

ROBUST SUMMARY FOR HFC-23 100 APR -7 11 72 02

Summary

HFC-23 is used as a refrigerant, as an etchant of silicon dioxide dielectric layers, and as a clean agent fire extinguishant. HFC-23 is a clear, colorless gas with a slight ethereal odor. It has a boiling point of -82°C and a melting point of -155.1°C (DuPont, 1992; Lide, 2001). HFC-23 has a density of 1.44 g/cm³ at -82°C and a vapor density of 2.4 (air=1) (DuPont, 2002a; 2002b). HFC-23 has a vapor pressure of 45850 hPa at 25°C, an experimental log Kow of 0.64, and water solubility of 0.1 wt% at 25°C (DuPont, 1992; DuPont, 2002a; 2002b; Hansch et al., 1995; SRC, 2004). HFC-23 has no flash point. No further testing for physical-chemical properties is recommended.

HFC-32, difluoromethane, was selected as an analogous chemical for repeated dose and reproductive toxicity, based on its close physical-chemical properties and similar acute toxicity to HFC-23. The major properties of HFC-23 and HFC-32 are compared in Table 1.

Table 1: Comparison of Properties of HFC-23 and HFC-32

Chemical Name	Trifluoromethane	Difluoromethane	
Synonym:	HFC-23	HFC-32	
CAS#	75-46-7	75-10-5	
Molecular Formula	CHF ₃	CH ₂ F ₂	
Molecular Weight	70.01	52.02	
Melting Point (°C)	-155.1	-136	
Boiling Point (°C)	-82	-52	
Vapor Pressure (25°C)	45850 hPa	16800 hPa	
Heavier than Air?	Yes	Yes	
4-hour ALC in rats study (ppm)	> 663,000	~760,000	

If released to the atmosphere, HFC-23 is expected to exist solely as a vapor in the ambient atmosphere with an estimated half-life of about 180 years (Atkinson, 1989; SRC, n.d.). Some HFC-23 may gradually diffuse into the stratosphere with a diffusion half-life for transport from the troposphere to the stratosphere in the order of 20 years (Dilling, 1982). HFC-23 has a global warming potential (GWP) of 11,700 compared to the GWP of carbon dioxide and methane, which are 1 and 23, respectively. Hydrolysis is not expected to be an important process based on estimated half-lives of 5.1 years and

190 days at pH values of 7 and 8, respectively (Mill et al., 1987; SRC, n.d.). HFC-23 is expected to have high mobility in soil and volatilization of HFC-23 from moist soil surfaces is expected to be an important fate process. The potential for volatilization of HFC-23 from dry soil surfaces may exist (SRC, n.d.). HFC-23 is not expected to adsorb to suspended solids and sediment in water and is expected to volatilize rapidly from water surfaces. The estimated volatilization half-life from a model river and lake are approximately 2.5 hours and 3.3 days, respectively (Lyman et al., 1990; SRC, n.d.). HFC-23 is not expected to biodegrade rapidly. No degradation was detected in a 24-hour bottle assay (King, 1997; Streger et al., 1999). A BCF of 3.2 was calculated for HFC-23, suggesting that bioconcentration in aquatic organisms is low (SRC, n.d.). No further environmental fate testing is recommended.

No ecotoxicological studies have been conducted with HFC-23 and there are very little or no ecotoxicology data for similar non-chlorinated, fluorocarbon compounds. The related compounds HFC-134a (1,1,1,2-tetrafluoroethane) and HCFC-123 (2,2-dichloro-1,1,1-trifluoroethane) have been included here for comparison with HFC-23 as they are related compounds or provide conservative estimates of effects for HFC-23. ECOSAR was used to predict the aquatic toxicity of these fluorocarbons to green algae, daphnids and fish (Table 2). ECOSAR predictions are based on actual toxicity test data for classes of compounds with similar modes of action, i.e., narcosis in the case of fluorocarbons. Predicted log₁₀ Kow values were used as input for the ECOSAR model.

Table 2: Ac	quatic Toxicity	Values
1 17 **	A1 061 EC	D 1

Compound	log ₁₀ Kow**	Algae, 96-hr EC ₅₀	Daphnid, 48-hr EC ₅₀	Fish, 96-hr LC ₅₀
		mg/L	mg/L	mg/L
HFC-23	0.58	-	-	-
		(112)	(437)	(956)
HCFC-123	2.17	67.8*	17.3	55.5
		(25)	(48)	(83)
HFC-134a	1.68	-	980	450
		(33)	(51)	(150)

 $[\]ast$ Experimental endpoints are indicated in bold text, ECOSAR (ver 1.00a) estimated endpoints are in parentheses

