Recommendation from Scientific Expert Group

on Occupational Exposure Limits

for Diethyl ether

8 hour TWA : 100 ppm (308 mg/m³)

STEL (15 mins) : 200 ppm (616 mg/m³)

Additional classification :

Substance:

Diethyl ether CH₃CH₂-O-CH₂CH₃

Synonyms: 1,1'-oxybis(ethane), diethyl oxide

EINECS N° : 200-467-2

EEC N° : 603-022-00-4 Classification : F+; R12 R19

CAS N° : 60-29-7

MWt : 74.12

Conversion factor (20°C, 101kPa) : $3.08 \text{ mg/m}^3 = 1 \text{ ppm}$

Occurrence/use:

Diethyl ether is a colourless, very volatile, highly flammable liquid with a sweetish, slightly pungent, characteristic odour. It has a MPt of -116°C, a BPt of 34.5°C and a vapour pressure of 56.28 kPa at 20°C. It has a vapour density of 2.55 times that of air and is explosive over the range 2 - 48% in air. The odour threshold is approximately 9 ppm (28 mg/m³).

Diethyl ether has a production rate in the EEC in excess of 10,000 tonnes per annum. It is widely

used as a solvent and as a reaction medium in the laboratory and in industry.

Health Significance:

Diethyl ether has a low acute toxicity and induces anaesthesia at concentrations in excess of 15,000 ppm (46.2 g/m³). The critical effect is nasal irritation. In human volunteers the first indication of irritation occurred at a calculated level of 200 ppm (616 mg/m³) diethyl ether for 3-5 minutes, and this became objectionable at a calculated level of 300 ppm (924 mg/m³). Most individuals tested felt that a concentration of 100 ppm (308 mg/m³) would be acceptable over an 8 hour exposure period (Nelson *et al*, 1943).

No animal inhalation studies are available. A 90 day gavage study in rats established a NOAEL for systemic toxicity of 500 mg/kg body weight per day (American Biogenics Corporation, 1988). This may correspond to an inhalation exposure concentration of 1000 ppm (3080 mg/m³).

Diethyl ether shows no evidence of mutagenicity, either in vitro or in vivo (Simmon et al, 1987; Baden and Simmon, 1980; De Flora et al, 1984). No long term animal studies or reproduction studies are available.

Recommendation:

Although the Nelson study has major limitations, it provides the only available basis for setting exposure limits. A NOAEL of 100 ppm (308 mg/m³) for nasal irritancy in human volunteers was indicated by this study. A 90 day gavage study in rats was suggestive of a NOAEL of 1000 ppm (3080 mg/m³) for systemic effects. The volatility of diethylether makes the calculation of the effective dose administered in the experiment difficult. However it seems that an 8-hour TWA of 100 ppm (308 mg/m³), derived from Nelson's study would also provide sufficient protection against systemic effects. To prevent short term exposure to irritant levels a STEL (15 mins) of 200 ppm (616 mg/m³) is recommended.

At the levels recommended, no measurement difficulties are foreseen.



Bibliography:

Principle reference

SEG/CDO/5A (1991). CEC Criteria document for occupational exposure limit values. Diethyl ether. Prepared by Dept of Environmental Technology, Danish Technological Institute.

Key studies

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Baden, J.M. and Simmon V.F. (1980). Mutagenic activity of inhalational aesthetics. Mutat. Res. <u>75</u>, 169-189.

De Flora, S., Zanacchi, P., Camoirano, A., Bennicelli, C. and Badolati, G. S. (1984). Genotoxic activity and potency of 135 compounds in the Ames reversion test and in a bacterial DNA-repair test. Mutat. Res. 133, 161-198.

Nelson, K.W., Ege, J.F., Woodmann, L.E., Silverman, L. (1943). Sensory response to certain industrial solvent vapors. J. Ind. Hyg. Toxicol. <u>25</u>, 282-285.

Simmon, V.F., Kauhanen, K. and Tardiff, R.G. (1977). Mutagenic activity of chemicals identified in drinking water. Prog. Genet. Toxicol. 2, 249-258.