

Figure 2. Effect of penicillic acid on liposomes.

penicillic acid, forming S-alkylated derivatives. Rapidly metabolising embryonic and developing systems are found to react with these adducts. In our experiment liposome carrying a net negative charge may act as a nucleophile; hence the affinity of penicillic acid will be greater towards the liposome carrying a net negative charge, resulting in the higher leakage of entrapped <sup>14</sup>C-glucose.

In other words membranes carrying a net negative charge may be more vulnerable to the action of penicillic acid.

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- Scott, P. M., Mycotoxic fungi, mycotoxins and mycotoxicoses,—An Encyclopedia Handbook, Vol. 1, (eds) D. Whyllie and L. G. Morehouse, Marcel Dekker, New York, 1978, p. 311.
- 2. Dickens, F. and Jones, H. E. H., Br. J. Cancer, 1965, 19, 392
- 3. Suzuki, S., Kimura, T., Saito, F. and Ando, K., Agric. Biol. Chem., 1971, 35, 287.
- 4. Kobayashi, H., Tsunoda, H. and Tatsuno, T., Chem. Pharm. Bull., 1971 19, 839.
- 5. Chan, P. K., Reddy, C. S. and Hayes, A. W., Toxicol. Appl. Pharmacol., 1980, 52, 1.
- 6. Bentley, R. and Keil, J. G., J. Biol. Chem.,

- 1962, 237, 867.
- 7. Gregoriadis, G., Neerunjun, E. D. and Hunt, R., Life Sci., 1977, 21, 357.
- 8. Imai, M., Inoue, K. and Nojima; S., Biochim. Biophys. Acta, 1975, 375, 137.
- 9. Imai, M. and Inoue, K., Biochim. Biophys. Acta, 1974, 352, 344.
- 10. Ganesan, M. G., Lakshmanan, M., Ravindran, K. V. and Damodaran, C., Z. Naturforsch., 1979, C34, 397.
- 11. Ciegler, A., Mintzlaff, H. F., Weisleder, D. and Leistner, L., Appl. Microbiol., 1972, 24, 114.

## A CONVENIENT LABORATORY SYNTHESIS OF METHYL ISOCYANATE

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METHYL isocyanate (MIC) is an important synthetic intermediate particularly in pesticide synthesis<sup>1</sup>. Amongst the several procedures<sup>2</sup> reported for the synthesis of aliphatic isocyanates, the Curtius rearrangement of acylazides<sup>3</sup> remains the method of choice due to the ready availability of acylazides. Acylazides are commonly prepared by reacting acylchloride with sodium azide. However, the insolubility of sodium azide in common organic solvents results in heterogeneous reaction conditions, leading to irreproducible results. The use of water-containing solvent mixture e.g. with aromatic acid chlorides is precluded due to the high sensitivity to hydrolysis of MIC. Several soluble azide equivalents for the preparation of aliphatic isocyanates have been described including tetramethylguanidinium azide<sup>4</sup>, trimethylsilyl azide<sup>5</sup>, tributylstannyl azide<sup>6</sup> and tetrabutylammonium azide<sup>7</sup>. Although these reagents are soluble in organic solvents, they have to be prepared separately prior to their reaction with acyl chlorides. A laboratory procedure is reported here for the preparation of MIC.

The reaction of acetyl chloride and sodium azide in refluxing toluene in the presence of tetrabutyl-ammonium bromide (TBAB) gave MIC in 70% yield. Comparable yield (73%) was obtained when reaction was conducted with 18-Crown-6 (18-C-6) in place of TBAB, while in their absence MIC resulted in only 20% yield, demonstrating definite advantage of phase transfer reagents in Curtius reaction. The

reaction appears to occur with exchange of TBAB with sodium azide yielding tetrabutylammonium azide in situ, which reacts with acetylchloride to form acetyl azide. As most aliphatic acylazides decompose to isocyanates<sup>8</sup> at 25-45°C, MIC results directly under the experimental conditions i.e. refluxing toluene. A similar phase-transfer mechanism may also operate in case of 18-C-6.

The method described here is a simple and convenient procedure for synthesis of monomeric MIC and the yields obtained are higher than the reported procedures? It may also be mentioned that although comparable yields of MIC are obtained using either TBAB or 18-C-6, the former has an advantage of low cost and is now being routinely used in our laboratory for synthesis of MIC. Furthermore, the method has also been adopted to prepare  $(1^{-14}C)$  methyl isocyanate using  $(1^{-14}C)$  acetyl chloride using microdistillation assembly for our studies on distribution and metabolism of MIC.

A 500 ml two-necked flask equipped with Leibig condensor, guard tube and dropping funnel was mounted on a magnetic stirrer. Freshly activated sodium azide<sup>3</sup> (13 g, 0.2 M) and tetrabutylammonium bromide (1.6 g,  $5 \times 10^{-3}$  M; or 18-C-6, 180 mg,  $7 \times 10^{-4}$  M) and dry toluene (100 ml) were charged into the flask. Acetyl chloride (7.8 g, 0.1 M) was added dropwise to the stirred mixture. After the addition was completed the mixture was refluxed for one hr after which the product was distilled through an efficient 25 cm spiral condensor. Yield 3.9 g (70%), b. p. 38°C (Lit<sup>7</sup> 38°C). The product was homogeneous on GLC (OV-17 column at 50°C, FID detector at 80°C), IR (neat) 2240, 2270 cm<sup>-1</sup> ( $\nu$  NCO) <sup>1</sup>H NMR (CDCl<sub>3</sub>)  $\delta$  3.03  $(CH_3\text{-NCO})$ , mass (E. I.) m/e 57 (M<sup>+</sup>).

