FOREWORD

INTRODUCTION

2-PROPANOL CAS N°: 67-63-0

SIDS INITIAL ASSESSMENT PROFILE

CAS NO.	67-63-0
CHEMICAL NAME	Isopropanol
STRUCTURAL FORMULA	CH3 - CH - CH3
	1/2
	ОН

RECOMMENDATION OF THE SPONSOR COUNTRY

[X]	currently of low priority for further work
[]	currently of low priority for further work, but avoid exposure to man and the environment
[]	requiring further information to assess identified concerns
[]	candidate for in-depth risk assessment with a view to possible risk reduction activities

SHORT SUMMARY OF THE REASONS WHICH SUPPORT THE RECOMMENDATION

Isopropanol is a high production volume chemical which is used as an industrial solvent, a component of industrial and consumer products and as a disinfectant. There is considerable potential for both occupational and consumer exposure.

There are estimates of significant fugitive emissions. Biodegradation in aquatic and terrestrial habitats, an physical degradation in the troposphere occur rapidly, indicating that isopropanol will not persist in the environment.

The mammalian/human toxicological properties of isopropanol have been well characterized in multiple animal species and humans for a variety of exposure routes, exposure durations and toxicity endpoints. High quality studies have been conducted that evaluate acute toxicity, skin and eye irritation, skin sensitization, subchronic and chronic toxicity, reproductive toxicity, developmental and developmental neurotoxicity, acute and subchronic neurotoxicity, genotoxicity and cancer. In addition, studies are available that characterize the disposition of isopropanol in mammals.

The information obtained from this database allows for the characterization of toxicity hazard of isopropanol for both human/mammalian and environmental effects. Taken together, these considerations support the conclusion that isopropanol is a low priority for further work.

FULL SIDS SUMMARY

CAS N	O: 67-63-0	SPECIES	PROTOCOL	RESULTS
DLIVCI	CALCUENICAL			
2.1	CAL-CHEMICAL Melting point		ASTM D97	90 °C
2.2	Boiling point		ASTM D1078	82 - 83 °C (at 1012 hPa)
2.3	Density		ASTM D4052	0.785 - 0.786 g/cm ³
2.4	Vapor pressure		not known	43 hPa at 20 °C
2.5	Partition coefficient (Log Pow)		not known	0.05 at 25 °C
2.6	Water solubility		not known	100 vol% at 20 °C (miscible)
	рН			at °C
	рКа			
2.7	Flash Point		IP 170	12°C (closed cup)
2.8	Auto Flammability		ASTM D2155	425°C
2.9	Flammability		84/449/EEC, A.10	Highly flammable
2.10	Explosive properties			Explosive
2.11	Oxidizing properties			No oxidizing properties
2.12	Adsorption coefficient (Log Koc)		Calculated	0.03
2.13	Henry's Law constant		calculated	7.52 x 10 ⁻⁶ atm • m³/mole 0.76 Pa • m³/mole
	RONMENTAL FATE PATHWAY			
3.1.1	Photodegradation			not subject to photolysis
	Atmospheric degradation (OH radical attack)		Various	in air $T1/2 = 18$ to 25 hours
3.1.2	Stability in water			not subject to hydrolysis
3.2	Monitoring data			In air = mg/ m ³ In surface water = ug/l In soil/sediment = ug/g
3.3				In biota = ug/g
-	Transport and Distribution		Calculated (Fugacity Level 1 type)	In air 22.3 % In water 77.7% In sediment 0.0 % In soil 0.0 %

			(Mackay)	In biota 0.0 %
3.5	Biodegradation	aerobic	(local exposure) APHA-219	49 % after 5 days at 20 °C
ECOT	OXICOLOGY			
4.1	Acute/Prolonged Toxicity to Fish	Pimephales promelas	Veith et al, 1983	LC50 (96 hr) = 9640 mg/l
4.2	Acute Toxicity to Aquatic Invertebrates	Daphnia Magna	Bringmann & Kuehn, 1977	EC50 (24 hr) > 10,000 mg/l
		Crangon Crangon		LC50 (48 hr) = 1400 mg/L
4.3	Toxicity to Aquatic Plants e.g. Algae	Scenedesmus Quadricuada	Bringmann & Kuehn, 1980	Toxicity Threshold = 1800 mg/l
4.4	Toxicity to Microorganisms e.g. Bacteria	Pseudomonas putida		Toxicity Threshold = 1050 mg/L
		Entosiphon Sulcatum		Toxicity Threshold = 4930 mg/L
		Microcystis Aeruginosa		Toxicity Threshold = 1000 mg/L
		Chilomonas paramecium (Protozoa)		Toxicity Threshold = 104 mg/L
4.5.2	Chronic Toxicity to Aquatic Invertebrates (Daphnia)			NOEC (16 day) = 141 mg/L (Growth) NOEC (21 day) = 30 mg/L
4.6.1	Toxicity to Soil Dwelling Organisms			LC50 (d) = mg/kg NOEL (d) = mg/l
4.6.2	Toxicity to Terrestrial Plants	Lactuca Sativa	Reynolds, 1977	EC50 (3 d) = 2100 mg/l (inhibition of seed germination)
4.6.3	Toxicity to Other Non- Mammalian Terrestrial Species (including Birds)	Drosophila	David & Bocquet, 1976	LC50 (48 hr) = 10200 - 13340 ppm
	Bioconcentration factor		calculated	1.0
TOXIC	COLOGY			
5.1.1	Acute Oral Toxicity	Rat	see Dossier	LD50 range = 4710 - 5840 mg/kg
		Mouse	Guseinov, 1985	LD50 = 4475 mg/kg
		Rabbit		LD50 = 5030 mg/kg

			Lehman, 1944	
		Dog	Lemman, 1944	LD50 = 4830 mg/kg
			Lehman, 1944	
5.1.2	Acute Inhalation	Rat	Cussinar	LC50 (4 hr) = 72.6 mg/l
	Toxicity	Rat	Guseinov, 1985	LC50 (8 hr) = 51 mg/l
		Mouse	Laham, 1979	LC50 (2 hr) = 53 mg/l
5.1.3	Acute Dermal Toxicity	Rabbit	Guseinov, 1985	LD50 = 12,870 mg/kg
5.2.1	Skin Irritation	Rabbit	Smyth, 1948	Not Irritating
5.2.2	Eye Irritation	Rabbit	·	Irritating
5.3	Sensitization	Guinea Pig	Nixon, 1975 Draize Test	Not Sensitizing
5.4	Repeated Dose	Rat and	Draize rest	NOEL = 500 ppm (13 weeks)
	Toxicity	Mouse, Inhl.	P&G unpubl.	LOEL = 1500 ppm
		Rat, Oral	EPA TSCA	NOEL=1% (870 mg/kg/day) (12weeks)
			Dil lo	LOEL = 2% (1280 mg/kg/day)
5.5	Genetic Toxicity in		Pilegaard & Ladefogel,	
	Vitro		1993	
	D	Salmonella		
	Bacterial Test	typhimurium		negative (with and without metabolic activation)
		СНО		
	Non-Bacterial Test	(HGPRT)	see Dossier	negative (with and without
		V79 (SCE)		metabolic activation)
			EPA TSCA	negative (with and without
		SA7/SHE		metabolic activation)
		SAT/SITE	see Dossier	negative (without metabolic
		Neurospora		activation)
		Crassa	see Dossier	negative (without metabolic
5.6		Mouse	300 2003101	activation)
	Genetic Toxicity in	(Micronucleus)	see Dossier	
5.7	vivo	Rat and		negative
		Mouse, Inhl.	EPA TSCA	
5.8	Carcinogenicity	Date 9 con		nogativo
J.0		Rats, 2-gen Oral	EPA	negative
	Toxicity to			NOTE DE LES TOPE AND A LES
	Reproduction		EPA TSCA	NOEL Parental = <500 mg/kg/day (BMDL10 =407 mg/kg/day)
				(Reproductive Effects)
				NOEL F1 Offspring = <500
5.9		Rat		mg/kg/day (BMDL5 = 449 mg/kg/day)
				NOEL F2 Offspring = <500

	Developmental Toxicity/Teratogenicit		EPA TSCA	mg/kg/day (BMDL5 = 418 mg/kg/day)
	у	Rabbit		NOEL = 400 mg/kg/day (Maternal)
				NOEL=400
				mg/kg/day(Developmental)
5.11				NOEL = 1200 mg/kg/day
			EPA TSCA	(Developmental neurotoxicity)
				NOEL = 240 mg/kg/day (Maternal)
	Experience with			NOEL=480
	Human			mg/kg/day(Developmental)
	Exposure			
				Accidental Exposure Information
				Voluntary Study Information
				Epidemiology Information

Table of Contents

Short Summary

Full SIDS Summary

1.0 Exposure Assessment

- **1.1 Production Volumes**
- 1.2 Major Uses
- 1.3 Manufacturing Process
- 1.4 Distribution
- 1.5 Fugitive Emissions
- 1.6 Workplace Monitoring

2.0 Summary of Environmental Effects

- 2.1 Environmental Fate
- 2.2 Toxicity to Aquatic Organisms
- 2.3 Toxicity to Plants

3.0 Summary of Health Effects

- 3.1 Acute Toxicity and Primary Irritancy
- 3.2 Effects Resulting from Repeated Exposure
- 3.3 Effects on Reproductive Capabilities
- 3.4 Effects on Developmental Toxicity
- 3.5 Genotoxic Effects
- 3.6 Carcinogenicity

4.0 Conclusions

5.0 References

1.0 Exposure Assessment

1.1 Production Volumes

Worldwide production of isopropanol in 1995 was reported to be 1.8 million metric tons, with the largest producing regions being North America (771 thousand tons), Western Europe (619 thousand tons), and Asia (280 thousand tons). (World Petrochemicals, SRI International, 1996). US production was 1.4 billion pounds in 1994. (Chem Eng. News, June, 1995).

1.2 Major Uses

Most isopropanol goes into the solvent market either directly or via conversion to acetone or one of acetone's derivatives -- methyl isobutyl ketone, methyl isobutyl carbinol, diacetone alcohol, or Isopropanol's major solvent isophorone. uses include inks, coatings, pharmaceuticals. Small percentages are used for esters and as rubbing alcohol. In Western Europe, some production of the acetone derivatives proceeds directly from isopropanol to the ultimate derivative, without separation of the acetone intermediate. IPA is used as a flavoring agent, foam inhibitor and a deicing agent for windshields in automobiles. IPA is used in the production of acetone, isopropyl acetate, isopropylamine, diisopropyl ether, isopropyl xanthate, herbicidal esters and aluminum isopropoxide. In contrast, Japan does not use isopropanol to produce acetone. The most important uses for isopropanol in Japan are as solvents in surface coatings and in inks. (World Petrochemicals, SRI International, 1996)

1.3 Manufacturing Process

Since the 1940s the dominant manufacturing process is the weak acid process in which propene gas is absorbed in, and reacted with, 60% sulfuric acid and the resulting sulfates hydrolyzed in a single step process. IPA is stripped and refined from the condensate which contains diisopropyl ether, acetone, and polymer oils of low molecular mass. Another major current manufacturing process is catalytic hydration of propene with water. Hydration can be gas-phase with a phosphoric acid catalyst, mixed phase with a cation-exchange resin catalyst or liquid phase using a tungsten catalyst.

1.4 Distribution

Tank cars and tank trucks are used for domestic clients. In plant distribution is via pipeline and tank truck.

1.5 Fugitive Emissions

It was estimated that 1.5% of the total US production in 1976 was lost to the environment (Dorigan et la., 1976). Emission registration data from the Netherlands from 1974-1979 indicated that 3.3% of the production volume was lost to the air and 0.2% was lost to water (IPCS, 2-Propanol, 1990). Approximately 0.6% of the 1985 worldwide production of IPA was estimated to be released to the environment. (DGEP, 1987).

1.6 Workplace Monitoring

Personnel exposures in isopropanol manufacturing facilities are very low because the process, storage and handling operations are enclosed. One Texas manufacturer reported personnel

exposures to be well below 10 ppm based on 1990-95 monitoring data. Workplace monitoring data, gathered from a Louisiana manufacturing facility between 1988-1994, showed most mean exposures to be below 5 ppm with peak part-time exposures below 20 ppm.

Less is known about customer workplace exposures, but they are expected to be higher due to evaporation of isopropanol in the various industrial and consumer product applications. OSHA and ACGIH have established workplace exposure limits for isopropanol of 400 ppm TWA-8hr and 500 ppm STEL.

Summary of Environmental Fate and Effects

2.1 Environmental Fate

2.0

Based on calculated results from a lever 1 fugacity model, isopropanol (IPA) is expected to partition primarily to the aquatic compartment (77.7%) with the remainder to the air (22.3%). IPA has been shown to biodegrade rapidly in aerobic, aqueous biodegradation tests and therefore, would not be expected to persist in aquatic habitats. IPA is also not expected to persist in surface soils due to rapid evaporation to the air. In the air, physical degradation will occur rapidly due to hydroxy radical (OH) attack. Overall, IPA presents a low potential hazard to aquatic or terrestrial biota.

IPA is expected to volatilize slowly from water based on a calculated Henry's Law constant of 7.52 x 10 -6 atm•m 3 /mole. The calculated half-life for the volatilization from surface water (1 meter depth) is predicted to range from 4 days (from a river) to 31 days (from a lake). Hydrolysis is not considered a significant degradation process for IPA. However, aerobic biodegradation of IPA has been shown to occur rapidly under non-acclimated conditions, based on a result of 49% biodegradation from a 5 day BOD test. Additional biodegradation data developed using standardized test methods show that IPA is readily biodegradable in both freshwater and saltwater media (72 to 78% biodegradation in 20 days).

IPA will evaporate quickly from soil due to it's high vapor pressure (43 hPa at 20° C), and is not expected to partition to the soil based on a calculated soil adsorption coefficient (log K_{∞}) of 0.03. IPA has the potential to leach through the soil due to it's low soil adsorption.

In the air, isopropanol is subject to oxidation predominantly by hydroxy radical attack. The room temperature rate constants determined by several investigators are in good agreement for the reaction of IPA with hydroxy radicals. The atmospheric half-life is expected to be 10 to 25 hours, based on measured degradation rates ranging from 5.1 to 7.1 x 10^{-12} cm³/molecule-sec, and an OH concentration of 1.5 x 10^6 molecule/cm³, which is a commonly used default value for calculating atmospheric half-lives. Using OH concentrations representative of polluted (3 x 10^6) and pristine (3 x 10^5) air, the atmospheric half-life of IPA would range from 9 to 126 hours, respectively. Direct photolysis is not expected to be an important transformation process for the degradation of IPA.

2.2 Toxicity to Aquatic Organisms

IPA has been shown to have a low order of acute aquatic toxicity. Results from 24- to 96-hour LC50 studies range from 1,400 to more than 10,000 mg/L for freshwater and saltwater fish and invertebrates. In addition, 16-hour to 8-day toxicity threshold levels (equivalent to 3% inhibition in

cell growth) ranging from 104 to 4,930 mg/L have been demonstrated for various microorganisms. Chronic aquatic toxicity has also been shown to be of low concern, based on 16- to 21-day NOEC values of 141 to 30 mg/L, respectively, for a freshwater invertebrate. Bioconcentration of IPA in aquatic organisms is not expected to occur based on a measured log octanol/water partition coefficient (log K_{ow}) of 0.05, a calculated bioconcentration factor of 1 for a freshwater fish, and the unlikelihood of constant, long-term exposures.

2.3 Toxicity to Plants

3.0

Toxicity of IPA to plants is expected to be low, based on a 7-day toxicity threshold value of 1,800 mg/L for a freshwater algae, and an EC50 value of 2,100 mg/L from a lettuce seed germination test.

Summary of Health Effects

3.1 Acute Toxicity and Primary Irritancy

Isopropanol has a low order of acute toxicity. It is irritating to the eyes, but not to the skin. Very high vapor concentrations are irritating to the eyes, nose, and throat, and prolonged exposure may produce central nervous system depression and narcosis. Human volunteers reported that exposure to 400 ppm isopropanol vapors for 3 to 5 min. caused mild irritation of the eyes, nose and throat. Although isopropanol produced little irritation when tested on the skin of human volunteers, there have been reports of isolated cases of dermal irritation and/or sensitization. The use of isopropanol as a sponge treatment for the control of fever has resulted in cases of intoxication, probably the result of both dermal absorption and inhalation. There have been a number of cases of poisoning reported due to the intentional ingestion of isopropanol, particularly among alcoholics or suicide victims. These ingestions typically result in a comatose condition. Pulmonary difficulty, nausea, vomiting, and headache accompanied by various degrees of central nervous system depression are typical. In the absence of shock, recovery usually occurred.

3.2 Effects Resulting from Repeated Exposure

The systemic (non-cancer) toxicity of repeated exposure to isopropanol has been evaluated in rats and mice by the inhalation and oral routes. The only adverse effects-in addition to clinical signs-identified from these studies were to the kidney. Rats exhibited an accumulation in hyaline (protein) droplets in kidney proximal tubule cells (males only, subchronic exposure) and an exacerbation of chronic progressive nephropathy, a spontaneous disease of unknown etiology common in aged rats (males and females, chronic exposure). In the mouse, minimal to mild effects to the kidney including renal tubular proteinosis and tubular dilation were observed following chronic exposure. The incidence of renal tubular proteinosis was generally significantly increased for all male and female treatment groups relative to controls; however, the majority of affected animals showed minimal degrees of tubular proteinosis (i.e., only a few tubules affected), there was no concentration-related gradient in either the frequency of severity of this change, and there was no corresponding evidence of alterations to the glomeruli. Mild to moderate degrees of tubular dilation were observed in a small number of females in the 2500 and 5000 ppm groups (significantly increased only for the 5000 ppm group). This finding, however, was not duplicated in male mice (a

significant increase was only seen for the 500 ppm group) nor was it accompanied by evidence of tubular cell degeneration or urinary outflow obstruction.

3.3 Effects on Reproductive Capabilities

A recent two-generation reproductive study characterized the reproductive hazard for isopropanol associated with oral gavage exposure. This study found that the only reproductive parameter apparently affected by isopropanol exposure was a statistically significant decrease in male mating index of the F1 males. It is possible that the change in this reproductive parameter was treatmentrelated and significant, although the mechanism of this effect could not be discerned from the results of the study. However, the lack of a significant effect of the female mating index in either generation, the absence of any adverse effect on litter size, and the lack of histopathological findings of the testes of the high-dose males suggest that the observed reduction in male mating index may not be biologically meaningful. Additional support for this conclusion is provided by the fact that most of the females became pregnant. Furthermore, male and female fertility, and female fecundity indices of rats dosed with isopropanol were not different from those of controls by statistical analysis and were within, or relatively close to, historical control values. No reproductive effects were noted in other studies in which rats were dosed up to 2% in the drinking water. Exposure to 1000 mg/kg/day and to a lesser extent 500 mg/kg/day did result in a reduction in postnatal survival in both F1 and F2 litters. Derivation of an appropriate NOAEL for offspring effects was made difficult because of conflicting interpretations of the reductions in postnatal survival for the 500 mg/kg/day treatment group. The U.S. EPA (1992) and Tyl (1996) concluded the reductions were treatment- and dose-related, a conservative interpretation that supports a NOAEL of 100 mg/kg/day. Alternatively, Bevan et al. (1995) and Harris (1995) deemed the observations not to be biologically significant and concluded the NOAEL to be 500 mg/kg/day. In order to clarify this issue a benchmark dose (BMD) assessment was conducted for the study's developmental and reproductive findings (Shipp et al., 1996). For the offspring developmental effects, BMD dosages (BMDLs) of 449 and 418 mg/kg/day were estimated for the F1 and F2 generations, respectively. Based upon the decrease in male mating index observations in the P2 males, a BMDL10 of 407 mg/kg/day was estimated for reproductive effects.

3.4 Effects on Developmental Toxicity

The developmental toxicity of isopropanol has been characterized in rat and rabbit developmental toxicity studies and in a rat developmental neurotoxicity study. The rats were dosed by oral gavage at 400, 800 or 1200 mg/kg from gestational days 6 through 15. The rabbits were dosed by oral gavage at 120, 240 or 480 mg/kg from gestational days 6 through 18. These studies indicate that isopropanol is not a selective developmental hazard. Isopropanol produced developmental toxicity in rats, but not in rabbits. In the rat, the developmental toxicity occurred only at maternally toxic doses and consisted of decreased fetal body weights, but no teratogenicity. These data suggest the developmental NOAEL is 400 mg/kg/day for rats and 480 mg/kg/day for rabbits.

Isopropanol has also been tested for developmental toxicity in rats via oral gavage. The rats were dosed at 200, 700 and 1200 mg/kg from gestational days 6 through 21. No exposure-related effects were noted on motor activity, weights of the four regions of the brain, developmental landmarks, or morphological changes to the tissues of the central nervous tissue. These data suggest the developmental neurotoxicity NOAEL for rats is 1200 mg/kg.

3.5 Genotoxic Effects

All genotoxicity assays reported for isopropanol have been negative. Characterization of the genotoxicity hazard for isopropanol is provided by both in vitro and in vivo mutation/chromosomal studies. Isopropanol was found to be negative in an in vitro CHO/HGPRT assay, was negative in vitro for aneuploidy in *Neurospora crassa*, and did not increase micronuclei in an in vivo micronuclei assay in mice. Mutagenicity studies also showed that isopropanol was not mutagenic in various Ames assays both in the presence or absence of an S9 metabolic activation system. In vitro sister chromatic exchange (SCE) assays on isopropanol using cultured V79 cells both with and without S9 activation, were also negative. Isopropanol did not induce transformation in Syrian hamster embryos infected with Simian SA7 virus. These studies demonstrate that isopropanol is not a hazard for genotoxic effects.

