

SYNTHESIS AND PROPERTIES OF SOME NEW 4-ALKOXY PYRIDINE -3-SULFONAMIDES DERIVATED FROM CLASSICAL ORTHOPRAMIDES.

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Since the discovery of the first orthopramides, many compounds have been synthesized. Different structural modifications as the nature of the amide side-chain, the benzene substituents and the carboxamide group itself (non classical "biosteric" replacement by a sulfonamide group with, in some case, saving an antiemetic and psychotropic effect) have been performed.

On one hand, some sulfonamides are more potent sedative, on the other hand, the introduction of a sulfonamide group in a tropapride analogues decrease the activity. Moreover, the replacement of the principal ring by a heterocycle as pyridine, pyrimidine or pyrazin led to compounds which retain some interesting pharmacological properties.

In the present work, two structural modifications have been intended by the synthesis of pyridine sulfonamide compounds. Several new products have been investigated by radioligand binding test, intestinal prokinetic activity and by structural analysis (X-RAY crystallography and conformational analysis). All the compounds are very low toxic ($LD_{50} > 1000 \text{ mg/kg}$). Most of them are devoided of significant activity in the different tests.

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