# TOXICOLOGICAL PROFILE FOR CHLOROETHANE

U.S. DEPARTMENT OF HEALTH AND HUMAN SERVICES
Public Health Service
Agency for Toxic Substances and Disease Registry

December 1998

# **DISCLAIMER**

The use of company or product name(s) is for identification only and does not imply endorsement by the Agency for Toxic Substances and Disease Registry.

# **UPDATE STATEMENT**

A Toxicological Profile for chloroethane was released in December 1989. This edition supersedes any previously released draft or final profile.

Toxicological profiles are revised and republished as necessary, but no less than once every three years. For information regarding the update status of previously released profiles, contact ATSDR at:

Agency for Toxic Substances and Disease Registry Division of Toxicology/Toxicology Information Branch 1600 Clifton Road NE, E-29 Atlanta, Georgia 30333

#### **FOREWORD**

This toxicological profile is prepared in accordance with guidelines\* developed by the Agency for Toxic Substances and Disease Registry (ATSDR) and the Environmental Protection Agency (EPA). The original guidelines were published in the Federal Register on April 17, 1987. Each profile will be revised and republished as necessary.

The ATSDR toxicological profile succinctly characterizes the toxicologic and adverse health effects information for the hazardous substance described therein. Each peer-reviewed profile identifies and reviews the key literature that describes a hazardous substance's toxicologic properties. Other pertinent literature is also presented, but is described in less detail than the key studies. The profile is not intended to be an exhaustive document; however, more comprehensive sources of specialty information are referenced.

The focus of the profiles is on health and toxicologic information; therefore, each toxicological profile begins with a public health statement that describes, in nontechnical language, a substance's relevant toxicological properties. Following the public health statement is information concerning levels of significant human exposure and, where known, significant health effects. The adequacy of information to determine a substance's health effects is described in a health effects summary. Data needs that are of significance to protection of public health are identified by ATSDR and EPA.

Each profile includes the following:

- (A) The examination, summary, and interpretation of available toxicologic information and epidemiologic evaluations on a hazardous substance to ascertain the levels of significant human exposure for the substance and the associated acute, subacute, and chronic health effects;
- (B) A determination of whether adequate information on the health effects of each substance is available or in the process of development to determine levels of exposure that present a significant risk to human health of acute, subacute, and chronic health effects; and
- (C) Where appropriate, identification of toxicologic testing needed to identify the types or levels of exposure that may present significant risk of adverse health effects in humans.

The principal audiences for the toxicological profiles are health professionals at the Federal, State, and local levels; interested private sector organizations and groups; and members of the public.

This profile reflects ATSDR's assessment of all relevant toxicologic testing and information that has been peer-reviewed. Staff of the Centers for Disease Control and Prevention and other Federal scientists have also reviewed the profile. In addition, this profile has been peer-reviewed by a nongovernmental panel and was made available for public review. Final responsibility for the contents and views expressed in this toxicological profile resides with ATSDR.

> Jeffrey Koplan Jeffrey P. Koplan, M.D., M.P.H.

> > Administrator

Agency for Toxic Substances and

Disease Registry

#### \*Legislative Background

The toxicological profiles are developed in response to the Superfund Amendments and Reauthorization Act (SARA) of 1986 (Public Law 99-499) which amended the Comprehensive Environmental Response, Compensation, and Liability Act of 1980 (CERCLA or Superfund). This public law directed ATSDR to prepare toxicological profiles for hazardous substances most commonly found at facilities on the CERCLA National Priorities List and that pose the most significant potential threat to human health, as determined by ATSDR and the EPA. The availability of the revised priority list of 275 hazardous substances was announced in the *Federal Register* on November 17, 1997 (62 FR 61332). For prior versions of the list of substances, see *Federal Register* notices dated April 29, 1996 (61 FR 18744); April 17, 1987 (52 FR 12866); October 20, 1988 (53 FR 41280); October 26, 1989 (54 FR 43619); October 17,1990 (55 FR 42067); October 17, 1991 (56 FR 52166); October 28, 1992 (57 FR 48801); and February 28, 1994 (59 FR 9486). Section 104(i)(3) of CERCLA, as amended, directs the Administrator of ATSDR to prepare a toxicological profile for each substance on the list.