3.6 Carcinogenicity

Two recent chronic exposure, rodent inhalation studies were conduct to evaluate isopropanol for cancer potential. One study was performed exposing Fischer 344 rats to 500, 2500 and 5000 ppm of IPA for 6 hours/day, 5 days/week for 24 months. The only tumor rate increase seen was for interstitial (Leydig) cell tumors in the male rats. Interstitial cell tumors of the testis is typically the most frequently observed spontaneous tumor in aged male Fischer 344 rats (Haseman et al., 1990). Nearly all male Fischer rats will develop these proliferative tumors if they are allowed to complete their lifespan (Boorman et al., 1990). A mouse inhalation study was performed exposing CD-1 mice to 500, 2500 and 5000 ppm of IPA for 6 hours/day, 5 days/week for 18 months. There was no increased frequency of neoplastic lesions in any of the treated groups. These studies demonstrate that isopropanol does not exhibit carcinogenic potential relevant to humans. Furthermore, there was no evidence from this study to indicate the development of carcinomas of the testes in the male rat, nor has isopropanol been found to be genotoxic. Thus, the testicular tumors seen in the isopropanol-exposed male rats are considered of no significance in terms of human cancer risk assessment.

4.0 Conclusions

Isopropanol (IPA) is a high production volume chemical which has wide use as an industrial solvent and as a component in numerous industrial and consumer products. It has a potential for widespread exposure to both workers and consumers.

Based upon physical and chemical properties, isopropanol is not expected to persist in the environment. Aerobic biodegradation of isopropanol occurs rapidly. IPA is not expected to persist in soil due to low soil adsorption and rapid evaporation to air. In the air, isopropanol is subject to rapid oxidation by hydroxyl radical attack. IPA has a low order of toxicity to aquatic organisms and plants, and bioconcentration in aquatic organisms is not expected to occur.

The mammalian/human toxicological properties of IPA have been well characterized in multiple animal species and humans for a variety of exposure routes, exposure durations and toxicity endpoints. High quality studies have been conducted that evaluate acute toxicity, skin and eye irritation, skin sensitization, subchronic and chronic toxicity, reproductive toxicity, developmental and developmental neurotoxicity, acute and subchronic neurotoxicity, genotoxicity and cancer. In addition, studies are available that characterize the disposition of IPA in mammals.

The information obtained from this database allows for the characterization of toxicity hazard of IPA for both human/mammalian and environmental effects. Taken together, these considerations support the conclusion that IPA is a low priority for further work.

5.0 References

Bevan C., Tyler T.R., Gardiner T.H., Kapp Jr. R.W., Andrews L. and Beyer B.K. (1995). Two-generation reproductive toxicity study with isopropanol in rats. *J. Appl. Toxicol.* 15(2):117-123.

Boorman, G.A., Chapin, R.E., and Mitsumori, K. (1990) Chapter 24. Testis and Epididymis. In: Pathology of the Fischer Rat, (Boorman, R.A., Eustis, S.L., Elwell, M.R., Montgomery, C.A., Jr., and Mackenzie, W.F., eds.), pp. 405-418, Academic Press, CA

Chemical and Engineering News, June 26, 1995, p.40.

DGEP - Review of literature data on 2-propanol, Leidschendam, Netherlands, Directorate General of Environmental Protection, Ministry of Housing, Physical Planning and Environment. (1987)

Dorigan, J., Fuller, B. and Duffy, R. Scoring of organic air pollutants. Chemistry production and toxicity of selected synthetic organic chemicals. The MITRE Corporation -- MITRE Technical Report MTR-7248, Rev. 1, Appendix III (1976)

Harris S.B. (1995). A review of the EPA comments regarding the study entitled "Multi generation rat reproduction study with isopropanol". Report prepared for the Chemical Manufacturers Association Isopropanol Panel.

Haseman, J.K., and Arnold, J. (1990) Chapter 35. Tumor Incidences in Fischer 344 Rats: NTP Historical Data. In: Pathology of the Fischer Rat., (Boorman, G.A., Eustis, S.L., Elwell, M.R., Montgomery, C.A., Jr., and MacKenzie, W.F., eds.), pp. 555-564, Academic Press, CA

IPCS - International Programme on Chemical Safety - Environmental Health Criteria 103 - 2-Propanol, World Health Organization (1990).

Shipp A.M., Allen B.C., Van Landingham C., Gentry P.R. and Crump K.S. (1996). Quantitative dose-response analysis - Application of the benchmark method to the multi-generation rat reproduction study for isopropanol. Report prepared for the Chemical Manufacturers Association Isopropanol Panel.

Tyl R.W. (1996). February 12, 1996 Letter to the Chemical Manufacturers Association Isopropanol Panel.

US EPA, Draft Final RM1 Risk Assessment of Isopropanol, OPPT,1996. TSCA Public Docket Number AR-141.

SIDS DOSSIER ON THE HPV CHEMICAL ISOPROPANOL

CAS. No. 67-63-0

DATE OF LAST UPDATE: March 1997

CONTENTS

1.	GENERAL INFORMATION
1.01	SUBSTANCE IDENTIFICATION
1.02	OECD INFORMATION
1.1	GENERAL SUBSTANCE INFORMATION
1.2	SYNONYMS
1.3	IMPURITIES
1.4	ADDITIVES
1.5	QUANTITY
1.7	USE PATTERNS
1.8	OCCUPATIONAL EXPOSURE LIMIT VALUES
1.9	SOURCES OF EXPOSURE
2.	PHYSICO-CHEMICAL DATA
2.1	MELTING POINT
2.2	BOILING POINT
2.3	DENSITY
2.4	VAPOR PRESSURE
2.5	PARTITION COEFFICIENT
2.6	WATER SOLUBILITY
2.7	FLASH POINT
2.8	AUTO FLAMMABILITY
2.9	FLAMMABILITY
	EXPLOSIVE PROPERTIES
2.11	OXIDIZING PROPERTIES
3.	ENVIRONMENTAL FATE AND PATHWAYS
3.1.1	PHOTODEGRADATION
3.1.2	STABILITY IN WATER
3.3.1	TRANSPORT BETWEEN ENVIRON. COMPART.
3.3.2	DISTRIBUTION
3.4	MODE OF DEGRADATION IN ACTUAL USE
3.5	BIODEGRADATION
4.	ECOTOXICITY
4.1	ACUTE/PROLONGED TOXICITY TO FISH
4.2	ACUTE TOX. TO AQUATIC INVERTEBRATES
4.3	TOXICITY TO AQUATIC PLANTS E.G. ALGAE
4.4	TOX. TO MICROORGANISMS E.G. BACTERIA
4.5.2	CHRONIC TOXICITY TO AQUATIC INVERTEBRATES
4.6.2	TOXICITY TO TERRESTRIAL PLANTS
4.6.3	TOX. TO OTHER NON-MAMM. TERR. SPECIES

5. TOXICITY

- 5.1.2 ACUTE INHALATION TOXICITY
- 5.1.3 ACUTE DERMAL TOXICITY
- 5.1.4 ACUTE TOXICITY, OTHER ROUTES
- 5.2.1 SKIN IRRITATION
- 5.2.2 EYE IRRITATION
- 5.3 SENSITIZATION
- 5.4 REPEATED DOSE TOXICITY
- 5.5 GENETIC TOXICITY IN VITRO
- 5.6 GENETIC TOXICITY IN VIVO
- 5.7 CARCINOGENICITY
- 5.8 TOXICITY TO REPRODUCTION
- 5.9 DEVELOPMENTAL TOXICITY/TERATOGENICITY
- 5.10 OTHER RELEVANT INFORMATION
- 5.11 EXPERIENCE WITH HUMAN EXPOSURE

6. REFERENCES

1. GENERAL INFORMATION

1.01 Substance Identification

CAS-No.: 67-63-0
EINECS-No.: 200-661-7
IUPAC-Name: Propan -2- ol
Molecular Formula: C3H8O
Molecular Weight: 60

Smiles Code: CC(OH)C

1.02 OECD Information

Sponsor Country: United States

Lead Organization: Environmental Protection Agency

Contact Person: Mr. Charles Auer

Name of Responder: Chemical Manufacturers Association

Name: Ms. Marion K. Stanley Address: 1300 Wilson Boulevard

Arlington, VA 22209

1.1 General Substance Information

Substance Type Physical Status Purity

organic liquid >99.7 % w/w

1.2 Synonyms

IPA

Isopropyl Alcohol Isopropanol 2-propanol

1.3 Impurities

<u>CAS-No.</u> <u>EINECS-No.</u> <u>IUPAC-Name</u> <u>Value</u>

water < 0.15 % w/w

1.4 Additives

<u>CAS-No.</u> <u>EINECS-No.</u> <u>IUPAC-Name</u> <u>Value</u>

none

1.5 Quantity

Production: 1.8 million metric tons: North America: 771 metric tons, Western

Europe: 619 metric tons, Asia: 280 metric tons

Reference: World Petrochemicals, 1996

1.7 Use Pattern

<u>Category</u> <u>Type of Use</u>

Industrial

Sectors/General Use adhesive

automotive construction cosmetics

electrical products leather industry metalurgical mining

ceramics/glass

industrial organic chemicals

paints photography pigments and dyes

plating and surface finishing printing and publishing petroleum refining soap production

textiles

water and waste treatment

Medical disinfectant

1.8 Occupational Exposure Limit Values

Type of Limit TLV (US) Value 983 mg/m³

Short Term Exposure Limit

Value 1230 mg/m^3

Time Schedule

Frequency 15 minute

Reference 1995-1996 Threshold Limit Values (TLVs) for Chemical Substances

and Physical Agents and Biological Exposure Indices (BEIs). American Conference of Governmental Industrial Hygienists

(ACGIH), Cinncinati, OH (1995).

1.9 Sources of Exposure

Remark Primary production process is the weak acid process in which propene

gas is absorbed in, and reacted with, 60% sulfuric acid and the resulting sulfates hydrolyzed in a single step process. IPA is stripped and refined from the condensate which contains diisopropyl ether,

acetone and polymer oils of low molecular mass.

Reference US NIOSH Criteria for a recommended standard: occupational

exposure to isopropyl alcohol, Cincinnati, OH. US NIOSH, US Dept of Health Education and Welfare, Public Health Service, Center for

Disease Control (DHEW Publication No. NIOSH 76-142.

Remark Emission registration data from the Netherlands from 1974-1979

indicated that 3.3% of the production volume was lost to the air and

0.2% was lost to water

Reference IPCS, 2-Propanol WHO (1990)

Remark Approximately 0.6% of the 1985 worldwide production of IPA was

estimated to be released to the environment.

Reference DGEP (1987).

2 PHYSICO-CHEMICAL DATA

2.1 Melting Point

Value = -90 degree C

Decomposition Sublimation

Method other

Year

GLP no

Test condition Method is ASTM D 97

Reference Shell Chemicals data sheet IS 3.2.4 dated June 1992

2.2 Boiling Point

Value ca. 82 - 83 degree C

Pressure 1012 hPa

Decomposition no

Method other

Year

GLP no

Test condition ASTM D1078, standard method for distillation range Reference Shell Chemicals data sheet IS 3.2.4 dated June 1992

2.3 Density

Type density

Value ca. 0.785 - 0.786 g/cm3

Temperature 20 degree C

Method other

Year

GLP no

Test condition ASTM D4052, Standard method for density

Reference Shell Chemicals data sheet IS 3.2.4 dated June 1992

2.4 Vapor Pressure

Value = 43 hPa Temperature 20 degree C

Method other

Year GLP

Reference Shell Chemicals data sheet IS 3.2.4 dated June 1992

2.5 Partition Coefficient

 $\begin{array}{ll} log \ Pow & = 0.05 \\ Temperature & 25 \ degree \ C \\ Method & other \ (measured) \end{array}$

Year 1973 GLP no data

Reference Dillingham, E.O. et al., J.Pharm.Sci., 62, 1973, 22. In Pomona Data

File on Log P and related parameters.

2.6 Water Solubility

Value = 100 vol% at 20 degree C pH Concentration at degree C

pKa at 25 degree C Descr. miscible Method other

Year GLP

Reference Shell Chemicals Data sheet IS 3.2.4 dated June 1992

2.7 Flash Point

Value = 12 degrees C
Type closed cup
Method other

Year

GLP no

Test condition IP 170, Standard method for flash point

Reference Shell Chemicals data sheet IS 3.2.4 dated June 1992

2.8 Auto Flammability

Value = 425 degree C

Pressure

Method other

Year

GLP no

Test condition ASTM D2155, Standard method for autoignition Reference Shell Chemicals data sheet IS 3.2.4 dated June 1992

2.9 Flammability

Result highly flammable

Method Directive 84/449/EEC, A.10

Year

GLP no

Reference Shell Chemicals data sheet IS 3.2.4 dated June 1992

2.10 Explosive Properties

Result explosive

Method Year GLP

Remark Vapor can form an explosive mixture with air at room temperature

Reference Shell Chemicals data sheet IS 3.2.4 dated June 1992

2.11 Oxidizing Properties

Result no oxidizing properties

Method Year GLP

Reference Shell Chemicals Data sheet IS 3.2.4 dated June 1992

3 ENVIRONMENTAL FATE AND PATHWAYS

3.1.1 Photodegradation

Type air Light Source Sun light

Light Spect. ca. 290 - 900 nm

Rel. Intens. = 100 based on Intensity of Sunlight

Spectrum of Substance

lambda (max) [>295nm] nm epsilon (max)

epsilon (295)

Conc. of Substance Temperature degree C

====== D I R E C T P H O T O L Y S I S ========

Halflife t1/2

Degradation % after

Quantum yield

======= I N D I R E C T P H O T O L Y S I S ========

Type air

Indirect photolysis

Sensitizer NO3

Rate Constant $<= .000000000000023 \text{ cm}^3/(\text{molecule*sec})$

Method other (measured): Flash-Photolysis-Visible Absorption

Year 1987

GLP no data

Testsubstance other TS: >= 99% pure

Remark Estimated troposphere half-life of 2-propanol for a "clean"

troposphere with 10 ppt (approximately 2.4* 10E8 radicals/cm³ of

NO3 radicals during night-time hours, t1/2 >= 14.5 d.

Test condition 25 degrees C, total pressure of 133.3 hPa of NO2 diluent Reference Wallington, T.J. et al. (1987): Int J Chem Kinet 19, 243-249.

Type air

Indirect photolysis

Sensitizer OH

Rate constant = .000000000051 cm3/(molecule*sec)

Method OECD Guide-line draft "Photochemical Oxidative Degradation in the

Year 1987 GLP no data Testsubstance no data

Remark Estimated troposheric half-life of 2-propanol with an average

tropospheric OH radical concentration of 5*10E5 radicals/cm³, t1/2 =

3.1 d.

Test condition smog-chamber: 27 degrees C, 0.105 - 0.110 Mpa, UV-A lamps (1.5 m

long, 65 W Hg low pressure lamps, TL 65-80 W/05), maximum volume-averaged UV intensity (k1 of NO2 photolysis amounted to about k1=0.9 min-1) corresponding to twice the solar UV intensity

at sea level and midlatitudes, OH source: HONO photolysis.

Reference Kloepffer, W. et al. (1988): Ecotoxicol Environ Safety 15, 298-319.

Type air

Indirect photolysis

Sensitizer OH

Rate constant = .0000000000521 cm3/(molecule*sec)

Method other (measured): Photodegradation by Indirect Photolysis

Year 1990 GLP no data Testsubstance no data

Remark Estimated tropospheric half-life of 2-propanol, with an OH radical

concentration of 5*10E5 radicals/cm3, t1/2 = 3.1 d.

Test condition 25 degrees C

Reference Atkinson, R. (1990): J Phys Chem Ref Data (1992), 21(6):1125-1568

Type air

Indirect photolysis

Sensitizer OH

Conc. of Sensitizer 500000 molecules/cm3

Rate constant = .0000000000521 cm3/(molecule*sec)

Degradation = 50% after 6.2 days

Method other (measured): AOP Computer Program, Vers. 1.53, Syracuse

Research Center (based upon reference).

Year 1994 GLP no data Testsubstance no data

Remark Half-life refers to 12 hour-days

Reference Atkinson, R.: A structure-activity relationship for the estimation of

rate constants for the gas-phase reactions of OH radicals with organic

compounds. Int J Chem Kinet 19, 799-828 (1987)

Type air

Indirect photolysis

Sensitizer OH

Rate constant = .0000000000532 cm3/(molecule*sec)

Method other (calculated): Prediction of the OH reaction rate constant by

correlation of the negative logarithm of OH reaction rate constants with the first vertical ionisation energy of organic compounds in the

gas phase.

Year 1984 GLP no data Testsubstance no data

Remark The correlation of the negative logarithm of OH reaction rate

constants with the first vertical ionisation energy of organic compounds in the gas phase enables the prediction of kOH with a probability of about 90%; estimated tropospheric half-life of 2 propanol, assessing a global diurnal mean of OH radical concentration

of 5*10E5 radicals/cm3, t1/2 = 3.0 d.

Test condition 27 degrees C

Reference Guesten, H. et al. (1984): J Atmos Chem 2, 83-93.

Type air

Indirect photolysis

Sensitizer OH

Rate constant = .0000000000548 cm3/(molecule*sec)

Method other (measured): Flash-Photolysis Resonance-Asorption Technique

Year 1978 GLP no data Testsubstance no data

Remark Estimated tropospheric half-life of 2 propanol, with an OH radical

concentration of 5*10E5 radicals/cm3, t1/2 = 2.9 d.

Test condition 23 degrees C, 200 hPa

Reference Overend, R., Paraskevopoulos, G. (1978): J Phys Chem 82, 1329-

1333.

Type air

Indirect photolysis

Sensitizer OH

Rate constant = .0000000000062 cm3/(molecule*sec)

Method other (measured): Photodegradation by Indirect Photolysis

Year 1985 GLP no data Test substance no data

Remark Estimated tropospheric half-life of 2-propanol, with an OH radical

concentration of 5*10E5 radicals /cm3, t1/2 = 2.6 d.

Test condition room temperature

Reference Atkinson, R. (1985): Chem Rev 85, pp. 69-75, 135-153, 182-201.

Type air

Indirect photolysis

Sensitizer OH

Rate constant = .000000000071 cm3/(molecule*sec)

Method other (measured): Environmental Chamber Photooxidation Study

Year 1976 GLP no data Testsubstance no data

Remark Estimated atmospheric half-life of 2 propanol, with an average

ambient OH radical concentration of 5/10E6 radicals/cm3, t1/2 - 5.4

h.

Test condition Smog-chamber: ca. 32 degrees C, 1013 hPa, OH source: HONO

photolysis and the reaction of NO2 with NO.

Reference Lloyd, A.C. et al. (1976): Chem Phys Lett. 42, 205-209.

3.1.2 Stability in Water

Type abiotic

Remark Propan-2-ol is not susceptible to hydrolysis.

Reference P.H. Howard, Handbook of Environmental Fate and Exposure Data

for OrganicChemicals, Lewis Publishers Inc., Chelsea Michigan USA,

1990, p.304 -309.

Type biotic

t1/2 pH4 at degree C

t1/2 pH7 = 5 days at 20 degree C

t1/2 pH9 at degree C t1/2 pH at degree C Degradation pH 7 at 20 degree C:

= 50 % after 5 days

Method other Year 1979 GLP no data

Test substance as prescribed by 1.1 - 1.4

Test condition APHA-219 (1971) at 20 degree C.

Reference Bridie, A.L., Wolff, C.J.M. & Winter, M., Water Research, 13, 1979,

627 - 630.

3.3.1 Transport between Environ. Compart.

Type volatility
Media water - air
Method other
Year 1982

Result The evaporation half-life of propan-2-ol in a model river with a depth

of 1 m and a current of 1 m/s at a wind velocity of 3 m/s is calculated

to be 85.2 hours.

Reference Lyman, W.J. et al., Chemical Property Estimation Methods.

Environmental behavior of organic compounds. McGraw-Hill Book Company, New York (USA), 1982, Chpt 15.

Type volatility
Media soil - air
Method other
Year 1990

Result Propan-2-ol is relatively volatile and would therefore readily

evaporate from dry soil and surfaces.

Reference Howard, P.H., Handbook of Environmental Fate and Exposure Data

for Organic Chemicals, Vol. 2. Lewis Publishers, Inc. Chelsea,

Michigan (USA), 1990, p. 304 309.

Method OECD Guide-line draft

Year 1990 GLP no data

Test substance as prescribed by 1.1 - 1.4

Type volatility

Medium other: domestic waste - air Method other: Evaporation Study

Year 1978 GLP no data

Remark The rate of evaporation of diluted (1:1 v/v mixture with water) and

undiluted 2-propanol from pulverized domestic waste (0.5 m deep, 1 m2) over the first 10 min was 1.90 and 3.22 kg.m2 per h, and from

200-440 min 0.65 and 0.15 kg/m² per h respectively

Test condition 2-Propanol (1:1 v/v mixture with water):

initial weight of 2-propanol 7.2 kg, initial weight of water 10 kg, weight of waste 174 kg, waste surface temperature 24-37 degrees C, ambient air temperature ca. 22 degrees C, mean wind speed ca. 4.5

m/sec

2-Propanol (undiluted):

initial weight of 2-propanol 19.5 kg, weight of waste 177 kg. waste surface temperature ca. 18 degrees C, ambient air temperature ca. 12

degrees C, mean wind speed ca. 0.5 m/sec

Test substance technical grade

Reference Jones, C.J. McGugan, P.J. (1977/78): J Hazard Mater. 2, 235-251.