Empirical test data are available for HFC-134a (fish, daphnia) and HCFC-123 (fish, daphnia, algae). Results of aquatic testing of HFC-134a with daphnids and fish indicated that the daphnid 48-hour EC₅₀ was 980 mg/L and the 96-hour fish LC₅₀ was 450 mg/L (Stewart and Thompson, 1991; Thompson, 1991) while the data for HCFC-123 indicate acute endpoint values of approximately 56, 17 and 68 mg/L for fish, daphnia and algae, respectively (Life Sciences Research 1992) ECOSAR generally over predicts the toxicity of HFC-134a and HCFC-123 and therefore, ECOSAR is also likely to over-estimate the toxicity for HFC-23. Based on the ECOSAR predictions, the actual toxicity test data for

^{**} Estimated values from KowWin

HFC-134a and HCFC-123 and the high Henry's Law Constant for these compounds, HFC-23 is considered a low hazard to aquatic organisms. No further ecotoxicological testing is recommended.

The 4-hour inhalation ALC of HFC-23 in male rats was >663,000 ppm, and the 4-hour ALC of HFC-32 (analogous chemical) was ~ 760,000 ppm (DuPont, 1975; 1980). In cardiac sensitization studies in dogs, HFC-23 was found to have no potential to cause cardiac sensitization in beagle dogs at concentrations up to 30% in air or 50% in air with auxiliary oxygen. There were no positive responses, no questionable positive responses, and no ventricular tachycardia ectopic bursts (DuPont, 1993). No further acute toxicity testing is recommended.

Neuro-behavioral and physiological effects of HFC-23 were examined in 6 male volunteers. Individuals were exposed to 10, 20, 40, or 60% of HFC-23 at escalating concentrations administered as 8 pulses of 3 minutes each, with 2-minute clearance periods between each pulse. The first subject exposed to HFC-23 completed the 8 pulses, but experienced an anesthetic effect and nausea at 60%. Although other physiologic parameters remained stable, the subject's response was considered intolerable. The second subject to inhale 40% HFC-23 experienced discomfort after 1 minute, and requested discontinuation of exposure. Both the 40 and 60% levels were then dropped from further evaluation. The remaining 4 subjects tolerated the 30% level of HFC-23. Therefore, 30% was considered to be the MTC (maximum tolerated dose). At 30% subjects reported anesthetic effects, but no effects were noted in blood pressure, heart rate or rhythm, oxygenation, respiratory rate, temperature, end tidal CO₂, or serum chemistries. However, when one subject received the 30% concentration during an NMR imaging study, an anesthetic effect with intolerable hyperacusis was demonstrated. In addition, exposure to 30% HFC-23 had systematic effects on neuropsychological functioning, mood, and subjective mental state (Fagan et al., 1995; Henry Ford, 1992; Rahill et al. 1998).

Although 90-day repeated dose inhalation studies with HFC-23 in rats and dogs reported LOAELs of > 10,000 and >5,000 ppm, respectively, there were insufficient details to assess the reliability of the studies (Leuschner et al., 1983). Therefore, data from an analogous chemical, HFC-32, are being presented to fulfill the repeated dose toxicity end point. A GLP 90-day inhalation toxicity study with HFC-32 was carried out in accordance with OECD Guideline No. 413. Groups of rats (20 per sex) were exposed whole body to HFC-32 for 6 hours per day, 5 days per week for 13 weeks. The treatment of rats with 4940, 14,600 and 49,100 ppm HFC-32 for 90 days resulted in a few minor and biologically insignificant changes (Ellis et al., 1996). Based on above data, no further repeated dose testing is recommended.

In a developmental toxicity study, pregnant rats were exposed by inhalation to 0, 5000, 20,000 or 50,000 ppm HFC-23 for 6 hours per day, on days 7-21 of gestation. The maternal and developmental no-observed-effect level (NOEL) was 50,000 ppm, which was the highest dose tested. No further developmental toxicity testing is recommended (DuPont, 1997).

Although no histological effects were reported in reproductive organs in 90-day inhalation studies in rats and dogs, there were insufficient details to assess the reliability of these studies (Leuschner et al., 1983). Therefore, data from an analogous chemical, HFC-32, are being presented to fulfill the reproductive toxicity end point. In the 90-day inhalation toxicity study with HFC-32 no effects on any reproductive organs examined or testis weights were observed (Ellis et al., 1996). Based on the available data, it is unlikely that HFC-23 will affect fertility. Therefore, no further reproductive toxicity testing is recommended.

