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Nucleosides, Nucleotides and Nucleic Acids

Publication details, including instructions for authors and subscription information: http://www.informaworld.com/smpp/title~content=t713597286

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To cite this Article Pompon, A. , Gosselin, G. , Bergogne, M. C. and Imbach, J. L.(1987) 'Anomeric Mixtures of D-Xylo-and D-Lyxo-Furanonucleosides: Correlations between their Structures and their Retention Times in HPLC Analysis', Nucleosides, Nucleotides and Nucleic Acids, 6: 1, 465 - 466

To link to this Article: DOI: 10.1080/07328318708056257 URL: http://dx.doi.org/10.1080/07328318708056257

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ANOMERIC MIXTURES OF \underline{D} -XYLO-AND \underline{D} -LYXO-FURANONUCLEOSIDES : CORRELATIONS BETWEEN THEIR STRUCTURES AND THEIR RETENTION TIMES IN HPLC ANALYSIS

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<u>ABSTRACT</u>. Rationalization of the intrinsic structural factors which can influence retention times of anomeric <u>D</u>-xylo- and lyxofuranonucleoside analogues in reversed-phase high-performance liquid chromatography (HPLC) has been established.

We have previously reported the synthesis and biological evaluation of α and β -D-xylo- 1 and lyxofuranonucleosides 2 of the five naturally occurring nucleic acid bases as chemotherapeutically active agents. Here we present recent results 3 concerning the HPIC analysis of reconstituted mixtures of these nucleoside analogues.

From a detailed study of various parameters (size of support particules, nature and pH of the mobile phase, temperature) we have been able to establish optimized separation conditions, using a Waters Associanstrument.

Thus the resolution of the twenty derivatives in the D-xylo- and D-lyxose series is optimum on a Radial-Pak C_{18} -5 μ m column (100 x 8 mm ID, in RCM 100), at 17.5°C, with the following program: first isocratic eluent (solvent A: 0.1M sodium phosphate buffer, pH 4.73) during 12 mm.;

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then linear gradient from solvent A to 80 % solvent A - 20% solvent B (solvent B: mixture water-methanol, 1/1, v/v) during 38 mm.

We found that : i) for a same base and anomery, all lyxo-derivatives eluted before their xylo-isomers ; and in each xylo- and lyxoseries : ii) α -anomer is eluted before the β -anomer for a given base ; iii) the elution order was always C < U < G < T < A.

From this study, a rationalization in term of the contributions due to the natures of base, sugar and anomeric configuration, allowed a comparison between the retention times of all studied compounds.

Utilization of these results should permit the prediction of the best separation conditions for the two other series of <u>D</u>-pentofuranonucleosides (ribo- and arabino-), as well as for various series of nucleoside analogues (with deoxy and halogeno sugar moieties or modified bases).

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