Type volatility Medium water-air

Method other: Evaporation Study

Year 1978 GLP no data

Remark The rate of evaporation of diluted (1:1 v/v mixture with water) and

undiluted 2-propanol from a shallow pool (1 m2) was 1.5 and 1.1

kg/m2 per h, respectively.

Test condition 2-Propanol (1:1 v/v mixture with water):

initial weight of 2-propanol 7.2 kg, initial weight of water 10 kg, pool temperature ca. 20 degrees C, ambient air temperature ca. 22 degrees

C, mean wind speed ca. 4.5 m/sec.

2-Propanol (undiluted):

initial weight of 2-propanol 18.25 kg, pool temperature ca. 13 degrees C, ambient air temperature ca. 12 degrees C, mean wind speed ca. 0.5

m/sec.

Test substance technical grade

Reference Jones, C.J. McGugan, P.J. (1977/78): J Hazard Mater. 2, 235-

2513.3.2

3.2.2 Distribution

Media air - biota - sediment(s) - soil - water Method Calculation according Mackay, Level I

Year 1981

Result Air 22.3 %m;

Water 77.7 %m; Soil 0.0 %m; Sediment 0.0 %m; Biota 0.0 %m.

Reference Mackay, D. & Paterson, S., Calculating Fugacity, Environm. Sci.

Technol., 15(9), 1981, 1006 - 1014.

3.4 Mode of Degradation in Actual Use

1 10/05/94 RS:1 RE:1

Result In air propan-2-ol will be rapidly transformed through reaction with

OH-radicals.

Reference P.H. Howard, Handbook of Environmental Fate and Exposure Data

for Organic Chemicals, Lewis Publishers Inc., Chelsea Michigan

USA, 1990, p 304 - 309

2 10/05/94 RS:1 RE:1

Result In water propan-2-ol will be lost by biodegradation.

Reference P.H. Howard, Handbook of Environmental Fate and Exposure Data for

Organic Chemicals. Lewis Publishers Inc., Chelsea Michigan USA,

1990, p. 304 - 309.

3.5 Biodegradation

Type aerobic

Inoculum predominantly domestic sewage, non-adapted

Concentration 1.5 mg/l related to Test substance

Degradation = 49 % after 5 day Results readily biodegradable

Kinetic

Method other
Year 1979
GLP no data

Testsubstance as prescribed by 1.1 - 1.4

Test condition APHA-219 (1971) at 20 degree C.

Reference A) Bridie, A.L., Wolff, C.J.M., & Winter, M. BOD and COD of some

Petrochemicals, Water Research, 13, 1979, p. 627 -630. B) Idem, Shell Group Research Report, AMGR.0224.74

Type aerobic

Inoculum domestic sewage, non-adapted

Concentration 3, 7 and 10 mg/l related to Test substance

Degradation = 77 % after 10 day Results readily biodegradable

Kinetic 5 day = 28

10 day = 77 15 day = 8020 day = 78

Method other Year 1974 GLP no data

Testsubstance as prescribed by 1.1 - 1.4
Test condition As described in reference.

Reference Price, K.S. Waggy, G.T. & Conway, R.A., Brine Shrimp Bioassay

and Seawater BOD of Petrochemicals, J. Water Pollut. Contr. Fed.,

46,1974, 63 - 77.

Type aerobic

Inoculum domestic sewage, non-adapted

Concentration 3, 7 and 10 mg/l related to Test substance

Degradation = 42 % after 10 day

Results

Kinetic 5 day = 13

10 day = 42 15 day = 6020 day = 72

Method other Year 1974 GLP no data

Testsubstance as prescribed by 1.1 - 1.4

Test condition As described in reference. The test medium is artificial seawater.

Reference Price, K.S., Waggy, G.T. & Conway, R.A., Brine Shrimp Bioassay

and Seawater BOD of Petrochemicals, J. Water Pollut. Contr. Fed.,

46, 1974, 63 - 77.

4. ECOTOXICITY

4.1 Acute/Prolonged Toxicity to Fish

Type flow through

Species Pimephales promelas

Unit mg/l Exposure Period 96 hour

NOEC LC0

LC50 = 9640

LC100

Analyt. Monitoring yes
Method other
Year 1983
GLP no data

Testsubstance as prescribed by 1.1 - 1.4
Test condition As described in reference.

Reference Veith, G.D., Call, D.J. & Brooke, L.T., Estimating the Acute Toxicity

of Narcotic Industrial Chemicals to Fathead Minnows. In: Bishop, W.E., Cardwell, R.D. & Heidolph, B.B. Eds. Aquatic Toxicology and Hazard Assessment: 6th Symp., ASTM STP 802, Philadelphia

(USA), 1983, 90 - 97.

4.2 Acute Tox. to Aquatic Invertebrates

Species Daphnia magna

Unit mg/l Exposure Period 24 hour

NOEC

EC0

EC50 > 10000

EC100

Analyt. Monitoring no
Method other
Year 1977
GLP no data

Testsubstance as prescribed by 1.1 - 1.4
Test condition As described in reference.

Reference Bringmann, G. & Kuehn, R., Results of the Damaging Effect of

Water Pollutants on Daphnia magna, Z. Wasser Abwasser Forsch.,

10(5), 1977, 161 - 166.

Species Crangon crangon

Unit mg/l Exposure Period 48 hour

NOEC EC0

EC50 EC100

LC50 = 1400
Analyt. Monitoring no
Method other
Year 1974
GLP no data

Testsubstance as prescribed by 1.1 - 1.4

Remark The species is a marine invertebrate.

Test condition As described in reference.

Renewal.

Reference Blackman, R.A.A., Toxicity of Oil-Sinking Agents, Mar. Pollut.

Bull., 5, 1974, 116 - 118.

4.3 Toxicity to Aquatic Plants e.g. Algae

Species Scenedesmus quadricauda

Endpoint growth rate

Unit mg/l Exposure Period 7 day

NOEC LOEC EC0 EC10 EC50

Analyt. Monitoring no
Method other
Year 1980
GLP no data

Testsubstance as prescribed by 1.1 - 1.4
Test condition As described in reference.

Remark Toxicity threshold concentration = 1800 mg/l.

Reference Bringmann, G. & Kuehn, R., Comparison of the Toxicity Thresholds

of Water Pollutants to Bacteria, Algae and Protozoa in the Cell

Multiplication Inhibition Test, Water Research, 14, 1980, 231 - 241.

4.4 Tox. to Microorganisms e.g. Bacteria

Type aquatic

Species Pseudomonas putida

Unit mg/l Exposure Period 16 hour

EC0 EC10 EC50 LOEC

Analyt. Monitoring no
Method other
Year 1980
GLP no data

Testsubstance as prescribed by 1.1 - 1.4
Test condition As described in reference.

Remark Toxicity threshold concentration = 1050 mg/l.

Reference Bringmann, G. & Kuehn, R. Water Research, 14, 1980, 231 - 241.

Type aquatic

Species Entosiphon sulcatum

Unit mg/l Exposure Period 72 hour

EC0 EC10 EC50

LOEC

Analyt. Monitoring no
Method other
Year 1980
GLP no data

Testsubstance as prescribed by 1.1 - 1.4
Test condition As described in reference.

Remark Toxicity threshold concentration = 4930 mg/l.

Reference Bringmann, G. & Kuehn, R., Water Research, 14, 1980, 231 - 241.

Type aquatic

Species Microcystis aeruginosa

Unit mg/l Exposure Period 8 day

EC0

EC10 EC50 LOEC

Analyt. Monitoring

Method other Year 1978 GLP no data

Testsubstance as prescribed by 1.1 - 1.4
Test condition As described in reference.

Remark Toxicity threshold concentration = 1000 mg/l.

Reference Bringmann, G. & Kuehn, R., Grenzwerte der Schadwirkung

wassergefaehdender Stoffe gegen Blaualgen (Microcystis aeruginosa) und Gruenalgen (Scenedesmus quadricauda) im

Zellvermehrungshemmtest, Vom Wasser, 50, 1978, 45 - 60.

Type aquatic

Species Chilomonas paramecium (Protozoa)

Exposure period 48 h
Unit mg/l
Analytical monitoring no
TT: = 104

Method other: Cell Multiplication Inhibition Test

Year 1980 GLP no data Testsubstance no data

Remark TT = toxicity thresholdTest condition pH 6.9; 20 degrees C

Reference Bringmann, G. et al. (1980): Z. Wasser Abwasser Forsch. 13, 170-

173.

4.5.2 Chronic Toxicity to Aquatic Invertebrates (Daphnia)

Species Daphnia magna (Crustacea)

Exposure period 16 day
Unit mg/l
Analyt. monitioring yes

NOEC = 141

Method other: Prolonged Toxicity Test

Year 1985 GLP no data Testsubstance no data

Remark NOEC on growth: highest concentration which did not result in a

significant reduction in growth at p < 0.001.

Reference Hermens, J. et al (1985): Aquatic Toxicol. 6, 209-217.

Species Daphnia magna (Crustacea)

 Exposure period
 21 day

 Unit
 mg/l

 Analyt. Monitoring
 no

 NOEC
 = 30

 EC50
 > 100

 EC29
 = 100

Method other: UBA-Verfahrensvorschlag: Verlaengerter Toxizitaetstest bei

Daphnia magna

Year 1984 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Huel-Bericht DL 106, 1988 (unveroeffentlicht)

4.6.2 Toxicity to Terrestrial Plants

Species Lactuca sativa

Endpoint other
Unit mg/l
Exposure Period 3 day

NOEC

EC50 = 2100

LC50

Method other Year 1977 GLP no data

Testsubstance as prescribed by 1.1 - 1.4

Remark The effect parameter reported is inhibition of seed germination.

Test condition As described in reference.

Reference Reynolds, T. An Anomalous Effect of Isopropanol on lettuce

germination. Plant Sci. Lett., 15, 1977, 25 - 28.

4.6.3 Tox. to Other Non-mamm. Terr. Species

Species other
Endpoint mortality
Unit ppm
Exposure Period 48 hour

NOEC

LC0

LC50 10200 - 13340

LC100

Method other Year 1976 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Remark The insects are exposed to the propan-2-ol in nutrient medium.

Remark The test species reported are Drosophila simulans and Drosophila

melanogaster

Test condition As described in reference.

Reference David, J. & Bocquet, C. Compared Toxicities of Different Alcohols

for Two Drosophila Sibling Species. Comp. Biochem. Physiol., 54C,

1976, 71 - 74.

5. TOXICITY

5.1.1 Acute Oral Toxicity

Type LD50 Species rat

Value = 5280 mg/kg

Method other Year 1944 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Lehman, A. J., Chase, H.F., (1944) J. Lab. Clin. Med. 29: 561 - 567.

Type LD50 Species rat

Value = 5840 mg/kg

Method other Year 1948 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Smyth, H.F., Carpenter, C.P., (1948), J. Ind. Hyg. Toxicol., 30: 63-70.

Type LD50 Species rat

Value = 4710 mg/kg

Method other Year 1971 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Remark Three LD50 values presented: 5.6, 6.0 and 6.8 ml/kg for neonates,

young adult and old rats respectively. Quoted value is a mean of the

three as presented in IPCS EHC document on 2-propanol (1990).

Reference Kimura, E.T., Ebert, D.M., Dodge, P.W., (1971), Toxicol. Appl.

Pharmacol., 19: 699 - 703.

Type LD50 Species rat

Value = 5500 mg/kg

Method other Year 1985 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Guseinov, V.G., (1985), Gig. Tr. Prof. Zabol. (7): 60-62.

Type LD50 Species mouse

Value = 4475 mg/kg

Method other Year 1985 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Guseinov, V.G., (1985), Gig. Tr. Prof. Zabol., (7): 60-62.

Type LD50 Species rabbit

Value = 5030 mg/kg

Method other Year 1944 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Lehman, A.J., Chase, H.F. (1944), J. Lab. Clin. Med., 29: 561 - 567.

Type LD50 Species rabbit

Value = 7990 mg/kg

Method other Year 1972 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Munch, J.C., (1972), Ind. Med. 41: 31 - 33.

Type LD50 Species dog

Value = 4830 mg/kg

Method other Year 1944 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Lehman, A.J., Chase, H.F., (1944), J. Lab. Clin. Med., 29: 561 - 567.

5.1.2 Acute Inhalation Toxicity

Type LC50
Species rat
Exposure Time 4 hour
Value = 72.6 mg/l
Method other
Year 1985

GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Guseinov, V.G., (1985), Gig. Tr. Prof. Zabol. (7): 60 - 62.

Type LC50 Species rat Exposure Time 8 hour

Value = 51.045 mg/l

Method other Year 1979 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Laham, S., Potvin, M., Schrader, K., Marino, I. (1979), Drug. Chem.

Toxicol. 3: 343 - 360.

Type LC50
Species mouse
Exposure Time 2 hour
Value = 53 mg/l
Method other
Year 1985
GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Guseinov, V.G., (1985), Gig. Tr. Prof. Zabol, (7), 60-62.

5.1.3 Acute Dermal Toxicity

Type LD50 Species rabbit

Value = 12870 mg/kg

Method other Year 1948 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Smyth, H.F., Carpenter, C.P., (1948), J. Ind. Hyg. Toxicol.,30: 63 -70.

5.1.4 Acute Toxicity, Other Routes

Type LC50 Species rat

Route of

Administration i.v.

Exposure Time

Value = 1088 mg/kg

Method other Year 1985 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Tichy, M., Trcka, V., Roth, Z., Krivucova, M., (1985), Environ.

Health. Perspect., 61: 321 328.

Type LC50 Species rat

Route of

Administration i.p.

Exposure Time

Value = 2830 mg/kg

Method other Year 1985 GLP no data

Testsubstance as prescribed by 1.1 - 1.4

Reference Tichy, M., Trcka, V., Roth, Z., Krivocova, M., (1985), Environ.

Health. Perspect., 61: 321 328.

5.2.1 Skin Irritation

Species rabbit
Result not irritating
Classification not irritating
Method other
Year 1975
GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Nixon, G.A., Tyson, C.A., Wertz, W.C., (1975), Toxicol. Appl.

Pharmacol. 31: 481 -490.

5.2.2 Eye Irritation

Species rabbit
Result irritating
Classification irritating
Method Draize-Test
Year 1973
GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Marzulli, F.N., Ruggles, D.I., (1973), J. Assoc. Off. Anal. Chem., 56:

905 - 914.

Species rabbit

Result moderately irritating

Classification irritating
Method Draize-Test
Year 1980
GLP no

Testsubstance as prescribed by 1.1 - 1.4

Reference Griffith, J.F., Nixon, G.A., Bruce, R.D., Reer, P.J., Bannan, E.A.,

(1980), Toxicol. Appl. Pharmacol. 55: 501 - 513.

Species rabbit Result irritating

Classification R36
Method Draize-Test
Year 1986
GLP Yes

Testsubstance as prescribed by 1.1-1.4

Reference Exxon Biomedical Sciences Inc. (1986) Ocular Irritation Study in

Rabbits (Isopropanol). EBSI Document No. 86MRL272.

Species rabbit

Result moderate irritating

Classification irritating
Method Draize-Test
Year 1987
GLP no data

Testsubstance as prescribed by 1.1 - 1.4

Remark Study gives maximum Draize score of 37 (out of 110) and provided

conflicting data from 2 tests, one moderate and the other severely

irritating.

Reference Morgan, R.L., Sorenson, S.S., Castles, T.R., (1987), Food. Chem.

Toxicol., 25: 609 -613.

5.3 Sensitization

Type Buehler-Test
Species guinea pig
Result not sensitizing
Classification not sensitizing

Method other Year 1980 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Remark Result given as "No sensitisation (0/20)". No other data on

sensitisation have been found.

Reference P & G, Unpublished data.

5.4 Repeated Dose Toxicity

Species rat

Strain Fischer 344
Sex male/female
Route of Administration inhalation
Exposure Period 13 weeks

Frequency of Treatment 6 hours/day, 5 days/week

Post Exposure

Observ. Period none

Doses 0, 100, 500, 1500 and 5000 ppm

Control Group yes NOEL 500 LOEL 1500

Method other (US EPA TCSA Test Guidelines)

Year 1994 GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Remark No exposure-related mortalities occurred. Narcotic effects were noted

during exposures to 1500 and 5000 ppm. Ataxia was observed

following exposure to 5000 ppm.

Decreases in body weight were also observed at the end of the first week of exposure to 5000 ppm. The only microscopic change observed was hyaline droplets within the kidneys of all male rats (including controls). The size and frequency of these droplets were increased in the exposed groups. Thus, repeated exposures produced toxic effects only at the highest concentration (5000 ppm) and a

kidney change in male rats of unknown biological significance.

Reference Burleigh-Flayer, H.D., et al. (1994). Fundam. Appl. Toxicol. 23, 421-

428.

Species rat
Strain Wistar
Sex male
Route of Administration inhalation
Exposure Period 13 or 20 weeks

Frequency of Treatment

4 hours/day, 5 days/week.

Post Exposure

Observ. Period none

Doses 400, 1000,4000, 8000 ppm for 12 weeks; 1000 and 8000ppm for 20

weeks

Control Group yes

 $\begin{array}{ll} \text{NOEL} & = 400 \text{ ppm} \\ \text{LOEL} & = 1000 \text{ ppm} \end{array}$

Method other Year 1991 GLP no data

Testsubstance as prescribed by 1.1 - 1.4

Remark No significant differences appeared between the groups exposed to

400 ppm and the control group in body weight and in hematological and serum chemistry tests. There was inhibition of body weight and marked local irritation in groups given 1000 ppm or more, decrease in erythrocyte and hemoglobin values in groups given 4000 ppm or more, and increases in serum GOT and GPT, and total cholesterol in the 8000 ppm group. Nakaseko et al. also exposed rats to 1000 and 8000 ppm isopropanol for 20 weeks for nerve conduction studies,

which was reported separately (see also section 5.10).

Reference Nakaseko, H., Teramoto, K., Horiguchi, S., Wakitani, F., Yamanoto,

T., Adachi, M., Tanaka, H., Hozu, S., Jpn. J. Ind.Hlth. 33(3): 200-

201, 1991.

Species rat
Strain F-344
Sex female
Route of Administration Inhalation

Exposure period 90 days or 63 days

Ferquency of exposure 1/2 6 hours/day for 5 days /week for 13 weeks

1/2 6 hours/day for 5 days /week for 9 weeks

Post exposure

observation period 2 weeks post exposure

Doses 5000 ppm

Control group yes

NOEL NA

LOEL 5000 ppm

Method other

Year 1994

GLP yes

Testsubstance as prescribed in 1.1-1.4

Remark Increases in motor activity were seen following exposure to 5000 ppm

of isopropanol. After 9 weeks of exposure, complete recovery was noted two days post exposure. After 13 weeks of exposure, complete

reversibility was noted at 2 weeks post exposure.

Reference Gill, M., Burleigh-Flayer, H., Bevan. C., Gardiner, T., Kapp, R.,

Isopropanol Ninety-Day Vapor Inhalation Neurotoxicity Study in

Female F-344 Rats. (Abstract) The Toxicologist, Vol. 14 (1), 1994.

Species rat
Strain no data
Sex male/female
Route of Administration drinking water
Exposure Period 27 weeks
Frequency of Treatment continuous

Post Exposure

Observ. Period none

Doses 600 & 2300 mg/kg for males, 1000 & 3900 mg/kg for females

Control Group yes

NOEL = 600 - 1000 mg/kg bw dLOEL = 2300 - 3900 mg/kg bw d

Method other Year 1944 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Remark The male rats showed some decreased body weight gains during the

first thirteen weeks of the study, and then increased body weight gain for the reminder of the study. The female rats showed decreased body weight gain throughout the study. No gross or microscopic

abnormalities were noted.

Reference Lehman, A.J., Chase, H.F., (1944), J. Lab. Clin. Med. 29: 561 - 567.

Species rat
Strain no data
Sex male

Route of Administration drinking water Exposure Period 12 weeks Frequency of Treatment continuous

Post Exposure

Observ. Period none

Doses 1,2,3 and 5 percent

Control Group yes

NOEL = 1% (870 mg/kg/day) LOEL = 2% (1280 mg/kg/day)

Method other Year 1993 GLP no data

Testsubstance as prescribed by 1.1 - 1.4

Remark The relative organ weights of liver, kidneys, and adrenals were

significantly increased in a dosedependent manner. No histological alterations could be attributed to the dosing, apart from a dose-dependent increase in formation of hyaline casts and droplets in the proximal tubules of the kidneys. Dorsal hippocampal glial fibrillary

acidic protein (GFAP) was unaffected after treatment.

Reference Pilegaard, K. and Ladefoged, O. (1993) In Vivo 7:325-330

Species mouse
Strain CD-1
Sex male/female
Route of Administration inhalation
Exposure Period 13 weeks

Frequency of Treatment 6 hours/day, 5 days/week

Post Exposure

Observ. Period none

Doses 0, 100, 500, 1500 and 5000 ppm

Control Group yes

NOEL = 500 ppmLOEL = 1500 ppm

Method other (US EPA TCSA Test Guidelines)

Year 1994 GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Remark No exposure-related mortalities occurred. Narcotic effects were noted

during exposures to 1500 and 5000 ppm. No exposure-related effects were noted on body weight of the male mice, but increased body weight and body weight gain were observed for the 5000 ppm female mice. No treatmentrelated effects were noted at gross necropsy or at

histopathologic examination.