HFC-23 was neither mutagenic in *Salmonella typhimurium* (Ames assay) nor in a gene mutation assay in Chinese hamster ovary (CHO) AS52/XPRT cells (gpt locus) (US Army, 1996a; 1996b). HFC-23 was mutagenic in an *in vitro* chromosome aberration assay in CHO-K1 cells at concentrations of 80% to 100% in the absence of metabolic activation. Damage, however, was probably due to a decreased oxygen level rather the activity of HFC-23, as under these same conditions 100% nitrogen also induced a significant increase in chromosomal damage to the same magnitude (US Army, 1996c). HFC-23 was positive in a *Drosophila* sex-linked recessive lethal (SLRL) test (Foltz and Fuerst, 1974). This study was included in the US EPA Report of the Gene-Tox Program, where HFC-23 could not be classified as positive or negative because of inadequate sample size. HFC-23 was negative in an *in vivo* mouse micronucleus assay (US Army, 1996d). No further genetic toxicity testing is recommended.

Human Exposure

HFC-23 is produced as a co-product in the reactors along with the production of HCFC-22 at the Louisville Works Plant site in Louisville, KY. The HFC-23 produced varies depending upon the age of the reactor catalyst, but is typically between 1% and 2% of the HCFC-22 produced. HCFC-22 is produced year round in the plant, so the HFC-23 is continuously produced along with the HCFC-22. The potential for exposure is greatest during loading of the material or during unscheduled maintenance. The primary concern for exposure is frostbite. The PPE for normal operations is Monogoggles and rubber gloves. If the exposure potential is likely due to line breaks containing residual material a Level "B" acid suit is worn (Greylite 20, a Poly Vinyl Chloride Material). For extreme potential exposure situations or emergency response a Greylite 20 Level "A" suit is worn.

In support of DuPont's commitment to reduce green house gas emission, the site has worked to minimize production of HFC-23 and improve HFC-23 capture technology. The Louisville plant has reduced emissions of HFC-23 about 70% from the 1990 base year. The HFC-23 is recovered into storage tanks and either sold into the fire extinguishant business or etchant gas business. More HFC-23 is produced than is sold, so the excess is shipped by railcar to Washington Works for destruction in their Thermal Converter. Losses of HFC-23 are minimized as much as possible by design. All of the HFC-23 processing equipment is located outdoors, so natural ventilation prevents accumulation of any high concentrations of HFC-23 where that material may leave the process (via small leaks or small vents necessary by design).

Because of its high vapor pressure, HFC-23 is handled as a liquefied gas during shipping and handling. Therefore, HFC-23 is handled in closed pressurized systems so that

exposure typically does not occur except for during maintenance or an abnormal situation, such as a leak.

HFC-23 is marketed as Freon[®] 23 and is a blend component in SUVA[®] 95, which are both refrigerants for very low temperature applications such as medical freezers and environmental chambers, Zyron[®] 23, an etchant gas for the semiconductor industry, and as FE-13, a clean agent fire extinguishant.

The DuPont Acceptable Exposure Limit for HFC-23 is 1000 ppm (8- and 12-hour TWA).

References:

Atkinson R (1989) Kinetics and mechanisms of the gas-phase reactions of the hydroxyl radical with organic compounds. <u>J. Phys. Chem. Ref. Data</u>. Monograph No.1, p 73 (cited in National Library of Medicine. Hazardous Substances Data Bank, HSDB/5207, accessed January 18, 2006).

Dilling WL (1982). In Conway RA, <u>Environmental Risk Analysis for Chemicals</u>, pp. 154-97 Van Nostrand Reinhold Co., New York, NY (cited in National Library of Medicine. Hazardous Substances Data Bank, HSDB/5207, accessed January 18, 2006).

DuPont Co. (1975). Unpublished Data, Report No. 698-75, Acute inhalation toxicity (November 18).

DuPont Co. (1980) Unpublished Data, Report No. 641-80, Inhalation approximate lethal concentration (ALC) (September 30).

DuPont Co. (1992). DuPont alternative fire extinguishants. FE-13 for total flooding agent applications. H-27337-2 (May).

DuPont Co. (1993) Unpublished Data, Huntington Research Center Report No. DPT273/921009, Halon 13B1, Freon 23, mixture of Freon 23 and HFC 125, assessment of cardiac sensitisation potential in dogs (March 15).

DuPont Co. (1997) Unpublished Data, Report No. 995-96, HFC-23: Inhalation developmental toxicity study in rats (February 27) (also in TSCATS Fiche OTS0573731).

DuPont Co. (2002) Material Safety Data Sheet 2025FR, Freon® 23 (November 4).

DuPont Co. (2002) Material Safety Data Sheet 6052FR, Suva® 23 (November 4).

Ellis MK, Trebilcock R, Naylor JL, Tseung K, Collins MA, Hext PM, and Green T (1996). The inhalation toxicology, genetic toxicology, and metabolism of difluoromethane in the rat. <u>Fundam. Appl. Toxicol.</u>, 31:243-251.

