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Preparation of Guanosines Modified by Carcinogenic Aromatic Amines

B. Robillard^a; M. F. Lhomme^a; J. Lhomme^a

^a Chimie organique Biologique, LA CNRS 351, Université de Lille 1, France

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PREPARATION OF GUANOSINES MODIFIED BY CARCINOGENIC AROMATIC AMINES

B. Robillard, M.F. Lhomme and J. Lhomme, Chimie organique Biologique, LA CNRS 351, Université de Lille 1, 59655 Villeneuve d'Ascq Cédex. France.

Summary: a general synthetic pathway has been developed to obtain aromatic amine-imidazole ring opened guanosine adducts.

Carcinogenic aromatic amines bind covalently to nucleic acids after metabolic activation into hydroxylamino derivatives. One of the major products obtained both <u>in vitro</u> and <u>in vivo</u> corresponds to an adduct in which the amino group of the carcinogen is bound to the C-8 position of guanine. More recently pyrimidine-type nucleosides <u>1</u> have been identified which could result from ring opening of the imidazole moiety in the latter adduct. This was observed notably in the case of 2-aminofluorene. In order to evaluate the physicochemical as well as the biological properties of this class of new compounds we have developed the general synthetic scheme as indicated:

This route was applied to the model aniline molecule and to the carcinogenic substance 2-aminofluorene.