Reference Burleigh-Flayer, H.D., et al. (1994). Fundam. Appl. Toxicol. 23, 421-

428.

Species rat

Strain Fischer 344
Sex male/female
Route of Administration inhalation
Exposure Period 13 weeks

Frequency of Treatment 6 hours/day, 5 days/week

Post Exposure

Observ. Period none

Doses 0, 500, 1500 and 5000 ppm

Control Group yes

 $\text{NOEL} = 1500 \\
 \text{LOEL} = 5000$

Method other (US EPA TCSA Test Guidelines)

Year 1994 GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Remark Neurobehavioral evaluations included a functional observation battery

(FOB), motor activity, and neuropathology. There were no changes in FOB, but increased motor activity was noted in female rats of the 5000 ppm group at week 9 and 13. Neuropathological examination

revealed no exposure-related lesions in the nervous system.

Reference Burleigh-Flayer, H.D., et al. (1994). Fundam. Appl. Toxicol. 23, 421-

428.

5.5 Genetic Toxicity in Vitro

Type Salmonella typhimurium reverse mutation assay

System of Testing TA 98, 100, 1535, 1537

Concentr. 180 mmol/plate Metabolic Activation with and without

Result negative
Method other
Year 1980
GLP no data

Testsubstance as prescribed by 1.1 - 1.4

Remark A report on screening of many chemicals which could be constituents

of tobacco smoke.

Reference Florin, I., Rutberg, L., Curvall, M., and Enzell, O.R. Toxicology

15:219-232, 1980

Type Sister chromatid exchange assay
System of Testing Chinese hamster V79 fibroblasts
Concentr. 3.3, 10, 33.3 and 100 mmol/l

Metabolic Activation with and without

Result negative
Method other
Year 1987
GLP no data

Testsubstance as prescribed by 1.1 - 1.4

Reference Von der Hude, W., Scheutwinkel, M., Gramlich, U., Fissler, B.,

Busler, A., In Vitro Environ. Mutagen., (1987), 9: 401 - 410.

Type Salmonella typhimurium reverse mutation assay System of Testing TA 97, 98, 100, 102, 104, 1535, 1537, 1538

Concentr. 100 mmol/plate Metabolic Activation with and without

Result negative
Method other
Year 1992
GLP Yes

Testsubstance as prescribed by 1.1 - 1.4

Reference Zeiger, E. et al, Env. Mol. Mut., 19 (Suppl.21): 2 - 141, 1992.

Type HGPRT assay

System of Testing Chinese hamster ovary

Concentr. 0.5 - 5.0 mg/ml Metabolic Activation with and without

Result negative

Method other (US EPA TCSA Test Guidelines)

Year 1993 GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Reference Kapp, R.W. et al. (1993). Environ. Mol. Mutagen., 22: 93-100.

Type Meiotic nondisjunction

System of Testing Neurospora crassa (Strain I x I)

Concentr. no data
Metabolic Activation without
Result negative
Method other
Year 1980
GLP no data

Test substance as prescribed by 1.1 - 1.4

Remark

Reference Griffiths A.J.F. (1980). NIEHS 263-77-C0604CC, Progress Report.

(Cited In: Brockman H.E., de Serres F.J., Ong T-M., DeMarini D.M., Katz A.J., Griffiths A.J.F. and Stafford R.S. (1984). Mutation tests in *Neurospora crassa*. A report of the U.S. Environmental Protection

Agency Gene-Tox Program. Mutat. Res. 133:87-134.)

Type Cell transformation

System of Testing SA7/Syrian Hamster Embryo

Concentr. 62 - 1000 µg/ml

Metabolic Activation without
Result negative
Method other
Year 1978
GLP no data

Test substance as prescribed by 1.1 - 1.4

Remark

Reference Casto B.C. and Hatch G.G. (1978). Progress Report NIH-NCI-N01-

CP-45615. pp. 62-75. (Cited In: Heidelberger C., Freeman A.E., Pienta R.J., Sivak A., Bertram J.S., Casto B.C., Dunkel V.C., Francis M.W., Kakunaga T., Little J.B. and Schechtman L.M. (1983). Cell transformation by chemical agents - a review and analysis of the literature. A report of the U.S. Environmental Protection Agency

Gene-Tox Program. Mutat. Res. 114:283-385.)

5.6 Genetic Toxicity in Vivo

Type Micronucleus Assay

Species mouse

Strain ICR random bred Sex male/female

Route of Administration i.p.

Exposure Period once, bone marrow examined after 24, 48 and 72 hours.

Doses 350, 1173, 2500 mg/kg

Result negative

Method other (US EPA TCSA Test Guidelines)

Year 1993 GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Reference Kapp, R.W. et al. (1993). Environ. Mol. Mutagen., 22: 93-100.

5.7 Carcinogenicity

SpeciesmouseStrainC3HSexmale/femaleRoute of AdministrationinhalationExposure Period5 - 8 months

Frequency of Treatment 3 - 7 hours/day, 5 days/week.

Post Exposure

Observ. Period none

Doses 7700 mg/m3

Control Group yes
Method other
Year 1952
GLP no

Testsubstance as prescribed by 1.1 - 1.4

Remark Two other strains used as well, ABC and C57/BL.
Result No excess of lung tumo rs noted among treated mice.

Reference Weil, C.S., Smyth, H.F., Nale, T.W., Arch. Ind. Hyg. Occup. Med.

(1952), 5: 535 -547.

Species mouse
Strain other
Sex no data
Route of Administration dermal
Exposure Period 52 weeks
Frequency of Treatment 3 times/week.

Post Exposure

Observ. Period none
Doses as supplied

Control Group yes
Method other
Year 1976
GLP no

Testsubstance as prescribed by 1.1 - 1.4

Result No treatment related skin tumo rs reported.

Reference US NIOSH (1976), Cincinnati, Ohio, US National Institute of

Occupational Safety and Health, US Dept. of Health, Education and

Welfare, Public Health Services, Centre for Disease Control (DHEW publication no. (NIOSH)76-142).

SpeciesmouseStrainC3HSexmaleRoute of Administrations.c.

Exposure Period 20 - 40 weeks. Frequency of Treatment once/week

Post Exposure

Observ. Period none
Doses 20 mg
Control Group yes
Method other
Year 1952
GLP no

Testsubstance as prescribed by 1.1 - 1.4

Remark Two other strains used as well, ABC and C57/BL.

Result No excessive incidence of lung tumo rs was noted in treated groups,

although incidence of lung tumours in control groups was high.

Reference Weil, C.S., Smyth, H.F., Nale, T.W., Arch. Ind. Hyg. Occup. Med.

(1952), 5: 535 -547.

Species rat

Strain Fischer 344
Sex male/female
Route of Administration inhalation
Exposure Period 24 months

Frequency of Treatment 6 hours/day, 5 days/week

Post Exposure

Observ. Period none

Doses 500, 2500, 5000 ppm

Control Group yes

NOEL(onco) >5000 ppm for oncogenetici effects

NOEL(toxicity) 500 ppm

Method other (US EPA TCSA Test Guidelines)

Year 1995 GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Remark The main cause of death for the 5000 ppm rats (both sexes) was

chronic renal disease. Chronic renal disease also accounted for much of the mortality of the 2500 ppm male rats. In contrast, the main cause of death for the control animals was large granular lymphocyte leukemia. During exposure, some male and female rats at 5000 ppm showed signs of hypoactivity, lack of startle reflex, and narcosis. Hypoactivity was also observed in some animals at 2500 ppm. Body weight and body weight gain were increased for animals in the 2500 and 5000 ppm groups. Urine chemistry changes indicative of impaired kidney function were noted in the 2500 (males) and 5000 (males and females) ppm groups. In the 2500 and 5000 ppm males and/or females, liver weights were increased, probably as a result of a

> metabolic response (enzyme induction), although there was no histopathological changes noted. There was also a concentrationrelated increase in absolute and relative testes weight seen in males. An excerbation of chronic renal disease was observed in the IPAexposed rats. Observations included mineralization, tubular dilation, glomerulosclerosis, interstitial nephritis and fibrosis, hydronephrosis, and transitional cell hyperplasia. These changes are common in aging rats. An increased incidence of interstitial cell adenomas occurred in the testes of male rats. The frequency of interstitial (Leydig) cell tumors of the testis was also increased in a concentration-related pattern for male rats in the study. The incidence of these tumors in IPA exposed groups was similar to that reported for unexposed rats in the National Toxicology Program and previous studies at this laboratory, while the incidence of these tumors in unexposed rats in this study was well below historical levels. There were no increases in the incidence of other tumor types in the exposed animals compared to the controls.

Reference

Garman, R. Bevan, C., Burleigh-Flayer, H., Gardiner, T., Kapp, R., Neptun, D., Tyler, T. and Wright, G. Isopropanol (IPA) Vapor Inhalation Oncogenicity Study in Fisher 344 Rats.(Abstract No. 979) The Toxicologist, Vol. 15 (1), 1995.

Species mouse Strain CD-1 male/female Sex Route of Administration inhalation **Exposure Period** 18 months 6 hours/day, 5 day/week

Frequency of Treatment

Post Exposure

Observ. Period None

Doses 500, 2500, 5000 ppm

Control Group

NOEL(onco) > 5000 ppm for oncogenetic effects

NOEL(toxicity) 500 ppm (see remarks)

other (US EPA TCSA Test Guidelines) Method

Year

GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Narcosis was observed during exposure in the 2500 and 5000 ppm Remark groups, and following exposure in the 5000 ppm groups. There was

no differences in mortality rate or mean survival time between IPAexposed and control animals. Clinical signs were observed for some male and female mice during exposures to 2500 and 5000 ppm and included hypoactivity, lack of a startle reflex, narcosis, ataxia (5000 ppm only), and prostration (5000 ppm only). Ataxia was the only exposure-related clinical sign that was noted for the 5000 ppm male and female animals following exposure. Increased body weight and/or body weight gain were observed for both sexes in the 2500 and 5000 ppm groups. The 2500 and 5000 ppm male mice from the recovery group continued to increase body weight and body weight gain to study termination even in the absence of IPA exposure. Absolute and

relative liver weights were increased in both male and female mice; relative testes weights were decreased in all IPA-exposed male mice; and absolute and relative brain weights were decreased in the 5000 ppm females. The only organ weight change noted for the recovery animals was a concentration-related absolute and relative increase in liver weight for the male mice. Hematologic parameters were unaffected by IPA exposure. The only gross lesion noted at study termination was seminal vesicle enlargement in the 5000 ppm males. However, microscopic evaluation showed increased incidence of dilation of the seminal vesicles in both the 2500 and 5000 ppm male mice. These effects were not present in the recovery group males. Since the seminal vesicles did not have any associated inflammatory or degenerative changes, the enlargement may have been a result of hormonal imbalance. It is not known whether the narcotic effects of isopropanol could have affected the secretion of the seminal vesicles. Minimal to mild effects to the kidney including renal tubular proteinosis and tubular dilation were observed following 78 weeks of exposure. The incidence of renal tubular proteinosis was generally significantly increased for all male and female treatment groups relative to controls; however the majority of affected animals showed minimal degrees of tubular proteinosis (i.e., only a few tubules affected), there was no concentration-related gradient in either the frequency of severity of this change, and there was no corresponding evidence of alterations to the glomeruli. Mild to moderate degrees of tubular dilation were observed in a small number of females in the 2500 and 5000 ppm groups (significantly increased only for the 5000 ppm group). This finding, however, was not duplicated in male mice (a significant increase was only seen for the 500 ppm group) nor was it accompanied by evidence of tubular cell degeneration or urinary outflow obstruction The increased incidence of kidney effects was not increased in animals in the recovery group. There was no increased frequency of neoplastic lesions in any of the IPA exposed groups, indicating a lack of carcinogenic activity by IPA. Thus, while the kidney findings may be considered treatment-related effects (female mice), their toxicological significance is unclear at this time. The uncertainty about kidney effects, in particular male mice, leatd to a NOEL of 500 ppm, and a LOEL of 2500 ppm (clinical signs), for non-cancer effects for this study.

Reference

Burleigh-Flayer, H., Bevan, C., Gardiner, T., Garman, R., Kapp, R., Neptun, D., Tyler, T. and Wright, G. Isopropanol Vapor Inhalation Oncogenicity Study in CD-10 Mice. (Abstract No. 1219) The Toxicologist, Vol. 14 (1), 1994.

5.8 Toxicity to Reproduction

Type One generation study

Species rat
Strain Wistar
Sex male/female
Route of Administration drinking water

Exposure Period To weaning (day 21 after birth).

Frequency of Treatment continuous

Premating Exposure Period

male 70 days
female 21 days
Duration of Test To weaning
Doses 0.5, 1.0 and 2.0%

Control Group yes
NOEL Parental = 1 %
NOEL F1 Offspring = 1 %

NOEL F2 Offspring

Method other Year 1986 GLP ves

Testsubstance as prescribed by 1.1 - 1.4

Remark Parental rats dosed with 2% isoproanol had decreased body weight

gain and corresponding reduced pup weight gain and decreased survival compared with controls. There was also a dose-related increase in relative liver weights of the F1 animals. There was no effect on reproductive parameters. There was no macroscopic or histopathological changes assoicated with isopropanol treatment. The reproductive NOEL of 1% corresponds to 825 and 625 mg/kg/day for females and males respectively. A further study by Lehman et al (J. Exp. Pharmacol. Expt. Therapy, 85, 61-69, 1945) studied the effects of IPA in drinking water. No effects were found at 2.5% but the

reported data are scant.

Reference British Industrial Biological Research Assoc., Report 0570/3/86.

Type Two generation study

Species rat

Strain Sprague-Dawley
Sex male/female
Route of Administration gavage

Exposure Period Prior to mating and to lactation and weaning of F1 and F2

generations.

Frequency of Treatment daily

Premating Exposure Period

male 10 weeks female 10 weeks

Duration of Test to lactation and weaning
Doses 100, 500 and 1000 mg/kg/day

Control Group yes

NOEL Parental < 500 mg/kg bw/day (BMDL10 = 407 mg/kg/day)(see remarks

below)

Reproduction

NOEL F1 Offspring < 500 mg/kg bw/day(BMDL5 = 449 mg/kg/day)(See remarks below)NOEL F2 Offspring < 500 mg/kg bw/day(BMDL5 = 418 mg/kg/day)(See remarks below)

Method other (US EPA TCSA Test Guidelines)

Year 1995 GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Remark

Remark

Reference

Thirty rats of each sex per group (P1) were dosed once daily by oral gavage with 0, 100, 500 or 1000 mg of isopropanol kg-1 for at least 10 weeks prior to mating. Findings in the parental animals included increased lactation body weight gain in the 500 and 1000 mg/kg females, increased liver and kidney weights in the 500 and 1000 sexes, and centrilobular hepatocyte groups of both hyperthrophy in some P2 males. There was microscopic findings in the kidneys from the mid- and high-dose P1 males and from all treated groups of the P2 males. Exposure to 1000 mg/kg day and to a lesser extent 500 mg/kg day resulted in a reduction in postnatal survival in both F1 and F2 litters. In addition, offspring body weight was reduced during the early postnatal period in the 1000 mg/kg F1 males and in the 1000 mg/kg F2 pups of both sexes. In the 1000 mg/kg group 18/70 F1 weanlings or were euthanized prior to P2 selection. No treatment-related postmortem findings were observed in offspring from either generation. A statistically significant reduction was observed in the male mating index of the 1000 mg/kg P2 males compared to controls. However, no treatment-related microscopic changes in reproductive tissues biologically meaningful differences in other reproductive parameters were noted.

The study-derived NOELs for the F1 and F2 offspring are contingent upon the biological significance ascribed to the effects observed for the 500 mg/kg /day treatment group. There are two perspectives on the interpretation of these observations. A conservative perspective is that the reductions in postnatal survival are treatment- and doserelated effects (U.S. EPA, 1992 U.S. EPA, 1996; Tyl, 1996). Consequently, the NOEL based on this interpretation would be set at 100 mg/kg/day. On the other hand, the NOEL may be set at 500 mg/kg/day if these observations are not deemed biologically significant (Bevan et al., 1995; Harris, 1995). A benchmark dose (BMD) assessment was conducted as a way of clarifying issues surrounding the derivation of effect levels for this study. As described below, this assessment resulted in calculated BMD dosages of 449 and 418 mg/kg low/day for the F1 and F2, respectively as appropriate descriptors for this endpoint. (See Benchmark study below.).

Bevan, C., Tyler, T.R., Gardiner, T.H., Kapp, R.W., Jr., Andrews, L. and Beyer, B.K. Journal of Applied Toxicology, Vol. 15(2) pp. 117-123 (1995).

Harris S.B. (1995). A review of the EPA comments regarding the study entitled "Multi-generation rat reproduction study with isopropanol". Report prepared for the Chemical Manufacturers Association Isopropanol Panel.

Tyl R. W. (1996) February 12, 1996 Letter to the Chemical Manufacturers Association Isopropanol Panel.

US EPA (1992). Review of Section 4 Data - A Two Generatio Reproductive Toxicity Study in Rats with Isopropanol.

US EPA, Draft Final RM1 Risk Assessment of Isopropanol, OPPT,

1996. TSCA Public Docket Number AR-141.

Remark Quantitative Dose-Rsponse Analysis -- Application of the Benchmark

Method to the Multi-Generation Rat Reproduction Study for

Isopropanol.

Remark Chemical Manufacturers Association Isopropanol Panel conducted a

quantitative dose-response analysis of the bioassay data from the multi-generation rat reproduction study noted above using the benchmark dose method to identify the relevant dosage to derive a toxicity value that may contribute in part to safety assessment

decisions for isopropanol.

Remark The reproductive/developmental effects were reported to have

NOAEL between 100 mg/kg/day (USEPA 1992) and 500 mg/kg/day (Bevan et al., 1995). Based upon decrease in mating index observed in the P2 males, a BMDL10 of 407 mg/kg/day was estimated for reproductive effects. A BMDL5 of 418 mg/kg/day was estimated for developmental effects based upon the F2 generation 4-day survival. For the F1 generation 4-day survival, 449 mg/kg bw/day was estimated as BMDL5. The corresponding MLE dosages were 786 (Polynomial model) and 771 mg/kg bw/day (Weibell model) for the reproductive effects, 656 mg/kg bw/day for the F1 postnatal effects,

and 804 mg/kg bw/day for the F2 postnatal effects.

Reference Shipp, A.M., Allen, B.C., Van Landingham, C., Gentry, P.R. and

Crump, K.S., Quantitative Dose-Response Analysis -- Application of the Benchmark Method to the Multi-Generation Rat Reproduction Study for Isopropanol. Final Report prepared by ICF Kaiser, KS Crump Division, 602, East Georgia Avenue, Ruston, Louisiana for Chemical Manufactureres Association Isopropanol Panel, April 1996.

Type One generation study

Species rat
Strain Wistar
Sex male/female
Route of Administration drinking water

Exposure Period prior to mating and to lactation and weaning of F1 and generation

Frequency of Treatment continuous

Premating Exposure Period

male 8 weeks female 8 weeks

Duration of Test

Doses 2 and 3% isopropanol in drinking water

Control Group yes NOEL Parental = 2% NOEL F1 Offspring = 2%

NOEL F2 Offspring

Method other Year 1977 GLP no

Testsubstance as prescribed by 1.1 - 1.4

Remark Isopropanol was administered as a 3% solution in the drinking water.

UNEP Publications

Reduced parental body weight gain, food and water consumption

were observed in the isopropanol-treated animals compared with controls. In addition, fertility, litter size, and pup weights at postnatal days 4 and 21 were reduced in the 3% treatment group compared with the controls. The dose was dropped to 2% isopropanol and the parental animals were re-mated to provide litters for a developmental toxicity evaluation. No parental toxicity or reproductive toxicity was

noted.

Reference Gallo, M.A., Oser, B.I., Cox, G.E., and Bailey, D.E. (1977) Toxicol.

Appl. Pharmacol. 41:35

5.9 Developmental Toxicity/Teratogenicity

Species rat
Strain Wistar
Sex female

Route of Administration drinking water

Exposure Period Days 6 - 16 of pregnancy.

Frequency of Treatment continuous

Duration of Test To day 20 of pregnancy.

Doses 0.5, 1.25, 2.5%

 $\begin{array}{lll} \text{Control Group} & \text{yes} \\ \text{NOEL Maternal Toxicity} & = 0.5 \% \\ \text{NOEL Teratogenicity} & = 0.5 \% \\ \text{Method} & \text{other} \\ \text{Year} & 1986 \\ \text{GLP} & \text{yes} \\ \end{array}$

Testsubstance as prescribed by 1.1 - 1.4

Remark Maternal body weights were significantly decreased from gestational

days 7-16. Animals in the 1.25% and 2.5% dose groups exhibited reduced food and water consumption during the treatment period. In the 1.25% and 2.5% dose groups, fetal body weights were reduced on a per fetus basis, but not on a per litter basis. No teratogenic effects were observed; but, delayed ossification of the skeleton was noted in the 1.25% and 2.5% dose groups, consistent with retarded

development as a result of maternal toxicity.

