Recommendation from the Scientific Expert Group on Occupational Exposure Limits for heptane

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8 hour TWA : $500 \text{ ppm } (2085 \text{ mg/m}^3)$

STEL (15 mins) : -

Additional classification : -

<u>Substance:</u>

Heptane CH₃-CH₂-CH₂-CH₂-CH₂-CH₃ Synonyms : n-Heptane, dipropylmethane

EINECS N° : 205-563-8

EEC N°: 601-008-00-2 Classification: F; R11

CAS N° : 142-82-5 MWt : 100.20

Conversion factor (20°C, 101kPa) : $4.17 \text{ mg/m}^3 = 1 \text{ ppm}$

1 Occurrence/use

Pure heptane is a colourless volatile liquid with a faint hydrocarbon odour. It has a MPt of 90.7° C, a BPt of 98.4° C and a vapour pressure of 5.33 kPa at 22.3° C. The vapour density is 3.5 times that of air and it is explosive in the range 1.1 - 6.7% in air. The odour threshold is about 400 ppm (1668 mg/m 3).

Heptane occurs in natural gas and crude oil (0.1 - 1.9%) and in the volatile oils of many kinds of conifer. A content of 2.5% has been reported in petroleum, and this results in the major proportion of heptane found in the atmosphere. Technical heptane is a mixture of isomers and some forms contain n-hexane, but highly purified heptane may be produced. It is used as an industrial solvent (for adhesives, lacquers and inks in gravure printing), as an extraction solvent and in manufacture of plastic foams and synthesis of toluene and alkylbenzenes. This summary refers to pure n-heptane and not to commercial forms that may contain n-hexane.

2 Health Significance

Heptane is absorbed from the lungs, and is excreted as metabolites in the urine. The major metabolite is 2-heptanol, but 2,5-heptanedione, which may cause peripheral polyneuropathy, has also been found in small amounts in urine of exposed humans and rats (Perbellini *et al.*, 1986).

No data concerning irritating effects on skin and eyes are available but, like other hydrocarbons, heptane could be expected to act as a weak irritant.

Heptane depresses the central nervous system, and is narcotic in rats and mice at concentrations greater than 8000 ppm (33,360 mg/m 3). Takeuchi *et al.* (1980, 1981) exposed male rats to 2960 ppm (12,343 mg/m 3) heptane for 12h/d, 7d/week for 16 weeks. No electrophysiological changes were observed at any time during the experiment, and the authors concluded that heptane failed to produce polyneuropathy. Decreased auditory sensitivity was seen in rats exposed to n-heptane for 6 h/day for 28 days at 4000 ppm (16,680 mg/m 3), but not at 800 ppm (3336 mg/m 3) (Simonsen and Lund, 1995).

Heptane did not show mutagenic activity in a number of *in vitro* assays with bacteria, yeast and cultured mammalian cells (Brooks *et al.*, 1988).

No long term animal studies are available.

Few good human data are available. Patty and Yant (1929) reported a subjective effect (slight vertigo) at 1000 ppm (4170 mg/m³) in volunteers exposed for 6 minutes, but that the exposure level was calculated not measured. The peripheral neurotoxic potencies of hexane and heptane in man have been compared by measuring the rates of 2,5-dexaniedione and 2,5-deptanedione formation in human volunteers (Störmer et al, 1994). These data indicate that the peripheral neurotoxic potency of heptane is more than 40 times lower than than of hexane.

3 Recommendation

The 16 week rat study of Takeuchi et al. (1980, 1981), indicating a NOAEL of about 3000 ppm (about 12,500 mg/m 3) was considered to be the best available basis for setting exposure limits. In view of the lack of human data and of long term animal data, an uncertainty factor of 5 was considered to be appropriate. Taking into account the preferred value approach, the recommended 8-hour TWA is 500 ppm (2085 mg/m 3).

No STEL or "skin" notation was considered to be necessary.

At the levels recommended, no measurement difficulties are foreseen.

4 Key Bibliography

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