Fagan SC, Rahill AA, Balakrishnan G, Ewing JR, Branch CA, and Brown GG (1995). Neurobehavioral and physiologic effects of trifluoromethane in humans. <u>J. Toxicol.</u> Environ. Health, 45:221-229.

Foltz VC and Fuerst R (1974). Mutation studies with *Drosophila melanogaster* exposed to four fluorinated hydrocarbon gases. <u>Environ. Res.</u>, 7(3):275-285.

Hansch C, Leo A, and Hoekman D (1995). <u>Exploring QSAR</u>. <u>Hydrophobic, Electronic, and Steric Constants</u>. ACS Prof Ref Book. Heller SR (consult ed.), p 3. American Chemical Society, Washington, DC (cited in National Library of Medicine. Hazardous Substances Data Bank, HSDB/5207, accessed January 18, 2006).

Henry Ford Hospital, Detroit, MI (1992) Data cited in a letter from JR Ewing to CF Reinhardt (October 6) (also in TSCATS Fiche OTS0000874).

King GM (1997) Stability of trifluoromethane in forest soils and methanotrophic cultures. <u>FEMS Microbiol. Ecol.</u>, 22:103-109.

Leuschner F, Neumann B-W, and Hubscher (1983). Report on subacute toxicological studies with several fluorocarbons in rats and dogs by inhalation. <u>Arzneim.-Forsch.</u>, 33(10):1475-1476.

Lide DR (2001). <u>CRC Handbook of Chemistry and Physics</u>, 82nd ed., p. 3-208, CRC Press, Boca Raton, FL.

Lyman WJ, Reehl WF, and Rosenblatt DH (1990). <u>Handbook of Chemical Property Estimation Methods</u>, pp. 4-9, 5-4, 5-10, 15-1 to 15-29, American Chemical Society, Washington, DC (cited in National Library of Medicine. Hazardous Substances Data Bank, HSDB/5207, accessed January 18, 2006).

Mill T, Haag W, Penwall P, Pettit T, and Johnson H (1987). Environmental Fate and Exposure Studies Development of a PC-SAR for Hydrolysis: Esters, Alkyl Halides and Epoxides. EPA Contract No. 68-02-4254. Menlo Park, CA: SRI International (cited in National Library of Medicine. Hazardous Substances Data Bank, HSDB/5207, accessed January 18, 2006).

Rahill AA, Brown GG, Fagan SC, Ewing JR, Branch CA, and Balakrishnan G (1998). Neuropsychological dose effects of a Freon, trifluoromethane (FC-23), compared to N_2O . Neurotox. Teratol., 20(6):617-626.

SRC (n.d.) Syracuse Research Corporation (cited in National Library of Medicine. Hazardous Substances Data Bank, HSDB/5207, accessed January 18, 2006).

SRC (2004). Syracuse Research Corporation, information supplied to ChemIDplus, U.S. National Library of Medicine, Bethesda, MD, Specialized Information Services (SIS).

Stewart KM and Thompson RS (1991). ICI Group Environmental Laboratory Report No. BL3908/B, ICI, UK (cited in Berends AG, de Rooij CG, Shin-ya S, and Thompson RS (1999). Biodegradation and ecotoxicity of HFCs and HCFCs. <u>Arch. Environ. Contam.</u> Toxicol., 36(2):146-151).

Streger SH, Condee CW, Togna P, and Deflaun MF (1999) Degradation of hydrohalocarbons and brominated compounds by methane- and propane-oxidizing bacteria. <u>Environ. Sci. Technol.</u>, 33:4477-4482.

Thompson, R. S. (1991). ICI Group Environmental Laboratory Report No. BL4035/B, ICI, UK (cited in Berends AG, de Rooij CG, Shin-ya S, and Thompson RS (1999). Biodegradation and ecotoxicity of HFCs and HCFCs. <u>Arch. Environ. Contam. Toxicol.</u>, 36(2):146-151).

US Army (1996a). Unpublished Data, Integrated Laboratory Systems [ILS] Project No. A073-001, Salmonella typhimurium microsome reverse mutation assay (March 20).

US Army (1996b). Unpublished Data, Integrated Laboratory Systems [ILS] Project No. A073-003, AS52/GPT mammalian mutagenesis assay (May 10).

US Army (1996c). Unpublished Data, Integrated Laboratory Systems [ILS] Project No. A073-004, In vitro chromosome aberrations study in Chinese hamster ovary (CHO) cells (May 24).

US Army (1996d). Unpublished Data, Integrated Laboratory Systems [ILS] Project No. A073-002, Repeated inhalation exposure of FE-13 in mice, *Mus musculus* (bone marrow micronucleus assay) (January 19).