Reference British Industrial Biological Research Assoc., Report no. 0570/2/86.

Species rat

Strain Sprague-Dawley

Sex female Route of Administration gavage

Exposure Period 6 - 15 of gestation

Frequency of Treatment daily

Duration of Test To day 20 of pregnancy

Doses 0, 400, 800 and 1200 mg/kg/day

Control Group yes

NOAEL Maternal Toxicity = 400 mg/kg bw/day NOAEL Developmental Toxicity = 400 mg/kg bw/day

Method other (US EPA TCSA Test Guidelines)

Year 1994

GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Remark No dams aborted or delivered early. Two dams died at 1200 mg/kg

and one dam died at 800 mg/kg. Reduced maternal gestational weight gain on gestational days 0 to 20 associated with significantly reduced gravid uterine weights were noted in the high-dose animals. All gestational parameters were equivalent across groups. Fetal body weights per liter were significantly reduced at the two highest doses. There were no adverse maternal or developmental effects at 400 mg/kg. No evidence of increased teratogenicity was observed at any

dose tested. Therefore, isopropanol was not teratogenic to CD rats.

Reference Tyl, R.W., Masten, L.W., Marr, M.C., Myers, C.B., Slauter, R.W.,

Gardiner, T.H., Strother, D.E., McKee, R.H., and Tyler, T.R. (1994)

Fundam. Appl. Toxicol. 22:139- 151.

Species rabbit

Strain New Zealand white

Sex female Route of Administration gavage

Exposure Period Days 6 - 18 of pregnancy

Frequency of Treatment daily

Duration of Test to day 28 of pregnancy
Doses 120, 240 and 480 mg/kg/day

Control Group yes

NOAEL Maternal Toxicity = 240 mg/kg bw/day NOAEL Developmental Toxicity = 480 mg/kg bw/day

Method other (US EPA TCSA Test Guidelines)

Year 1994 GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Remark No does aborted or delivered early. Four does died at 480 mg/kg.

Maternal body weights were significantly reduced during treatment and clinical signs of toxicity were observed at 480 mg/kg. No adverse maternal effects were noted at 120 or 240 mg/kg. All gestational parameters were equivalent across groups. No evidence of increased teratogenicity was observed at any dose tested. Therefore, isopropanol

was not teratogenic to NZW rabbits.

Reference Tyl, R.W., Masten, L.W., Marr, M.C., Myers, C.B., Slauter, R.W.,

Gardiner, T.H., Strother, D.E., McKee, R.H., and Tyler, T.R. (1994)

Fundam. Appl. Toxicol. 22:139- 151.

Species rat

Strain Sprague-Dawley

Sex female Route of Administration gavage

Exposure Period day 6 of pregnancy to day 21 postnatal.

Frequency of Treatment daily

Duration of Test Day 68 post natal.

Doses 200, 700 and 1200 mg/kg/day

Control Group yes

NOEL Maternal Toxicity = 700 mg/kg bw/day

NOEL Developmental Neurotoxicity = 1200 mg/kg bw/day

Method other (US EPA TCSA Test Guidelines)

Year 1994 GLP yes

Testsubstance as prescribed by 1.1 - 1.4

Remark This study was specifically designed to investigate developmental

neurotoxicity. One high-dose dam died on postnatal day 15, but there were no other clinical observations or effects on maternal weight, food consumption, or gestation length. Pup survival, weight, sex ratio, and sexual maturation were unaffected. There were no biologically significant findings in the behavioral tests, no changes in organ weights, and no pathological findings that could be attributed to

isopropanol exposure.

Reference Bates, H.K., McKee, R.H., Bieler, G.S., Gardiner, T.H., Gill, M.W.,

Strother, D.E., and Masten, L.W. (1994) Fundam. Appl. Toxicol.

22:152-158.

Species rat

Strain Sprague-Dawley

Route of Administration inhalation

Exposure Period gestational days 1-19

Frequency of Treatment 7 hours/day
Duration of Test gestational day 20

Doses 3500, 7000 and 10000 ppm

Control Group yes

NOEL Maternal Toxicity = 3500 ppm NOEL Teratogenicity < 3500 ppm Method not specified

GLP yes

Remark The animals showed unsteady gait and narcotization during initial

exposures in the mid- and high-dose groups; reduced food consumption and reduced weight gain were also noted in both the mid- and high-dose groups. Fetal body weights per litter were reduced in all dose groups. Exposure to 10000 ppm also resulted in failure of implantation, fully resorbed litters, increased resportions per litter and

increased incidence of cervical ribs.

Reference Nelson, B.K., Brightwell, W.S., MacKenzie-Taylor, D.R., Khan, A.,

Burg, J.R., and Weigel, W.W. (1988) Fd. Chem. Toxicol. 26: 247-

254.

5.10 Other Relevant Information

Type METABOLISM

Remark Numerous studies on the absoption, distribution, metabolism and

excretion of IPA have been performed. These indicate that IPA is readily absorbed in animals and man through the lungs, skin and GI tract. There is evidence for a delay in absorption through the GI tract at high dose levels and an extension in half life suggesting limited metabolic capability. IPA is rapidly distributed throughout the body and has been shown to cross the blood/brain barrier. Elimination from

the blood follows first order kinetics. Approximately 64 - 84% of an intravenous dose has been shown to be oxidised to acetone in rabbit. Elimination of IPA is retarded by ethanol and it has been shown that IPA is a poorer substrate for alcohol dehydrogenase than ethanol. Excretion occurs mainly through the expired air either as unchanged IPA or as acetone. Quantities of acetone and IPA are excreted in the urine together with the glucuronide conjugate of IPA. There is evidence in man that sulphonation may occur.

World Health Organisation, Environmental Health Criteria 103, 2-Propanol, WHO, Geneva, 1990.

In rats, a similar relationship was established for excretion of acetone and IPA in expired air following a single IP injection of 0.1 ml IPA. 2% of the administered dose was excreted in the expired air as IPA and 38.8% as acetone.

Teramoto, K., Horiguchi, S., Adachi, M., Wakitani, F., Fukui, M., Osaka City Med. J., 33 (2): 153 - 160, 1987.

IPA has been shown to be readily absorbed and distributed through rat tissue following intravenous, gavage or inhalation routes of exposure. IPA is mainly metabolised to acetone by a saturable pathway. The main route of excretion is via the expired air as acetone, CO2 and unchanged IPA. Over 80% of a radiolabel was excreted in expired air following these routes of exposure. There were no substantial differences between males and females.

Slauter, R. W. et al. Disposition and Pharmacokinetics of Isopropanol in F-344 Rats and B6C3F1 Mice. (Abstract 809) The Toxicologist Vol 12 (1) 1992.

absorption. metabolism. disposition. isopropanol (IPA) were studied in male and female rats and mice. Animals were exposed by i.v. (300 mg/kg) and inhalation (500 and 5000 ppm for 6 hours) routes; additionally IPA was given by gavage to rats only in single and multiple 300 and 3000 mg/kg doses. In the rat approximately 81-89% of the administered dose was exhaled (as acetone, CO2, and unmetabolized IPA); approximately 76% of the dose in mice was exhaled after iv bolus but 92% was exhaled following inhalation. Approximately 3-8% of the administered dose was excreted in urine as IPA, acetone, and a metabolite tentatively identified as isopropyl glucuronic acid. Small amounts of radiolabel were found in feces and in the carcass. There were no major differences in the rates or routes of excretion observed either between sexes or between routes of administration. Additionally, repeated exposure had no effect on excretion. However, both the route of administration and the exposure or dose level influenced the form in which material was exhaled. Following exposure to 5000 ppm, a greater percentage of unmetabolized IPA was recovered in the expired air than following exposure to 500 ppm, implying saturation of metabolism".

Slauter, R.W. et al, (1994). Fundam. Appl. Toxicol. 23, 407-420.

PHARMACOKINETICS

Male and female rats and mice were exposed to 300 mg/kg isopropanol intravenously, and to 500 and 5000 ppm isopropanol by

Reference

Remark

Reference

Remark

Reference

Remark

Reference

Type Remark

inhalation for 6 hours. Additionally, isopropanol was given to rats by gavage (300 and 3000 mg/kg) in single and multiple doses. Exhalation was the major route of excretion with acetone as the major metabolite along with unmetabolized isopropanol and CO2. Urinary excretion accounted for 5-8% of the administered dose which included isopropanol, acetone, and the glucuronide conjugate of isopropanol. A small amount was excreted in the feces. Excretion and distribution patterns were similar by all routes of administration. High doses exceeded the metabolic threshold as evidenced by greater proportion of isopropanol excretion. There was no bioaccumulation, and distribution was similar for single and repeated administration. No major differences in absorption, distribution, metabolism, and excretion between species were observed.

Slauter, R.W., Coleman, D.P., Gaudette, N.F., McKee, R.H., Masten, L.W., Gardiner, T.H., Marino, D.J., Tyler, T.R., and Jeffcoat, A.R. (1992) Toxicologist 12: 219.

Following inhalation exposure of rats to isopropanol for 4 hours, blood levels of isopropanol and its metabolite acetone were directly related to airborne concentrations in the range of 500 and 8000 ppm. Following inhalation, the acetone/isopropanol ratio in blood decreased indicating saturation of the oxidative metabolic pathway above 4000 ppm.

Laham, S., Potvin, M., Schrader, K., and Marino, I. (1980) Drug Chem. Toxicol. 3: 343-360.

NEUROTOXICITY

Groups of male and female rats (Fischer 344) received a single inhalation exposure of IPA at dose levels of 0, 500, 1500, 5000 or 10000 ppm for 6 hours. A spectrum of behavioral effects indicative of narcosis, defined as a generalized loss of neuromotor and reflex function, was observed in animals of the 10000 ppm group, and to a lesser extent in the 5000 ppm animals. Recovery from these effects was observed by 24 hours for the 10000 ppm animals, and by 6 hours for the 5000 ppm animals. A concentration-dependent decrease in motor activity was observed for males exposed to > 1500 ppm and for females exposed to > 5000 ppm. The results showed that exposure of rats to isopropanol vapor produces transient, concentration-related narcosis and/or CNS sedation. A NOEL for clinical signs of intoxication (narcosis and neurobehavioural function) was 1500 ppm. A NOEL for neurobehavioural effects was 500 ppm.

- (1) Gill, M.W., Burleigh-Flayer, H.D., Marino, D.J., Masten, L.W., McKee, R.H., Tyler, T.R. and Gardiner, T. Isopropanol Single Exposure Vapor Unhalation Neurotoxicity Sttudy in Rats. (Abstract 862) The Toxicologist Vol. 12 (1), 1992.
- (2) Gill, M.W., Burleigh-Flayer, H.D., Strother, D.E., Masten, L.W., McKee, R.H., Tyler, T.R. and Gardiner, T.H. Isopropanol: Acute Vapor Inhalation neuroticity Study in Rats. J. Appl. Toxicol. 15 (2) 77-84, 1995.

A vapor inhalation study was performed on isopropanol on F-344 rats and CD-1 Mice. Exposures were 0, 100, 500, 1500, or 5000 ppm for

Reference

Remark

Reference

Type Remark

Reference

Remark

6 hours/day, 5 days per week, for 13 weeks. Effects of narcosis were observed only during exposure to 5000 ppm. Neurobehavioural evaluations indicated no changes in the functional observation battery; however, increased motor activity was noted in female rats of the 5000 ppm group at Weeks 9 and 13. Neuropathological examination revelaed no exposure-related lesions in the nervous system.

Reference

Burleigh-Flayer, H.D., Gill, M.W., Marino, D.J., Masten, L.W., McKee, R.H., Tyler, T.R., and Gardiner, T.H. (Abstract No. 1057) Toxicologist Vol. 12 (1) 1992.

Remark

See M. Gill's review of case studies for an overview of neurotixicty testing of isopropsanol.

Reference

Gill, M.W., "Case Studies in Industrial Neurotoxicology" In Neurobehavioral Toxicity: Analysis and Interpretation, B. Weiss and J. O'Donoghue (Eds.) Raven Press, New York, 1994.

Remark

See Section 5.4 for details of 12 week study in rats receiving isopropanol in drinking water. No indication of neurotoxicity, as assayed as GFAP in the dorsal hippocampal region, was detected.

Remark

Jcl-Wistar rats were exposed to 1000 and 8000 ppm isopropanol 8 hours/day, 5 days/week for 20 weeks. Sensory nerve conduction velocity was reported to have been affected in the 8000 ppm dose group.

Reference

Teramoto, K., Wakitani, F., Horiguch, S., Jo, T., Yamamoto, T., Misutake, H., and Nakaseko, H. (1993) Environ.Res 62:148

Type Remark

NERVE CONDUCTION STUDY

Jcl-Wistar rats were exposed to 1000 and 8000 ppm isopropanol 8 hours/day, 5 days/week for 20 weeks. Sensory nerve conduction velocity was reported to have been affected in the 8000 ppm dose group.

Reference

Teramoto, K., Wakitani, F., Horiguchi, S., Jo, T., Yamamoto, T., Mitsutake, H., and Nakaseko, H. (1993) Environ. Res. 62: 148.

Type Remark

IMMUNOTOXICITY

It has been suggested that IPA may interfere with the normal inflammatory response. Inhibition of histamine induced increase in vascular permeability has been demonstrated as has experimentally induced plasma exudation into the pleuralcavity. Administration of antiinflammatory drugs however increased the pleural inflammatory reaction.

Reference

Kasuga, F., Inoue, S., Asano, T., Kumagi, S., Fd. Cos. Tox. 30 (7): 631-634, 1992.

Type Remark

IMMUNOHISTOCHEMISTRY

Kidney section prepared from paraffin blocks of kidney tissues obtained following a 9- day vapor inhalation study were examined for immunohistochemical staining with alpha2u-globulin. The section examined included 5/sex from the control and 5000 ppm exposure groups and 5 males from a 10,000 ppm exposure group. There were no substantive differences noted between the exposure groups for the male rats in either intensity or distribution of the alpha2u-globulin

reactivity. The females did not have any alpha2u-globulin positive

reaction in the kidney sections.

Reference Fowler, E.H. and Martin, C. Immunohistochemical Evaluation of Alpha2u-Globulin in Kidneys of Fischer 344 Rats from the Isopropanol Nine-Day Vapor Inhalation Study. BRRC Report No.

93N1229 dated April 29, 1994.

Type SENSORY IRRITATION

Remark Isopropanol produced sensory irritation in mice, with RD50 values of

 $5000\ ppm$ and $17693\ ppm.$

Reference (1) DeCeaurriz et al, 1981.

(2) Kane et al, 1980.

Remark Guinea pigs exposed by inhalation to 400 ppm isopropanol for 24

hours showed epithelial changes in the respiratory mucosa and a decline in tracheal ciliary activity which returned to normal within 2 weeks following exposure. Recovery did not occur within 2 weeks in

animals exposed to 5500 ppm for 24 hours.

Reference (1) Ohashi et al, 1987a.

(2) Ohashi et al, 1987b.

Type DERMAL ABSORPTION

Remark Dermal absorption and Pharmacokinetics of Isopropanol in the Male

and Female F-344 Rat.

Remark Chemical Manufacturers Association Isopropanol Panel conducted a

dermal absorption study in which the rats were exposed dermally for a period of 4 hours. First order elimination half-lives for IPA was 0.8 hours and 2.6 hours for acetone. ¹⁴C-IPA studies showed excellent agreement with dermal absorption rates found via the analytical method. The study showed that IPA is rapidly absorbed through rat

skin when applied under occlusion.

Reference Boatman, R.J., Perry, L.G., Fiorica, L.A., English, J.C., Kapp, R.W.,

Jr., Bevan, C., Tyler, T.R., Banton, M.I. and Wright, G.A.. Dermal Absorption and PharmacoKinetics of Isopropanol in the Male and Female F-344 Rats. The Toxicologist (Abstract) Vol 30, No.1, (2) No.

210 (1996).

Type HAZARD IDENTIFICATION

Remark US EPA, Draft Final RM1 Risk Assessment of Isopropanol, OPPT,

1996. TSCA Public Docket Number AR-141.

Type OTHER

Remark Deaths occurred in 6 of 10 rats following aspiration of 0.2 ml of

100% isopropanol and in 1 of 10 rats following aspiration of 70%

isopropanol.

Reference Gerarde, H.W., and Ahlstrom, D.B. (1986) Arch. Environ. Health 13:

457-461.

Remark Acute or chronic treatment of rats with isopropanol caused a

significant increase in hepatic and renal cytochrome P450 content.

Reference (1) Uneng, T.H., Moore, L., Elves, R.G., and Alvares, A.P. (1983)

Toxicol. Appl. Pharmacol. 71: 204-214.

(2) Zahlsen, K., Aasted, K., and Nilsen, O.G. (1985) Toxicology 34: 57-66.

Remark Acute oral pretreatment of rats with isopropanol resulted in a dose-

related potentiation of carbon tetrachloride hepatotoxicity.

Reference Plaa, G.L., Hewitt, W.R., du Souich, R., Caille, G., and Lock, S.

(1982) J. Toxicol. Environ. Health 9: 235-250.

Remark The cutaneous penetration of IPA was evaluated usinf excised, full-

thickness skin from male and female F-344 rats, B6C3F1 mice and female humans in a flow-through in vitro skin penetration apparatus. Steady state penetration rates were 1.3, 1.88 and 1.65 mg/cm2 for rat, mouse and human, respectively. Permeability constants were similar among the three tissue types while over a 6 hour treatment period, more IPA penetrated the mouse skin (11.6%) than through rat (6.4%) or through human (7.1%). The rat skin was determined to be a good human skin model while the mouse skin was found to be the most

permeable.

Reference Morris, E.M., Sun, J.D., Frantz, S.W., Beskitt, J.L, Bevan, C.,

Gardiner, T., Kapp, R., Tyler, T. and Wright, G. (Abstract No. 1727)

The Toxicologist Vol. 15 (1), 1995.

5.11 Experience with Human Exposure

Type ODOR THRESHOLD

Remark The air odor threshold for isopropanol has been reported as 22 ppm. Reference Amoore, J.E., and Hautala, E. (1983) J. Appl. Toxicol. 3: 272-290.

Type ACCIDENTAL EXPOSURE

Intoxications have been reported following ingestion. rectal administration and. in children, following inhalation and absorption. Signs of intoxication are CNS depression, leading to coma, respiratory arrest and death. GI effects and hypothermia may occur. Cardiac effects include severe hypotension, shock and cardiac arrest with tachycardia as a secondary effect. The lowest dose reported to be life threatening was 170 ml in an 18 month old child. Acetone can be detected in the blood, breath and urine after intoxication with IPA, but acidosis does not usually occur. Occasional reports of allergic reaction to IPA have appeared in the literature.

World Health Organisation, Environmental Health Criteria 103, 2-

Propanol WHO, Geneva, 1990.

Type VOLUNTEER STUDIES

Remark

2.6 or 6.4 mg/kg/day IPA for 6 weeks was well tolerated by human male volunteers, there being no adverse effects on hematology, blood chemistry, urinalysis or ophthalmoscopy. Application of 0.5 ml

undiluted IPA in an open patch test did not result in skin irritation. 10 minute covered patches produced transient erythema following immersion in water. Premature infants however were found to be

particularly susceptible.

Reference World Health Organisation, Environmental Health Criteria 103, 2-

Propanol, WHO, Geneva, 1990.

Remark

Reference

Remark

Although isopropanol was found to produce little irritation when tested on the skin of six human subjects, there have been reports of dermal irritation and/or skin sensitization. With the exception of three reports, the positive reactions were observed on patch testing of patients with a contact dermatitis due to ethanol. These patients also had a positive reaction to ethanol.

Reference

Lington, A.W., and Bevan, C. (1994) Patty's Industrial Hygiene and Toxicology, 4th Edition.

Remark

Exposure to 400 ppm isopropanol vapors for 3 to 5 min caused mild irritation of the eyes, nose and throat of human volunteers. At 800 ppm, the effects were not severe but the majority felt the atmosphere was unsuitable.

Reference

Nelson, K.W., Ege, J.F., Jr., Ross, M., Woodman, L.E., and Silverman, L. (1943) J. Ind. Hyg. Toxicol. 9:93-97

Remark

The production of acetone following IPA ingestion in man was demonstrated in a study involving three human volunteers. There was a correlation between blood levels of IPA and acetone with the initial peak of IPA recorded half an hour after dosing and reducing as the acetone levels increased over a 24 hour period.

Reference

Lacouture, P.G., Heldreth, D.D., Vet. Hum. Toxicol., 29 (6): 486, 1987.

Type Remark

EPIDEMIOLOGY

There is epidemiological evidence indicating that the manufacture of isopropanol by the strong-acid process is associated with an excess of upper respiratory tract cancer in workers. IARC has concluded that there is sufficient evidence for carcinogenicity to humans in the manufacture of isopropanol by this process. Although the use of this strong-acid process has raised carcinogenicity concerns, these are related to sulfuric acid and to by-products such as dialkyl sulfates that are formed during this manufacturing process, not to isopropanol itself.

