phenoxyalkylamines described by Mátyus et al [2], which exerted antiarrhythmic activity through the simultaneous block of Na⁺ and K⁺ channels.

The novel compounds are studied using isolated guinea pig dx atria spontaneously beating and Langendorff retrogradely perfused heart. Functional assays on adrenergic receptor subtypes are also performed.

[1] A. Sparatore, F. Sparatore. *Farmaco* 50 (1995) 153-166

[2] P. Mátyus, T. Rettegí, A. Varró, J. G. Papp. *Med. Res. Rev.* 20 (2000) 294-303