SOME CYCLIZATION REACTIONS IN THE NITROSATION OF UREA DERIVATIVES

Makoto Miyahara and Shozo Kamiya

National Institute of Hygienic Sciences

1-18-1 Kamiyoga Setagaya, Tokyo 158, Japan

In the course of the study to prepare new, antitumor nitrosourea derivatives, we found some cyclization reactions in the nitrosation of monosubstituted ureas. 1) Treatment of N-(2-hydroxyethyl)ureas (Ia-e) with sodium nitrite and dil. hydrochloric acid, gave 1-nitroso-2-oxazolidinone derivatives (IIa-e), which were easily denitrosated to give 2-oxazolidinone derivatives

2) Treatment of N-(2-pyridylmethyl)ureas (IVa,b) with nitrosyl chloride gave 1,2,4-oxadiazole derivatives (Va,b). However, ureas VIa,b, which are substituted at the methylene group of compound IVa,b, gave 2,5-diketopiperadine derivatives (VIIa,b).

3) N,N'-ethylenbis(ureas) (VIIIa,b) cyclized to give 1-carbamoyl-3-nitroso-2-imidazolidinone derivatives (IXa,b) under nitrosating conditions. Compounds IXa,b were denitrosated to give 1-carbamoyl-2-imidazolidinones (XIa,b) by treatment with acidic methanol.