Reference

- (1) World Health Organisation, Environmental Health Criteria 103, 2-Propanol, WHO, Geneva, 1990.
- (2) International Agency for Research on Cancer, Monographs on the evaluation of carcinogenic risk to man, No. 15, IARC, Lyon, France, 1977 and update, 1987.
- (3) Weil, C.S., Smyth, H.F., Jr., and Nale, T.W. (1952) Arch. Ind. Hyg. Occup. Med. 5:535-547.
- (4) Lynch, J., Hannis, N.M., Bird, M.G., Murray, K.J., and Walsh, J.P. (1979) J. Occup. Med. 21:333341.
- (5) Soskolne, C.L., Zeighami, E.A., Hanis, N.M., Kupper, L.L., Herrman, N., Amsel, J., Mausner, J.S., and Stellman, J.S. (1984) Am. J. Epidemiol. 120:358-369

A single study was reported showing that IPA was associated with a higher risk of breast cancer. However, there was a combined exposure with freon and solder flux and aspects of lifestyle (such as smoking

habits and alcohol consumption) were not taken into account.

Spiritas, R., Stewart, P.A., Lee, J.S., Marano, D.E., Forbes, C.D., Grauman, D.J., Pettigrew, H.M., Blair, A., Hoover, R.N., Cohen, J.L., Brit. J. Ind. Med., 48: 515 - 530, 1991.

Remark

Reference

Remark In a population based referent study, the occupational experince of 86

men with oral or oropharyngeal cancer and 373 referents was analyzed with respect to employment and exposure to 16 chemicals, including IPA, as estimated by a job exposure matrix. There was no evidence of association between IPA exposure and oral or

oropharangeal exposure in this study.

Reference Merletti, F., Boffetta, P., Ferro, G., Pisani, P., Terrachini, B., Scand.

J. Work. Env. Hlth., 17: 248 - 254, 1991.

Remark The lung uptake was studied in 12 printing workers exposed to

workplace levels in the range of 8 to 647 mg/m3 (3.3 to 264 ppm) isopropanol. The alveolar isopropanol concentration was highly correlated with the exposure level at any time of exposure. Acetone, but not isopropanol, was detected in the blood or urine. The acetone concentration ranged between 0.76 and 15.6 mg/liter in the blood and between 3 and 93 mg/m3 in the alveolar air. The acetone levels in the alveolar air and blood increased with the increasing exposure period and were linearly related to the alveolar isopropanol levels. Elimination of acetone was mainly via the lungs, varying from 10.7 to

39.8% of the uptake, and was inversely related to the exposure level.

Brugnone, F., Perbellini, L., Apostoli, P., Bellomi, M., and Caretta,

D. (1983) Br. J. Ind. Med. 40:160-168.

6. REFERENCES

Reference

• Atkinson, R. A structure-activity relationship for the estimation of rate constants Amoore, J.E., and Hautala, E. (1983) J. Appl. Toxicol. 3: 272-290.

- Atkinson, R. (1985): Chem Rev 85, pp. 69-75, 135-153, 182-201.
- for the gas=phase reactions of OH radicals with organic compounds. Int J Chem Kinet 19, 799-828 (1987).
- Atkinson, R. (1990): J Phys Chem Ref Data (1992), 21(6):1125-1568
- Bates, H.K., McKee, R.H., Bieler, G.S., Gardiner, T.H., Gill, M.W., Strother, D.E., and Masten, L.W. (1994) Fundam. Appl. Toxicol. 22:152-158.
- Bevan, C., Tyler, T.R., Gardiner, T.H., Kapp, R.W., Jr., Andrews, L. and Beyer, B.K. Journal of Applied Toxicology, Vol. 15(2) pp. 117-123 (1995).
- Blackman, R.A.A., Toxicity of Oil-Sinking Agents, Mar. Pollut. Bull., 5, 1974, 116-118.
- Boatman, R.J., Perry, L.G., Fiorica, L.A., English, J.C., Kapp, R.W., Jr., Bevan, C., Tyler, T.R., Banton, M.I. and Wright, G.A.. Dermal Absorption and PharmacoKinetics of Isopropanol in the Male and Female F-344 Rats. The Toxicologist (Abstract) Vol 30, No.1, (2) No. 210 (1996).
- Bridie, A.L., Wolff, C.J.M. & Winter, M., Water Research, 13, 1979, 627-630.
- Bridie, A.L., Wolff, C.J.M., & Winter, M. BOD and COD of some Petrochemicals, Water Research, 13, 1979, p.627- 630.
- Bringmann, G. & Kuehn, R., Comparison of the Toxicity Thresholds of Water Pollutants to Bacteria, Algae and Protozoa in the Cell Multiplication Inhibition Test, Water Research, 14, 1980, 231-241.
- Bringmann, G. et al. (1980): Z. Wasser Abwasser Forsch. 10, 87-98.
- Bringmann, G. & Kuehn, R., Grenzwerte der Schadwirkung waserfefaehdender Stoffe gegen Blaualgen (Microcystis aeruginosa) und Gruenalgen (Scenedesmus quadricauda) im Zellvermehrungshemmtest, Vom Wasser, 50, 1978, 45-60.

• Bringmann, G. & Kuehn, R., Results of the Damaging Effect of Water Pollutants on Daphnia magna, Z. Wasser Abwasser Forsch., 10(5), 1977, 161-166.

- British Industrial Biological Research Assoc., Report 0570/3/86.
- British Industrial Biological Research Assoc., Report 0570/2/86.
- Brugnone, F., Perbellini, L., Apostoli, P., Bellomi, M., and Caretta, D. (1983) Br. J. Ind. Med. 40:160-168.
- Burleigh-Flayer, H.D., Gill, M.W., Marino, D.J., Masten, L.W., McKee, R.H., Tyler, T.R., and Gardiner, T.H. (1992) Toxicologist 12: 227
- Burleigh-Flayer, H., Bevan, C., Gardiner, T., Garman, R., Kapp, R., Neptun, D., Tyler, T. and Wright, G. Isopropanol Vapor Inhalation Oncogenicity Study in CD-1 Mice.(Abstract No. 1219) The Toxicologist, Vol. 14 (1), 1994.
- Burleigh-Flayer, H.D., et al. (1994). Fundam. Appl. Toxicol. 23, 421-428.
- David, J. & Bocquet, C. Compared Toxicities of Different Alcohols for Two Drosophila Sibling Species. Comp. Biochem. Physiol., 54C, 1976, 71-74.
- DeCeaurriz, J.C., Micillino, J.C., Bonnet, P., and Guenier, J.P. (1981) Toxicol. Lett. 9: 137-144..
- Dillingham, E.O. et al., J. Pharm. Sci., 62, 1973, 22. In Pomona Data File on Log P and related parameters. DGEP Review of literature data on 2-propanol, Leidschendam, Netherlands, Directorate-General of Environmental Protection, Ministry of Housing, Physical Planning and Environment. (1987)
- Exxon Biomedical Sciences Inc. (1986) Ocular Irritation Study in Rabbits (Isopropanol). EBSI Document No. 86MRLZ72.
- Fowler, E.H. and Martin, C. Immunohistochemical Evaluation of Alpha2u-Globulin in Kidneys of Fischer 344 Rats from the Isopropanol Nine-Day Vapor Inhalation Study.BRRC Report No. 93N1229 dated April 29, 1994.
- Gallo, M.A., Oser, B.I., Cox, G.E., and Bailey, D.E. (1977) Toxicol. Appl. Pharmacol. 41:35.
- Garman, R. Bevan, C., Burleigh-Flayer, H., Gardiner, T., Kapp, R., Neptun, D., Tyler, T. and Wright, G. Isopropanol (IPA) Vapor Inhalation Oncogenicity Study in Fisher 344 Rats.(Abstract No. 979) The Toxicologist, Vol. 15 (1), 1995.
- Gerarde, H.W., and Ahlstrom, D.B. (1986) Arch. Environ. Health 13: 457-461.
- Gill, M.W., Burleigh-Flayer, H.D., Marino, D.J., Masten, L.W., McKee, R.H., Tyler, T.R. and Gardiner, T. Isopropanol Single Exposure Vapor Inhalation Neurotoxicity Sttudy in Rats. (Abstract 862) The Toxicologist Vol. 12 (1), 1992.
- Gill, M., Burleigh-Flayer, H., Bevan. C., Gardiner, T., Kapp, R., Isopropanol Ninety-Day Vapor Inhalation Neurotoxicity Study in Female F-344 Rats. (Abstract) The Toxicologist, Vol. 14 (1), 1994.
- Gill, M.W., Burleigh-Flayer, H.D., Strother, D.E., Masten, LW., McKee, R.H., Tyler, T.R. and Gardiner, T.H. Isopropanol: Acute Vapor Inhalation neuroticity Study in Rats. J. Appl. Toxicol. 15 (2) 77-84, 1995.
- Griffith, J.F., Nixon, G.A., Bruce, R.D., Reer, P.J., Bannan, E.A, (1980), Toxicol. Appl. Pharmacol. 55: 501-513.
- Guesten, H et al. (1984): J Atmos Chem 2, 83-93.
- Guseinov, V.G., (1985), Gig. Tr. Prof. Zabol. (7): 60-62.
- Harris, S.B. (1995). A review of the EPA comments regarding the study entitled "Multigeneration rat reproduction study with isopropanol". Report prepared for the Chemical Manufacturers Association Isopropanol Panel.
- Howard, P.H., Handbook of Environmental Fate and Exposure Data for Organic Chemicals, Lewis Publishers, Inc., Chelsea Michigan USA, 1990, p.304-309.

 Huels-Bericht DL 106, 1988 (unveroeeffentlicht). Idem, Shell Group Research Report, AMGR.0224.74

- International Agency for Research on Cancer, Monographs on the evaluation of carcinogenic risk to man, No. 15, IARC, Lyon, France, 1977 and update, 1987.
- IPCS International Programme on Chemical Safety Environmental Health Criteria 103 2- Propanol, World Health Organization (1990).
- Jones, C.J., McGugan, P.J. (1977/78): J Hazard Mater. 2, 235-251.
- Kane, L.E., Dombroske, R., and Alerie, Y. (1980) Am. Ind. Hyg. Assoc. J. 41: 451-455.
- Kapp, R.W. et al. (1993). Environ. Mol. Mutagen., 22: 93-100.
- Kasuga, F., Inoue, S., Asano, T., Kumagi, S., Fd. Cos. Tox. 30(7):631-634, 1992.
- Kimura, E.T., Ebert, D.M., Dodge, P.W., (1971), Toxicol. Appl. Pharmacol., 19: 699-703.
- Kloepffer, W. et al. (1988): Ecotoxicol Environ Safety 15, 298-319.
- Laham, S., Potvin, M., Schrader, K., Marino, I. (1979), Drug. Chem. Toxicol. 3: 343-360.
- Laham, S., Potvin, M., Schrader, K., and Marino, I. (1980) Drug Chem. Toxicol. 3: 343-360.
- Lacouture, P.G., Heldreth, D.D, Vet. Hum. Toxicol., 29 (6):486, 1987
- Lehman, A.J., Chase, H.F., (1944) J. Lab. Clin. Med. 29: 561-567.
- Lington, A.W., and Bevan, C. (1994) Patty's Industrial Hygiene and Toxicology, 4th Edition.
- Lloyd, A.C. et al. (1976): Chem Phys. Lett 42, 205-209.
- Lyman, W.J. et al., Chemical Property Estimation Methods. Environmental behaviour of organic compounds. McGraw-Hill Book Company, New York (USA), 1982, Chpt. 15.
- Lynch, J., Hannis, N.M., Bird, M.G., Murray, K.J., and Walsh, J.P. (1979) J. Occup. Med. 21:333341.
- Mackay, D. & Paterson, S., Calculating Fugacity, Environm. Sci. Technol,. 15(9), 1981, 1006-1014
- Marzulli, F.N., Ruggles, D.I., (1973), J. Assoc. Off. Anal. Chem., 56: 905-914.
- Merletti, F., Boffetta, P., Ferro, G., Pisani, P., Terrachini, B., Scand. J. Work. Env. Hlth., 17:248-254, 1991.
- Morgan, R.L., Sorenson, S.S., Castles, T.R., (1987), Food. Chem. Toxicol., 25: 609-613.
- Morris, E.M., Sun, J.D., Frantz, S.W., Beskitt, J.L, Bevan, C., Gardiner, T., Kapp, R., Tyler, T. and Wright, G. (Abstract No. 1727) The Toxicologist Vol. 15 (1), 1995.
- Munch, J.C., (1972), Ind. Med. 41: 31-33.
- Nakaseko, H., Teramoto, K., Horiguchi, S., Wakitani, F., Yamanoto, T., Adachi, M., Tanaka, H., Hozu, S., Jpn. J. Ind. Hlth. 33(3): 200-201, 1991.
- Nelson, K.W., Ege, J.F., Jr., Ross, M., Woodman, L.E., and Silverman L. (1943) J. Ind. Hyg. Toxicol. 9:93-97.
- Nelson, B.K., Brightwell, W.S., MacKenzie-Taylor, D.R., Khan, A., Burg, J.R., and Weigel, W.W. (1988) Fd. Chem. Toxicol. 26: 247-254.
- Nixon, G.A., Tyson, C.A., Wertz, W.C., (1975), Toxicol. Appl. Pharmacol. 31: 481-490.
- Ohashi, Y., Nakai, Y., Ikeoka, H., Koshimo, H., Esaki, Y., Horiguchi, S., Teramoto, K., and Nakaseko, H. (1987a) J. Appl. Toxicol. 8: 67-71.
- Ohashi, Y., Nakai, Y., Ikeoka, H., Koshimo, H., Esaki, Y., Horiguchi, S., Teramoto, K., and Nakaseko, H. (1987b) Arch. Toxicol. 61: 12-20.
- Overend, R., Paraskevopoulos, G. (1978): J phys Chem 82, 1329-1333...
- P&G, Unpublished data. For further information, contact Dr. C. Lally, Proctor & Gamble, Brussels. Phone 32-2-456-3056, Fax 32-2-456-2845.
- Pilegaard, K. and Ladefoged, O. (1993) In Vivo 7:325-330.
- Plaa, G.L., Hewitt, W.R., du Souich, R., Caille, G., and Lock, S. (1982) J. Toxicol. Environ. Health 9: 235-250.

• Price, K.S., Waggy, G.T. & Conway, R.A., Brine Shrip Bioassay and Seawater BOD of Petrochemicals, J. Water Pollut. Contr. Fed., 46, 1974, 63-77.

- Reynolds, T. An Anomalous Effect of Isopropanol on lettuce germination Plant Sci. Lett., 15, 1977, 25-28.
- Shell Chemicals data sheet IS 3.2.4 dated June 1992.
- Shipp A.M., Allen B.C., Van Landingham C., Gentry P.R. and Crump K.S. (1996). Quantitative dose-response analysis Application of the benchmark method to the multigeneration rat reproduction study for isopropanol. Report prepared for the Chemical Manufacturers Association Isopropanol Panel.
- Slauter, R.W., Coleman, D.P., Gaudette, N.F., McKee, R.H., Masten, L.W., Gardiner, T.H., Strother, D.E., Tyler, T.R. and Jeffcoat, A.R. (1994). Disposition and Pharmacokinetics of Isopropanol in F-344 Rats and B6C3F1 Mice, Fundam. Appl. Toxicol. 23, 407-420.
- Slauter, R.W., Coleman, D.P., Gaudette, N.F., McKee, R.H., Masten, L.W., Gardiner, T.H., Marino, D.J., Tyler, T.R., and Jeffcoat, A.R. (1992) Toxicologist 12:219.
- Slauter, R. W. et al. Disposition and Pharmacokinetics of Isopropanol in F-344 Rats and B6C3F1 Mice. (Abstract 809) The Toxicologist Vol 12 (1) 1992.
- Soskolne, C.L., Zeighami, E.A., Hanis, N.M., Kupper, L.L., Herrman, N., Amsel, J., Mausner, J.S., and Stellman, J.S. (1984) Am. J. Epidemiol. (120:358-369.
- Smyth, H.F., Carpenter, C.P., (1948), J. Ind. Hyg. Toxicol., 30: 63-70.
- Spiritas, R., Sewart, P.A., Lee, J.S., Marano, D.E., Forbes, C.D., Grauman, D.J., Pettigrew, H.M., Blair, A., Hoover, R.M., Cohen, J.L., Brit. J. Ind. Med., 48:515-530, 1991.
- Teramoto, K., Horiguchi, S., Adachi, M., Wakitani, F., Fukui, M., Osaka City Med. J., 33(2): 153-160, 1987.
- Teramoto, K., Wakitani, F., Horiguch, S., Jo, T., Yamamoto, T., Misutake, H., and Nakaseko, H. (1993) Environ. Res. 62:148.
- Tichy, M., Trcka, V., Roth, Z., Krivucova, M., (1985), Environ. Health. Perspect., 61: 321-328.
- 1995-1996 Threshold Limit Values (TLVs) for Chemical Substances and Physical Agents and Biological Exposure Indices (BEIs). American Conference of Governmental Industrial Hygienists (ACGIH), Cinncinati, OH (1995).
- Tyl, R.W., Masten, L.W., Marr, M.C. Myers, C.B., Slauter, R.W., Gardiner, T.H., Strother, D.E., McKee, R.H., and Tyler, T.R. (1994) Fundam. Appl. Toxicol. 22:139-151.
- Tyl R.W. (1996). February 12, 1996 Letter to the Chemical Manufacturers Association Isopropanol Panel.
- Uneng, T.H., Moore, L., Elves, R.G., and Alvares, A.P. (1983) Toxicol. Appl. Pharmacol. 71: 204-214.
- U.S. EPA, Draft Final RM1 Risk Assessment of Isopropanol, OPPT, 1996. TSCA Public Docket AR-141.
- U.S. EPA (U.S. Environmental Protection Agency). (1992). Review of Section 4 Data A two Generation Reproductive Toxicity Study in Rats with Isopropanol.
- US NIOSH (1976), Cincinnati, Ohio, US National Institute of Occupational Safety and Health, US Dept. of Health, Education and Welfare, Public Health Services, Centre for Disease Control (DHEW publication no. (NIOSH) 76-142).
- Veith, G.D., Call, D.J. & Brooke, L.T., Estimating the Acute Toxicity of Narcotic Industrial Chemicals to Fathead Minnows. In: Bishop, W.E., Cardwell, R.D. & Heidolph, B.B. Eds. Aquatic Toxicology and Hazard Assessment: 6th Symp., ASTM STP 802, Philadelphia (USA), 1983, 90-97.
- Von der Hude, W., Scheutwinkel, M., Gramlich, U., Fissler, B., Busler, A., In Vitro Environ. Mutagen., (1987), 9:401-410.
- Wallington, T.J. et al. (1987): Int J Chem Kinet 19, 243-249.

• Weil, C.S., Smyth, H.F., Nale, T.W., Arch. Ind. Hyg. Occup. Med. (1952), 5:535-547.

- World Petrochemicals, SRI International, 1996.
- Zahlsen, K., Aasted, K., and Nilsen, O.G. (1985) Toxicology 34: 57-66.

• Zeiger, E. et al, Env. Mol. Mut., 19 (Suppl.21):2-141, 1992.

