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## Synthesis of Acyclic Nucleoside Using Potassium Iodide as Catalyst

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# SYNTHESIS OF ACYCLIC NUCLEOSIDE USING POTASSIUM IODIDE AS CATALYST

H.B. Lazrek \*\*, N. Redwane +, J.-L. Barascut \*\* and J.-L. Imbach \*\*

Abstract. An effective method for the synthesis of acyclonucleosides is reported. It is based on the use of potassium iodide as catalyst which enables mild condensation conditions.

Recent progress in the application of acyclonucleosides as antiviral agents creates the need for a simple and more general approach to their synthesis. We have recently reported (1) a rapid method based on the use of phase transfert catalysis. When we used alkylating agents having a base labile protecting group we observed the formation of a side product due to a partial deprotection (2). In order to overcome this problem we have developed milder conditions using potassium iodide as the catalyst (Scheme 1).

The condensation between various heterocycles (pyrimidines or purines) with acyclic chains (allyl bromide or (2-acetoxyethoxy)methyl bromide) has been made in the presence of potassium iodide. This selec-

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$$R_{2} \xrightarrow{N} R - X \xrightarrow{DMF/KI} \text{ isomer } N_{1} \xrightarrow{R_{2}} N_{1}$$

$$\frac{1}{2} R_1 = R_2 = OH$$
  
 $\frac{1}{2} R_1 = NH_2 R_2 = OH$ 

$$\frac{3}{4} R_1 = R_2 = OH$$
  
 $\frac{4}{4} R_1 = NH_2 R_2 = OH$ 

$$\begin{array}{c|c}
 & NH_2 \\
 & N \\
 & N$$

l R3

8

R = Allyl $R_3 = (2 - Acetoxy-ethoxy)methyl$ 

-Table 1-

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Base	Solvant	temp: •C	time hours	(a) yield(b) catalyseur	yield(b) %	isomers	(HO2M) AN
n	DMF	09	8	KI	36	ž	263 nm
C	DMF	90	2	ΚΙ	30	ź	272 nm
٨	DMF	80	2	X I	<sub>50(c)</sub>	N9+N3	(N3)=273nm (N9)=261nm
G	DMF	130	7	1	(P) <sup>07</sup>	N9+N7	260 nm 280 nm

<sup>(</sup>a) Imol base —— Imol KI.
(b) all yield are given after column chromatography.
(c) 40 % (Ng), 10% (Ng)
(d) 26 % (Ng), 14% (Ng)

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tively afforded N  $_{\rm l}$  isomers with pyrimidines but N  $_{\rm 9}$  , N  $_{\rm 3}$  or N  $_{\rm 9}$  , N  $_{\rm 7}$  isomer mixture with purines (Table !).

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