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- 1. Kuhr, R. J. and Dorough, H. W., Carbamate insecticides: Chemistry, biochemistry and toxicology, CRC Press, Ohio, 1976, p. 15.
- 2. Richter, R. and Ulrich, H., The chemistry of cyanates and their thioderivatives, (ed.) S. Patai, John Wiley, New York, 1977, Vol. II. p. 619.
- 3. Smith, P. A. S., Organic reactions, (ed.) R. Adam, John Wiley, New York, 1947, Vol. III. p. 337.
- 4. Sakai, K. and Anselme, J. P., J. Org. Chem., 1971, 36, 2387.
- 5. MacMillan, J. H. and Washburne, S. S., J. Org. Chem., 1973, 38, 2982.
- 6. Kricheldorf, H. R. and Leppert, E., Synthesis,

- 1976, 329.
- 7. Brandstrom, A., Lamm, B. and Palmertz, I., Acta Chem. Scand., 1974, B28, 699.
- 8. Labbe, G., Chem. Rev., 1969, 69, 345.

SYMPLECTITIC INTERGROWTHS OF SAPPHIRINE IN CORDIERITE FROM BIOTITE-HYPERSTHENE-SILLIMANITE GNEISSES OF KASIPATNAM, VISAKHAPATNAM DISTRICT, ANDHRA PRADESH

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MIDDLEMISS<sup>1</sup>, Walker and Collins<sup>2</sup> have reported the sapphirine-bearing rocks from the Eastern Ghats of India. Since then, many investigators have found and described sapphirine-bearing rocks from the high grade granulite belt. Grew<sup>3</sup> reviewed the earlier works on sapphirine-bearing rocks of metasedimentary origin in the Eastern Ghats of India, and described the textures of some of these samples collected by him in the field, or received from others or from the Royal Ontario Museum, Canada. Geological work in Kasipatnam area in Visakhapatnam District (figure 1) revealed an occurrence of

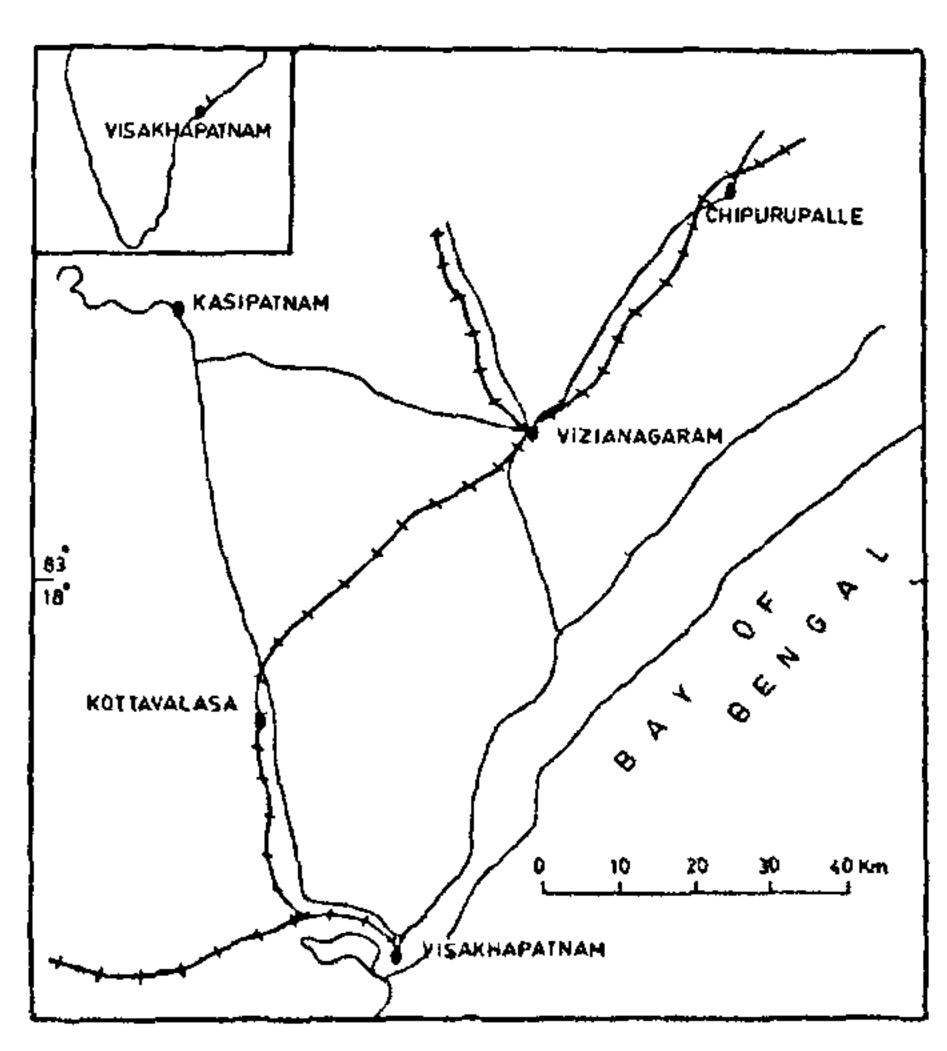


Figure 1. Location map of Kasipatnam area in Visakhapatnam District, Andhra Pradesh.