OECD SIDS			2-PROPANOL
EXTRACT FROM	IRPTC	LEGAL	FILES

```
rn: 6729
File: 17.01 LEGAL
    systematic name: 2-Propanol
    common name :2-propanol
    reported name :ISOPROPYL ALCOHOL
          : b / .
                                 rtecs no :NT8050000 type : REC
               :67-63-0
     _____
    |subject|specification|descriptor|
    |-----
    AIR OCC TLV
    ______
    TWA: 980MG/M3 (400PPM) STEL: 1225MG/M3 (500PPM)
    entry date: MCH 1985
    {\tt original} \; : \; {\tt ILO} \;\; , \;\; , \;\; , \;\; , \;\;
    amendment: AOHGN*, APPROVED OCCUPATIONAL HEALTH GUIDE THRESHOLD LIMIT
             VALUES, , , , 1983
                              *****
File: 17.01 LEGAL
                                                      rn: 15004
    systematic name: 2-Propanol
    common name :2-propanol
    reported name :ISOPROPYL ALCOHOL
                                rtecs no
type
   cas no :67-63-0
                                             :NT8050000
              : BEL
                                            : REC
    area
    ______
    |subject|specification|descriptor|
    |-----
    AIR OCC
                    TLV
     _____
    TWA: 980MG/M3 (400PPM); STEL: 1225MG/M3 (500 PPM). SKIN ABSORPTION.
    entry date: JUL 1987
    original : ILO , , , ,
    amendment: TLVBE*, THRESHOLD LIMIT VALUES(TOLERABLE LIMIT VALUES), , , ,
            1984
                              *****
File: 17.01 LEGAL
                                                      rn: 15640
    systematic name: 2-Propanol
    common name :2-propanol
    reported name :ISOPROPYL ALCOHOL
                                rtecs no :NT&
               :67-63-0
                                             :NT8050000
    cas no
              : FIN
    area
     ______
    |subject|specification|descriptor|
    |-----|
    AIR OCC MPC
     -----
    TWA: 500MG/M3 (200PPM) STEL: 625MG/M3 (250PPM) (SKIN) (APPLIES TO ALL
    ISOMERS)
    entry date: MAY 1989
    original : ILO , , , ,
    amendment: APWFI*, HTP-ARVOT (LIST OF LIMIT VALUES FOR CONCENTRATIONS OF
             TOXIC SUBSTANCES KNOWN TO BE HARMFUL TO HEALTH), 25 , , 10 ,
```

1988

```
File: 17.01 LEGAL
                                                  rn: 16325
   systematic name: 2-Propanol
   rtecs no :NT8050000 type : REC
         :67-63-0
: ITA
   cas no
   area
    |subject|specification|descriptor|
    |-----
    AIR OCC TLV
    500MG/M3 (200PPM) (SKIN ABSORPTION)
    entry date: MCH 1985
   original : ILO , , , ,
   amendment: TLVIT*, VALORI LIMITE PONDERATI(APPRAISED LIMIT VALUES), , ,
                            *****
File: 17.01 LEGAL
                                                 rn: 16707
   systematic name: 2-Propanol
   common name :2-propanol
   reported name :ISOPROPYL ALCOHOL
   cas no :67-63-0
                              rtecs no :NT8050000
             : NLD
                                         : REC
                               type
    _____
    |subject|specification|descriptor|
    |-----
    AIR OCC MXL
    _____
   TWA: 980MG/M3 (400PPM) (SKIN ABSORPTION)
    entry date: JUN 1987
   original : ILO , , , ,
   amendment: NMACN*, NATIONALE MAC-LIST(NATIONAL MAC-LIST), , , , 1986
                            *****
File: 17.01 LEGAL
                                                 rn: 17374
   systematic name:2-Propanol
   common name :2-propanol
   reported name :ISOPROPYL ALCOHOL
                              rtecs no :NT80
   cas no :67-63-0
                                          :NT8050000
             : ROM
   area
    _____
    |subject|specification|descriptor|
    |-----|
    AIR OCC MPC
    -----
   TWA: 400MG/M3; CLV: 600MG/M3
   entry date: MCH 1985
   original : ILO , , , , ,
   amendment: OMHRO*, ORDINANCE OF THE MINISTRY OF HEALTH, 60 , , , 1975
```

```
File: 17.01 LEGAL
                                                rn: 17824
   systematic name: 2-Propanol
   rtecs no :NT8050000
type : REG
         :67-63-0
: CHE
   cas no
   area
    |subject|specification|descriptor|
    |-----
    AIR OCC MAK
    -----
   TWA: 980MG/M3 (400PPM)
   entry date: DEC 1987
   original : ILO , , , , ,
   amendment: ZWACH*, ZULAESSIGE WERTE AM ARBEITSPLATZ(PERMITTED VALUES IN
           THE WORKPLACE), , , 1987
                           *****
File: 17.01 LEGAL
                                                rn : 18341
   systematic name: 2-Propanol
   common name :2-propanol
                             rtecs no :NT80 : REG
   reported name : ISOPROPYL ALCOHOL
   cas no :67-63-0
                                        :NT8050000
             : YUG
    _____
   |subject|specification|descriptor|
    |-----
    AIR OCC MAC
   TWA: 980MG/M3 (400PPM)
   entry date: MCH 1985
   original : ILO , , , ,
   amendment: ORYUG*, ORDINANCE, 24-3698/1 , , , 1971
                           *****
File: 17.01 LEGAL
                                                rn : 50882
   systematic name: 2-Propanol
   common name :2-propanol
   reported name :2-Propanol
   cas no :67-63-0
                              rtecs no :NT80
                                        :NT8050000
             : IMO
   area
    _____
    |subject|specification|descriptor|
    |-----|
```

This substance is presently considered to present no harm to human health, marine re sources, amenities or other legitimats uses of the sea when discharged into the sea from tank cleaning or deballasting operations (applies to n-propyl alcohol)

entry date: APR 1993

original : IMODC*, , , , 1992

File: 17.01 LEGAL rn: 100030

systematic name: 2-Propanol common name :2-propanol reported name :ISOPROPYL ALCOHOL

:67-63-0 rtecs no type cas no :NT8050000

area : ARG : REG

______ |subject|specification|descriptor| |-----AIR OCC MPC

8H-TWA: 980MG/M3 (400PPM); 15MIN-STEL: 1225MG/M3 (500PPM) (MAXIMUM 4

TIMES/DAY WITH INTERVALS OF AT LEAST 60 MINUTES).

entry date: OCT 1991 effective date: 29MAY1991

title: LIMIT VALUES FOR CHEMICAL SUBSTANCES IN THE WORKING ENVIRONMENT-RESOLUTION NO. 444/1991 OF THE MINISTRY OF WORK AND SOCIAL SECURITY (AMENDING REGULATION DECREE NO. 351/1979 UNDER LAW NO.

19587/1972: HYGIENE AND SAFETY AT WORK)

original: ARGOB*, Boletin Oficial de la Republica Argentina(Argentinian

Official Bulletin), 24170 , I , 1 , 1979

amendment: ARGOB*, Boletin Oficial de la Republica Argentina(Argentinian

Official Bulletin), 27145 , I , 4 , 1991

File: 17.01 LEGAL rn: 300124

systematic name: 2-Propanol common name :2-propanol

reported name :ISOPROPYL ALCOHOL

:67-63-0 rtecs no :NT8050000 cas no : REG

: CAN type area

|subject|specification|descriptor| |-----| AIR OCC TLV _____

TWA: 400 PPM, 980 MG/M3; STEL: 500 PPM, 1,225 MG/M3. PRESCRIBED BY THE CANADA OCCUPATIONAL SAFETY AND HEALTH REGULATIONS, UNDER THE CANADA LABOUR CODE (ADMINISTERED BYTHE DEPARTMENT OF LABOUR). THE REGULATIONS STATE THAT NO EMPLOYEE SHALL BE EXPOSED TO A CONCENTRATION OF AN AIRBORNE CHEMICAL AGENT IN EXCESS OF THE VALUE FOR THAT CHEMICAL AGENT ADOPTED BY ACGIH (AMERICAN CONFERENCE OF GOVERNMENTAL INDUSTRIAL HYGIENISTS) IN ITSPUBLICATION ENTITLED: "THRESHOLD LIMIT VALUE AND BIOLOGICAL EXPOSURE INDICES FOR 1985-86".

entry date: MCH 1991 effective date: 13MCH1986

amendment: CAGGAK, , 120 , 6 , 1105 ,

File: 17.01 LEGAL rn: 300832

systematic name: 2-Propanol

rtecs no :NT8050000 type : REG :67-63-0 : CAN cas no

area

|subject|specification|descriptor| |----+-----PACK | AGRIC | CLASS | LABEL | PESTI USE |

HISTORICAL STATUS. DATE OF LAST OCCURRENCE: 1988. CODE IAL. THE PEST CONTROL PRODUCTS ACT AND REGULATIONS ARE ADMINISTERED BY THE DEPARTMENT OF AGRICULTURE. THEY ESTABLISH A REGISTRATION, CLASSIFICATION, PACKAGING AND LABELLING SYSTEM FOR PEST CONTROL PRODUCTS. ONLY PEST CONTROL PRODUCTS THAT ARE CURRENTLYREGISTERED WITH THE DEPARTMENT OF AGRICULTUREAND PRODUCTS THAT HAVE BEEN REMOVED FROM THATLIST SINCE 1983 ARE INCLUDED; OTHER HISTORICAL RECORDS ARE NOT.

entry date: JUN 1991 effective date: 11AUG1988

amendment: CAGAAK, Canada Gazette Part II, 122 , 18 , 3601 ,

File: 17.01 LEGAL rn: 302142

systematic name: 2-Propanol common name :2-propanol reported name : ISOPROPANOL

cas no :67-63-0 :NT8050000 rtecs no : REG area

: CAN type ______

|subject|specification|descriptor| |------TRNSP | CLASS | LABEL | RQR PACK |

PIN (PRODUCT IDENTIFICATION NO.): UN1219. CLASS (3.2): FLAMMABLE LIQUID. PACKING GROUPII, (I=GREAT DANGER, III=MINOR DANGER). MAXIMUM AMOUNT PER PACKAGE THAT MAY BE TRANSPORTED ON A PASSENGER AIRCRAFT OR VEHICLE: 5 L. MAXIMUM AMOUNT PER PACKAGE THAT MAY BE TRANSPORTED ON A CARGO AIRCRAFT: 60 L. PRESCRIBED BY THE TRANSPORTATION OF DANGEROUS GOODS REGULATIONS, UNDER THE TRANSPORTATION OF DANGEROUS GOODS ACT (ADMINISTERED BY THE DEPARTMENT OF TRANSPORT). THE ACT AND REGULATIONS ARE INTENDED TO PROMOTE SAFETY IN THE TRANSPORTATION OF DANGEROUS GOODS IN CANADA, AS WELL AS PROVIDE ONE COMPREHENSIVE SET OF RULES APPLICABLE TO ALL MODES OF TRANSPORT ACCROSS CANADA. THESE ARE BASED ON UNITED NATIONS RECOMMENDATIONS. THE ACT AND REGULATIONS SHOULD BE CONSULTED FOR DETAILS. RECORDS ARE ENTERED UNDER THE PROPER SHIPPINGNAME FOUND IN THE REGULATIONS; THIS MAY INCLUDE VERY GENERAL GROUPS OF CHEMICAL SUBSTANCES.

entry date: OCT 1991 effective date: 06DEC1990

amendment: CAGAAK, Canada Gazette Part II, 124 , 26 , ,

File: 17.01 LEGAL rn: 303389

systematic name: 2-Propanol common name :2-propanol reported name :ISOPROPANOL

:67-63-0 : CAN rtecs no :NT8050000 cas no

: REG type area

|subject|specification|descriptor| -----USE occ l ROR | STORE | LABEL |

INGREDIENT DISCLOSURE LIST CONCENTRATION 1% WEIGHT/WEIGHT. THE WORKPLACE HAZARDOUS MATERIALS INFORMATION SYSTEM (WHMIS) IS A NATIONAL SYSTEM TO PROVIDE INFORMATION ON HAZARDOUS MATERIALS USED IN THE WORKPLACE. WHMIS IS IMPLEMENTED BY THE HAZARDOUS PRODUCTS ACT AND THE CONTROLLED PRODUCTS REGULATIONS (ADMINISTERED BY THE DEPARTMENT OF CONSUMER AND CORPORATE AFFAIRS). THE REGULATIONS IMPOSE STANDARDS ON EMPLOYERS FORTHE USE, STORAGE AND HANDLING OF CONTROLLED PRODUCTS AND ADDRESS LABELLING AND IDENTIFICATION, EMPLOYEE INSTRUCTION AND TRAINING, AS WELL AS THE UPKEEP OF A MATERIALS SAFETY DATA SHEET (MSDS). THE PRESENCE IN A CONTROLLED PRODUCT OF AN INGREDIENT IN A CONCENTRATION EQUAL TO OR GREATER THAN SPECIFIED IN THE INGREDIENT DISCLOSURE LIST MUST BE DISCLOSED IN THE SAFETY DATA SHEET.

entry date: APR 1991 effective date: 31DEC1987

amendment: CAGAAK, Canada Gazette Part II, 122, 2, 551,

File: 17.01 LEGAL rn: 401113

systematic name: 2-Propanol common name :2-propanol

reported name : ISOPROPYL ALCOHOL

rtecs no :NToo: REG cas no :67-63-0 :NT8050000

: CSK area

|subject|specification|descriptor| |-----FOOD MPC _____

LIMIT OF ADDITIVE PRESENT DUE TO PRODUCTION, PACKING, TRANSPORT AND

STORAGE OF FOOD PRODUCTS: 5G/KG.

entry date: DEC 1991 effective date: 1JUL1986

title: DIRECTIVE NO. 50/1978 ON FOREIGN SUBSTANCES IN FOODSTUFFS original : HPMZC*, HYGIENICKE PREDPISY MINISTERSTVA ZDRAVOTNICTVI CSR(HYGIENIC REGULATIONS OF MINISTRY OF HEALTH OF CSR), 43,

, , 1978

amendment: HPMZC*, HYGIENICKE PREDPISY MINISTERSTVA ZDRAVOTNICTVI

CSR(HYGIENIC REGULATIONS OF MINISTRY OF HEALTH OF CSR), 61 ,

, , 1986

File: 17.01 LEGAL rn: 402145

systematic name: 2-Propanol

: o, : CSK rtecs no :NT8050000 type : REG cas no :67-63-0

|subject|specification|descriptor| |----+-----| MANUF OCC CLASS RQR RSTR

PRODUCTION OF ISOPROPANOL BY SULFONATION IS CLASSIFIED AS PROCESS WITH OCCUPATIONAL CARCINOGENIC RISK. REQUIREMENTS AND RESTRICTIONS ON THE PROCESS ARE GIVEN.

effective date: 1APR1985 entry date: DEC 1991

title: DIRECTIVE NO. 4/1985 ON HYGIENIC PRINCIPLES FOR WORK WITH CHEMICAL CARCINOGENS

original: VMZSR*, Directive no 4/1985 on hygiene principles for work

with chemicalcarcinogens, , 3 , , 1985

amendment: VMZSR*, Directive no 4/1985 on hygiene principles for work

with chemicalcarcinogens, , 10 , 98 , 1990

File: 17.01 LEGAL rn: 500482

systematic name: 2-Propanol common name :2-propanol reported name :ISOPROPANOL

rtecs no :NT80 : REC cas no :67-63-0 :NT8050000

: DEU

|subject|specification|descriptor| AQ CLASS RQR USE INDST

THIS SUBSTANCE IS CLASSIFIED AS SLIGHTLY HAZARDOUS TO WATER (WATER-HAZARD CLASS: WGK 1). (THE DIFFERENT CLASSES ARE: WGK 3 = VERY HAZARDOUS; WGK 2 = HAZARDOUS; WGK 1 = SLIGHTLY HAZARDOUS; WGK 0 = IN GENERAL NOT HAZARDOUS.) THE CLASSIFICATION FORMS THE BASIS FOR WATER-PROTECTION REQUIREMENTS FOR INDUSTRIAL PLANTS IN WHICH WATER-HAZARDOUS SUBSTANCES ARE HANDLED.

entry date: DEC 1991

title: ADMINISTRATIVE RULES CONCERNING WATER-HAZARDOUS SUBSTANCES (VERWALTUNGSVORSCHRIFT WASSERGEFAEHRDENDE STOFFE)

original : GMSMA6, Gemeinsames Ministerialblatt. Joint Ministerial Papers, , 8 , 114 , 1990

rn: 502805 File: 17.01 LEGAL

systematic name: 2-Propanol common name :2-propanol

reported name :ISOPROPYL ALCOHOL

:67-63-0 :NT8050000 cas no rtecs no

: DEU : REC area type ______ |subject|specification|descriptor| |-----OCC MAK

8H-TWA: 400 ML/M3 (PPM); 980 MG/M3 (20C, 101.3 KPA). SUBSTANCE WITH SYSTEMIC EFFECTS. ONSET OF EFFECT <= 2H. HALF-LIFE < 2H. 30MIN-STEL: 800 ML/M3 (PPM); 1960 MG/M3; AVERAGE VALUE; 4X/SHIFT. PREGNANCY GROUP D: AVAILABLE DATA ARE NOT SUFFICIENT FOR A FINAL EVALUATION WITH REGARD TO EMBRYOTOXIC AND/OR FETOTOXIC EFFECTS. VAPOUR PRESSURE: 4.0 KPA AT 20C. entry date: JAN 1992

title: MAXIMUM CONCENTRATIONS AT THE WORKPLACE AND BIOLOGICAL TOLERANCE VALUES FOR WORKING MATERIALS (MAXIMALE ARBEITSPLATZKONZENTRATIONEN UND BIOLOGISCHE ARBEITSSTOFFTOLERANZWERTE)

original: MPGFDF, MITTEILUNG DER SENATSKOMMISSION ZUR PRUEFUNG GESUNDHEITSSCHAEDLICHER ARBEITSSTOFFE (DEUTSCHE FORSCHUNGSGEMEINSCHAFT), XXVII , , 17 , 1991

File: 17.01 LEGAL rn: 503265

systematic name: 2-Propanol common name :2-propanol

reported name :ISOPROPYL ALCOHOL

cas no :67-63-0 rtecs no :NT8050000

area : DEU type : REC

|subject|specification|descriptor| |-----AIR OCC BAT ______

BLOOD: 50 MG/L, AS ACETONE. URINE: 50 MG/L, AS ACETONE. SAMPLING TIME:

END OF EXPOSURE OR END OF SHIFT.

entry date: FEB 1992

title: MAXIMUM CONCENTRATIONS AT THE WORKPLACE AND BIOLOGICAL TOLERANCE VALUES FOR WORKING MATERIALS (MAXIMALE ARBEITSPLATZKONZENTRATIONEN UND BIOLOGISCHE ARBEITSSTOFFTOLERANZWERTE)

original : MPGFDF, MITTEILUNG DER SENATSKOMMISSION ZUR PRUEFUNG GESUNDHEITSSCHAEDLICHER ARBEITSSTOFFE (DEUTSCHE FORSCHUNGSGEMEINSCHAFT), XXVII, , 99, 1991

File: 17.01 LEGAL rn: 613032

systematic name: 2-Propanol common name :2-propanol reported name :ISOPROPANOL

rtecs no :67-63-0 :NT8050000 cas no

: REG : GBR area type

_____ |subject|specification|descriptor| |-----| | TRNSP | CLASS RQR LABEL |

LABELLING OF ROAD TANKERS: FLAMMABLE LIQUID. EMERGENCY ACTION CODE:

2(S)E

entry date: JAN 1983 effective date: 28MCH1979

title: HAZARDOUS SUBSTANCES (LABELLING OF ROAD TANKERS) REGULATIONS 1978

original: GBRSI*, STATUTORY INSTRUMENTS, 1702 , , , 1978

File: 17.01 LEGAL rn: 650703

systematic name:2-Propanol

common name :2-propanol
reported name :ISOPROPYL ALCOHOL

cas no :67-63-0 rtecs no type :NT8050000

: GBR area : REG

|subject|specification|descriptor| |------TRNSP | MARIN | RQR AQ | MARIN RQR EMI RQR

CLASSIFIED AS A NON-POLLUTING LIQUID SUBSTANCE. DOCUMENTARY EVIDENCE OF ASSESSMENT AND APPROVAL REQUIRED BY A CARRIER. DISCHARGE INTO THE SEA IS NOT PROHIBITED.

entry date: 1992 effective date: 06APR1987

title: THE MERCHANT SHIPPING (CONTROL OF POLLUTION BY NOXIOUS LIQUID

SUBSTANCES IN BULK) REGULATIONS 1987, SCHEDULE 2

original : GBRSI*, STATUTORY INSTRUMENTS, 551 , , 15 , 1987 amendment: GBRSI*, STATUTORY INSTRUMENTS, 2604 , , 2 , 1990

File: 17.01 LEGAL rn: 800152

systematic name: 2-Propanol common name :2-propanol

reported name :ISOPROPYL ALCOHOL

:67-63-0 rtecs no :NT8050000 cas no

: REC : JPN area type

|subject|specification|descriptor| |-----AIR OCC MAC ______

CLV: 980MG/M3 (400PPM). TENTATIVE VALUE.

entry date: DEC 1991

title: MAXIMUM ALLOWABLE CONCENTRATIONS RECOMMENDED BY THE JAPANESE ASSOCIATION OF INDUSTRIAL HEALTH.

original : SAIGBL, Sangyo Igalu (Japanese Journal of Industrial Health), 33 , 4 , 277-287 , 1991

File: 17.01 LEGAL rn: 911959

systematic name: 2-Propanol

common name :2-propanol

reported name :ISOPROPYL ALCOHOL

:67-63-0 : KEN rtecs no type :NT8050000 cas no

: REG

_____ |subject|specification|descriptor| |-----| FOOD | ADDIT | PRMT ______

FOOD ADDITIVE PERMITTED FOR EXTRACTING MOISTURE, FAT AND OTHER SOLUBLE COMPONENTS FROM FISH. FOOD PRODUCTS IN OR UPON WHICH IT IS PERMITTED AND MAXIUM LEVELS OF USE ARE LISTED

entry date: SEP 1982

title: THE FOOD, DRUGS AND CHEMICAL SUBSTANCES (FOOD LABELLING,

ADDITIVES AND STANDARDS) REGULATIONS, 1978

original : GSKEN*, KENYA GAZETTE SUPPLEMENT NO. 40, SPECIAL ISSUE (LEGISLATIVE SUPPLEMENT NO. 27), 40 , , 363 , 1978

rn : 1010068 File: 17.01 LEGAL

systematic name: 2-Propanol common name :2-propanol

reported name :ISOPROPYL ALCOHOL

rtecs no cas no :67-63-0 :NT8050000

area : MEX type : REG

______ |subject|specification|descriptor| |-----AIR OCC MXL ______

AT ANY WORKPLACE WHERE THIS SUBSTANCE IS PRODUCED, STORED OR HANDLED A MAXIMUM PERMISSIBLE LEVEL OF 980MG/M3 (400PPM) MUST BE OBSERVED FOR A PERIOD OF 8 HOURS OR 1225MG/M3 (500PPM) FOR 15 MINUTES FOUR TIMES A DAY WITH INTERVALS OF A LEAST 1 HOUR.

entry date: DEC 1991 effective date: 28MAY1984

title: INSTRUCTION NO.10 RELATED TO SECURITY AND HYGIENIC CONDITIONS AT WORKPLACES. (INSTRUCTIVO NO. 10, RELATIVO A LAS CONDICIONES DE SEGURIDAD E HIGIENE DE LOS CENTROS DE TRABAJO).

original : DOMEX*, Diario Oficial, , , , 1984 amendment: DOMEX*, Diario Oficial, , , , 1989

File: 17.01 LEGAL rn: 1122015

systematic name: 2-Propanol common name :2-propanol

reported name :ISOPROPYL ALCOHOL

rtecs no :67-63-0 :NT8050000 cas no

: REG : RUS area type

_____ |subject|specification|descriptor| |-----AIR AMBI MAC

0.6MG/M3 1X/D, 0.6MG/M3 AV/D.

entry date: SEP 1985 effective date: AUG1984

amendment: PDKAV*, PREDELNO DOPUSTIMYE KONTSENTRATSII (PDK)

ZAGRYAZNYAYUSHCHIKH VESHCHESTV V ATMOSFERNOM VOZDUKHE NASELENNYKH MEST (MAXIMUM ALLOWABLE CONCENTRATIONS (MAC) OF

CONTAMINANTS IN THE AMBIENT AIR OF RESIDENTIAL AREAS),

3086-84 , , , 1984

File: 17.01 LEGAL rn: 1122514

systematic name: 2-Propanol

cas no :67-63-0 rtecs no type :NT8050000

: REG : RUS area

|subject|specification|descriptor| |-----AIR OCC MAC CLASS _____

CLV: 10.0MG/M3 (VAPOUR) HAZARD CLASS: III

entry date: MAY 1990 effective date: 01JAN1989

amendment: GOSTS*, GOSUDARSTVENNYI STANDART SSSR(STATE STANDARD OF

USSR), 12.1.005 , , , 1988

File: 17.01 LEGAL rn: 1123251

systematic name: 2-Propanol common name :2-propanol

reported name : ISOPROPYL ALCOHOL

rtecs no :NT80 type : REG cas no :67-63-0 :NT8050000

: RUS area

|subject|specification|descriptor| |-----AQ SURF MAC CLASS

0.25MG/L HAZARD CLASS: IV

entry date: JUL 1990 effective date: 1JAN1989

amendment: SPNPV*, SANITARNYE PRAVILA I NORMY OKHRANY POVERKHNOSTNYKH VOD OT ZAGRIAZNENIA (HEALTH REGULATION AND STANDARDS OF

SURFACE WATER PROTECTION FROM CONTAMINATION), 4630-88 , , ,

1988

File: 17.01 LEGAL rn : 1200149

systematic name: 2-Propanol common name :2-propanol reported name : ISOPROPANOL

cas no :67-63-0 :NT8050000 rtecs no

: SWE : REG area type

______ |subject|specification|descriptor| |-----AIR OCC HLV

1D-TWA: 350MG/M3 (150PPM); 15MIN-STEL: 600MG/M3 (250PPM). (APPLIES TO

ALL ISOMERS OF PROPANOL).

entry date: 1992 effective date: 01JUL1991

title: HYGIENIC LIMIT VALUES.

original: AFS***, ARBETARSKYDDSSTYRELSENS FOERFATTNINGSSAMLING, 1990:13 , , 5-64 , 1990

File: 17.01 LEGAL rn: 1301002

systematic name: 2-Propanol common name :2-propanol reported name :2-Propanol

cas no :67-63-0 rtecs no :NT8050000

area : USA type : REG

|subject|specification|descriptor| |----+----| MANUF | REQ PRMT USE OCC PRMT MXL

; Summary - THE FOLLOWING CHEMICAL IS INCLUDED ON A LIST OF CHEMICALS AND MIXTURES FOR WHICH REPORTING IS CURRENTLY REQUIRED UNDER THE TOXIC SUBSTANCES CONTROL ACT SECTION 2607A. THIS TOXIC SUBSTANCE IS SUBJECT TO PRELIMINARY ASSESSMENT INFORMATION RULES ON PRODUCT ION QUANTITIES, USES, EXPOSURES, AND ADVERSE EFFECTS. MANUFACTURERS INCLUDING IMPORTERS MUST SUBMIT A REPORT FOR THIS LISTED CHEMICAL MANUFACTURED AT EACH SITE. entry date: OCT 1991 effective date:

title: PRELIMINARY ASSESSMENT INFORMATION RULES original: FEREAC, Federal Register, 47 , , 26998 , 1982 amendment: CFRUS*, Code of Federal Regulations, 40 , 712 , 30 , 1990

File: 17.01 LEGAL rn: 1302342

systematic name: 2-Propanol common name :2-propanol

reported name :ISOPROPYL ALCOHOL

rtecs no :67-63-0 :NT8050000 cas no

: REG : USA area type

_____ |subject|specification|descriptor| |----+----ADDIT | FOOD | RSTR RSTR TRANS RSTR | STORE | RSTR PACK

; Summary - THIS SUBSTANCE IS INCLUDED ON A LIST OF SUBSTANCES USED TO PREPARE ADHESIVES WHICH MAY BE SAFELY USED AS COMPONENTS OF ARTICLES INTENDED FOR USE IN PACKAGING, TRANSPORTATION, OR HOLDING FOOD IN

ACCORDANCE WITH THE FOLLOWING PRESCRIBED CONDITIONS: SUBSTA NCE MUST BE SEPARATED FROM THE FOOD BY A FUNCTIONAL BARRIER, MUST NOT EXCEED LIMITS OF GOOD MANUFACTURING PRACTICE USED WITH DRY FOODS, OR NOT EXCEED TRACE AMOUNTS AT SEAMS AND EDGE EXPOSURES WHEN USED WITH FATTY AND AOUEOUS FOODS. ALSO REGULATED BY SEA M INTEGRITY, LABELING STANDARDS, AND ANY PROVISION UNDER 21 CFR 175

entry date: NOV 1991 effective date: 1977

title: SUBSTANCES FOR USE ONLY AS COMPONENTS OF ADHESIVES original: FEREAC, Federal Register, 42 , , 14534 , 1977

amendment: CFRUS*, Code of Federal Regulations, 21 , 175 , 105 , 1988

File: 17.01 LEGAL rn: 1314568

systematic name: 2-Propanol common name :2-propanol reported name :ISOPROPANOL

cas no :67-63-0 rtecs no :NT8050000

area : USA type : REG

_____ |subject|specification|descriptor| |-----TRNSP | PRMT CNTRL PACK | LABEL RQR

MAY BE TRANSPORTED IN PASSENGER AIRCRAFT AND PASSENGER RAILCAR NOT TO EXCEED 1 QUART/PACKAGE. MAY BE TRANSPORTED IN CARGO AIRCRAFT NOT TO EXCEED 10 GALLONS/PACKAGE. MAY BE TRANSPORTED IN CARGO VESSELS ON AND BELOW DECK AND IN PASSENGER VESSELS ON DECK. ALL SHIPMENTS MUST BE LABELED FLAMMABLE LIQUID.; Summary - THIS REGULATION LISTS AND CLASSIFIES THOSE MATERIALS WHICH THE DEPARTMENT OF TRANSPORTATION HAS DESIGNATED AS HAZARDOUS MATERIALS FOR SHIPPING PAPERS, PACKAGE MARKING, LABELING, AND TRANSPORT VEHICLE PLACARDING APPLICABLE TO THE SHIPMENT AND TRANSPORT OF THOSE HAZARDOUS MATERIALS.

entry date: NOV 1991 effective date: OCT1991

title: HAZARDOUS MATERIALS REGULATIONS, PART 172--HAZARDOUS MATERIALS TABLES AND HAZARDOUS MATERIALS COMMUNICATIONS REGULATIONS

original: CFRUS*, Code of Federal Regulations, 49 , 172 , 101 , 1984 amendment: CFRUS*, Code of Federal Regulations, 49 , 172 , 101 , 1990

rn: 1323007 File: 17.01 LEGAL

systematic name: 2-Propanol common name :2-propanol reported name :ISOPROPANOL

rtecs no :67-63-0 :NT8050000 cas no

: REG : USA area type

______ |subject|specification|descriptor| |-----PESTI | RQR PESTI | PRMT | CLASS | PESTI MANUF | ADDIT RQR FOOD |

CASE NAME ALIPHATIC ALCOHOLS, C1-C5; Summary - THIS SUBSTANCE IS

INCLUDED ON A LIST OF ACTIVE INGREDIENTS CONTAINED IN A PRODUCT FIRST REGISTERED BEFORE NOVEMBER 1, 1984, FOR WHICH A REGISTRATION STANDARD HAS NOT BEEN ISSUED. PUBLICATION OF THIS LIST INITIATES AN ACCELERATED REREGISTRATION AND DATA C ALL-IN FOR PRODUCTS CONTAINING THE LISTED ACTIVE INGREDIENTS.

entry date: JAN 1992 effective date: 1989

title: FEDERAL INSECTICIDE, FUNGICIDE, AND RODENTICIDE ACT PESTICIDES

REQUIRED TO BE REREGISTERED; LIST D

original : FEREAC, Federal Register, 54 , 204 , 43388 , 1989 amendment: FEREAC, Federal Register, 54 , 204 , 43388 , 1989

File: 17.01 LEGAL rn: 1325037

systematic name: 2-Propanol common name :2-propanol reported name :ISOPROPANOL

cas no :67-63-0 rtecs no :NT8050000

area : USA type : REC

______ |subject|specification|descriptor| |-----SAFTY | OCC MXL MXL USE | OCC

12000 PPM

entry date: OCT 1991 effective date: JUN1990

title: POCKET GUIDE TO CHEMICAL HAZARDS

original: XPHPAW, US PUBLIC HEALTH SERVICE PUBLICATION, 90, 117, 132

, 1990

amendment: XPHPAW, US PUBLIC HEALTH SERVICE PUBLICATION, 90 , 117 , 132

, 1990

File: 17.01 LEGAL rn: 1336031

systematic name: 2-Propanol common name :2-propanol reported name :2-Propanol

rtecs no cas no :67-63-0 :NT8050000

: REG : USA area type

_____ |subject|specification|descriptor| |-----EMI EMI AIR RQR RQR SOIL EMI RQR | AQ EMI RQR MANUF _____

; Summary - FACILITIES THAT EXCEEDED A MANUFACTURING, IMPORTATION, OR PROCESSING THRESHOLD OF 25,000 LBS OR THE USE OF 10,000 LBS FOR THIS CHEMICAL MUST REPORT TO EPA ANY RELEASES OF THE CHEMICAL (OR CATEGORY CHEMICAL) TO AIR, LAND, WATER, POTW, UNDERGROUND INJECTIO N, OR OFF SITE TRANSFER. THIS REGULATION COVERS STANDARD INDUSTRIAL CLASSIFICATION

(SIC) CODES 20-39 ONLY).

entry date: OCT 1991 effective date: 1987

title: SUPERFUND AMENDMENTS AND REAUTHORIZATION ACT, TITLE III. EPCRA

SECTION 313 LIST OF TOXIC SUBSTANCES

original: CFRUS*, Code of Federal Regulations, 40 , 372 , 65 , 1988 amendment: CFRUS*, Code of Federal Regulations, 40 , 372 , 65 , 1988

File: 17.01 LEGAL rn: 1340603

systematic name: 2-Propanol

cas no :67-63-0 rtecs no type :NT8050000

: USA area : REC

_____ |subject|specification|descriptor| |-----AIR OCC TLV _____

Time Weighted Avg (TWA) 400 ppm, 985 MG/M3; Short Term Exposure Limit (STEL) 500 ppm, 1230 MG/M3; Summary - THIS THRESHOLD LIMIT VALUE IS INTENDED FOR USE IN THE PRACTICE OF INDUSTRIAL HYGIENE AS A GUIDELINE OR RECOMMENDATION IN THE CONTROL OF POTENTIAL HEALTH HAZARDS.

entry date: DEC 1991 effective date:

title: THRESHOLD LIMIT VALUES

original : ACGIH*, Threshold Limit Values and Biological Exposure

Indices, , , 11 , 1989

amendment: ACGIH*, Threshold Limit Values and Biological Exposure

Indices, , , 11 , 1991

File: 17.01 LEGAL rn: 1345002

systematic name: 2-Propanol common name :2-propanol reported name :2-Propanol

cas no :67-63-0 rtecs no :NT8050000

: REG : USA area type

|subject|specification|descriptor| |-----| MONIT | RQR _____

; Summary - THIS IS A CHEMICAL OR MIXTURE FOR WHICH REPORTING IS CURRENTLY REQUIRED UNDER THE TOXIC SUBSTANCE CONTROL ACT HEALTH AND SAFETY STUDIES SECTION 2607D. PERSONS WHO CURRENTLY MANUFACTURE OR PROCESS CHEMICAL SUBSTANCES OR MIXTURES FOR COMMERCIAL PURPOSES, THOSE WHO PROPOSE TO DO SO, AND THOSE WHO ARE NOT CURRENTLY INVOLVED WITH A LISTED CHEMICAL BUT WHO MANUFACTURED OR PROCESSED IT OR PROPOSED TO DO SO ANY TIME DURING THE TEN YEAR PERIOD PRIOR TO THE TIME IT BECAME LISTED MUST SUBMIT TO THE ADMINISTRATOR OF THE U.S. EPA STUDIES OR LISTS OF HEALTH AND SAFETY STUDIES CONDUCTED ON THIS SUBSTANCE FOR EVALUATION.

entry date: OCT 1991 effective date:

title: HEALTH AND SAFETY DATA REPORTING RULES SECTION 8(D) original : FEREAC, Federal Register, 51 , , 32726 , 1986

amendment: CFRUS*, Code of Federal Regulations, 40 , 716 , 120 , 1990

File: 17.01 LEGAL rn: 1346073

systematic name: 2-Propanol

: USA rtecs no :NT8050000 type : REG :67-63-0 cas no

area

|subject|specification|descriptor| -----FOOD | ADDIT | RSTR ADDIT RSTR PACK | GL | MANUF | ADDIT RSTR USE

-----; Summary - THIS SUBSTANCE INCLUDED ON A LIST OF SUBSTANCES USED TO PREPARE BASE SHEET OR COATING SUBSTANCES FOR CELLOPHANE MUST BE OF A GRADE OF PURITY SUITABLE FOR USE IN FOOD PACKAGING TO IMPART THE DESIRED TECHNOLOGICAL PROPERTIES. ACRYLONITRILE COPOLYMER SUBST ANCES MUST ABIDE UNDER THE CONDITIONS GIVEN IN 21 CFR 180.22 1988.

entry date: NOV 1991 effective date:

title: INDIRECT FOOD ADDITIVES; POLYMERS-CELLOPHANE. original : FEREAC, Federal Register, 42 , , 14572 , 1977

amendment: CFRUS*, Code of Federal Regulations, 21 , 177 , 1200 , 1988

File: 17.01 LEGAL rn: 1400094

systematic name: 2-Propanol common name :2-propanol

reported name : ISOPROPYL ALCOHOL

rtecs no :NT80 cas no :67-63-0 :NT8050000 : EEC

_____ |subject|specification|descriptor| |-----FOOD ADDIT PRMT ______

THIS SUBSTANCE MAY BE USED AS A DILUTANT OR SOLVENT FOR COLOURING MATTERS AUTHORIZED FOR USE IN FOODSTUFFS INTENDED FOR HUMAN CONSUMPTION. entry date: FEB 1986

title: COUNCIL DIRECTIVE ON THE APPROXIMATION OF THE RULES OF THE MEMBER STATES CONCERNING THE COLOURING MATTERS AUTHORISED FOR USE IN FOODSTUFFS INTENDED FOR HUMAN CONSUMPTION.

 $\verb|original: OJEC**|, Official Journal of the European (Communities)/Union,\\$ 115 , , 2645/62 , 1962

amendment: OJEC**, Official Journal of the European (Communities)/Union, L43 , , 11 , 1981

File: 17.01 LEGAL rn: 1402098

systematic name: 2-Propanol common name :2-propanol

reported name :ISOPROPYL ALCOHOL

rtecs no type cas no :67-63-0 :NT8050000

: EEC : REG

_____ |subject|specification|descriptor| |------FOOD MXLRSTR FOOD

THE SUBSTANCE MAY BE USED FOR THE MANUFACTURE OF REGENERATED CELLULOSE FILM WHICH IS INTENDED TO OR DOES COME INTO CONTACT WITH FOODSTUFFS. IT MAY BY USED AS SOLVENT; MAXIMUM TOTAL QUANTITY OF ALL SOLVENTS:

0.6MG/DM2 ON THE SIDE IN CONTACT WITH FOODSTUFFS.

entry date: OCT 1987 effective date: 01APR1987

title: COUNCIL DIRECTIVE OF 25 APRIL 1983 ON THE APPROXIMATION OF THE LAWS OF THE MEMBER STATES RELATING TO MATERIALS AND ARTICLES MADE OF REGENERATED CELLULOSE FILM INTENDED TO COME INTO CONTACT WITH

FOODSTUFFS. (83/229/EEC).

original: OJEC**, Official Journal of the European (Communities)/Union,

L123 , , 31 , 1983

amendment: OJEC**, Official Journal of the European (Communities)/Union,

L228 , , 32 , 1986

rn: 1402213 File: 17.01 LEGAL

systematic name: 2-Propanol common name :2-propanol

reported name : ISOPROPYL ALCOHOL

:67-63-0 :NT8050000 rtecs no

: EEC : REG type

|subject|specification|descriptor| |-----GOODS | FUEL FUEL RQR GOODS

MEMBER STATES MAY NOT PREVENT OR DISCOURAGE, ON GROUNDS OF OXYGENATE CONTENT, THE PRODUCTION, MARKETING AND FREE MOVEMENT OF BLENDED PETROL CONTAINING ISOPROPYL ALCOHOL UP TO 5% (VOL.). PUMPS WHICH DISPENSE MOTOR FUELS CONTAINING MORE THAN 10% (VOL.) OF ISOPROPYL ALCOHOL SHALL BE VERY CLEARLY MARKED ACCORDINGLY TO MAKE ACCOUNT IN PARTICULAR OF VARIATIONS IN THE CALORIFIC VALUE OF SUCH FUELS

entry date: FEB 1986 effective date: 01JAN1988

title: COUNCIL DIRECTIVE OF 5 DECEMBER 1985 ON CRUDE-OIL SAVINGS THROUGH THE USE OF SUBSTITUTE FUEL COMPONENTS IN PETROL. (85/536/EEC).

original: OJEC**, Official Journal of the European (Communities)/Union, L334 , , 20 , 1985

File: 17.01 LEGAL rn: 1402713

systematic name: 2-Propanol common name :2-propanol

reported name :ISOPROPYL ALCOHOL

:67-63-0 :NT8050000 cas no rtecs no

area : EEC type : REG

STRONG ACID PROCESS IN THE MANUFACTURE OF ISOPROPYL ALCOHOL IS A PROCESS BY WHICH WORKERS ARE OR ARE LIKELY TO BE EXPOSED TO CARCINOGENS AS A RESULT OF THEIR WORK. IN THIS TYPE OF ACTIVITY, THE RISK TO THE HEALTH AND SAFETY OF WORKERS MUST BE ASSESSED. EMPLOYERS' OBLIGATIONS INCLUDE REDUCTION OF THE USE OF THE PROCESS AND WHERE POSSIBLE ITS REPLACEMENT BY A LESS DANGEROUS PROCESS. WHERE THE REPLACEMENT OF THE PROCESS IS NOT POSSIBLE, MEASURES TO ENSURE THE PROTECTION OF THE WORKERS AND TO REDUCE THE LEVEL OF THEIR EXPOSURE MUST BE TAKEN. EMPLOYERS SHALL WHEN REQUESTED, MAKE AVAILABLE TO THE COMPETENT AUTHORITY APPROPRIATE INFORMATION. THE WORKERS AND/OR THEIR REPRESENTATIVES MUST BE INFORMED ABOUT THE HEALTH RISKS, HYGIENIC REQUIREMENTS AND PRECAUTIONS TO BE TAKEN AND THEY CAN CHECK THAT THIS DIRECTIVE IS APPLIED. RELEVANT HEALTH SURVEILLANCE OF WORKERS IS REQUIRED. THE MEDICAL RECORDS SHALL BE KEPT FOR AT LEAST 40 YEARS.

entry date: JUN 1992 effective date: 31DEC1992

title: COUNCIL DIRECTIVE OF 28 JUNE 1990 ON THE PROTECTION OF WORKERS FROM THE RISKS RELATED TO EXPOSURE TO CARCINOGENS AT WORK (SIXTH INDIVIDUAL DIRECTIVE WITHIN THE MEANING OF ARTICLE 16(1) OF DIRECTIVE 89/391/EEC). (90/394/EEC).

original : OJEC**, Official Journal of the European (Communities)/Union, 196 , , 1 , 1990

File: 17.01 LEGAL rn: 1646178

systematic name:2-Propanol
common name :2-propanol
reported name :ISOPROPANOL

cas no :67-63-0 rtecs no :NT8050000

area : IMO type : REC

| subject | specification | descriptor | |------| | TRNSP | MARIN | CLASS | | LABEL | | |

HAZARD CLASS: 3 = INFLAMMABLE LIQUID. PACKING GROUP: II = MEDIUM DANGER

(I=GREAT DANGER - III=MINOR DANGER). UN NO. 1219

entry date: JAN 1991

amendment: !IMCOC*, International Maritime Dangerous Goods Code, , ,

10004 , 1990

File: 17.01 LEGAL rn : 1744419

systematic name:2-Propanol
common name :2-propanol
reported name :ISOPROPANOL

cas no :67-63-0 rtecs no :NT8050000

HAZARD CLASS: 3 = INFLAMMABLE LIQUID. PACKING GROUP: II = MEDIUM DANGER

(I=GREAT DANGER - III=MINOR DANGER). UN NO. 1219

entry date: AUG 1990

 $\verb|amendment: !UNTDG*|, UN Transport of Dangerous Goods|, Recommendation|\\$

prepared by the Committee of Experts on the Transport of

Dangerous Goods, , , 15 , 1989