CHLOROETHANE vii

# **QUICK REFERENCE FOR HEALTH CARE PROVIDERS**

Toxicological Profiles are a unique compilation of toxicological information on a given hazardous substance. Each profile reflects a comprehensive and extensive evaluation, summary, and interpretation of available toxicologic and epidemiologic information on a substance. Health care providers treating patients potentially exposed to hazardous substances will find the following information helpful for fast answers to often-asked questions.

# Primary Chapters/Sections of Interest

**Chapter 1: Public Health Statement:** The Public Health Statement can be a useful tool for educating patients about possible exposure to a hazardous substance. It explains a substance's relevant toxicologic properties in a nontechnical, question-and-answer format, and it includes a review of the general health effects observed following exposure.

**Chapter 2: Health Effects:** Specific health effects of a given hazardous compound are reported by route *of exposure*, by *type of health effect* (death, systemic, immunologic, reproductive), and by *length of exposure* (acute, intermediate, and chronic). In addition, both human and animal studies are reported in this section.

**NOTE:** Not all health effects reported in this section are necessarily observed in the clinical setting. Please refer to the Public Health Statement to identify general health effects observed following exposure.

**Pediatrics:** Four new sections have been added to each Toxicological Profile to address child health issues:

Section 1.6 How Can (Chemical X) Affect Children?

Section 1.7 How Can Families Reduce the Risk of Exposure to (Chemical X)?

Section 2.6 Children's Susceptibility

Section 5.6 Exposures of Children

#### Other Sections of Interest:

Section 2.7 Biomarkers of Exposure and Effect

Section 2.10 Methods for Reducing Toxic Effects

## ATSDR Information Center

**Phone:** 1-800-447-1544 (to be replaced by 1-888-42-ATSDR in 1999)

The following additional material can be ordered through the ATSDR Information Center:

Case Studies in Environmental Medicine: Taking an Exposure History--The importance of taking an exposure history and how to conduct one are described, and an example of a thorough exposure history is provided. Other case studies of interest include Reproductive and Developmental Hazards; Skin Lesions and Environmental Exposures; Cholinesterase-Inhibiting Pesticide Toxicity; and numerous chemical-specific case studies.

Managing Hazardous Materials Incidents is a three-volume set of recommendations for on-scene (prehospital) and hospital medical management of patients exposed during a hazardous materials incident. Volumes I and II are planning guides to assist first responders and hospital emergency department personnel in planning for incidents that involve hazardous materials. Volume III-Medical Management Guidelines for Acute Chemical Exposures-is a guide for health care professionals treating patients exposed to hazardous materials.

Fact Sheets (ToxFAQs) provide answers to frequently asked questions about toxic substances.

# Other Agencies and Organizations

The National Centerfor Environmental Health (NCEH) focuses on preventing or controlling disease, injury, and disability related to the interactions between people and their environment outside the workplace. Contact: NCEH, Mailstop F-29,4770 Buford Highway, NE, Atlanta, GA 30341-3724
• Phone: 770-488-7000 • FAX: 770-488-7015.

The National Institute for Occupational Safety and Health (NIOSH) conducts research on occupational diseases and injuries, responds to requests for assistance by investigating problems of health and safety in the workplace, recommends standards to the Occupational Safety and Health Administration (OSHA) and the Mine Safety and Health Administration (MSHA), and trains professionals in occupational safety and health. Contact: NIOSH, 200 Independence Avenue, SW, Washington, DC 20201 • Phone: 800-356-4674 or NIOSH Technical Information Branch, Robert A. Taft Laboratory, Mailstop C-19,4676 Columbia Parkway, Cincinnati, OH 45226-1998 • Phone: 800-35-NIOSH.

The National Institute of Environmental Health Sciences (NIEHS) is the principal federal agency for biomedical research on the effects of chemical, physical, and biologic environmental agents on human health and well-being. Contact: NIEHS, PO Box 12233, 104 T.W. Alexander Drive, Research Triangle Park, NC 27709 • Phone: 919-541-3212.

#### Referrals

The Association of Occupational and Environmental Clinics (AOEC) has developed a network of clinics in the United States to provide expertise in occupational and environmental issues. Contact: AOEC, 1010 Vermont Avenue, NW, #513, Washington, DC 20005 • Phone: 202-347-4976 • FAX: 202-347-4950 • e-mail: aoec@dgs.dgsys.com • AOEC Clinic Director: http://occ-env/med.mc.duke.edu/oem/aoec.htm.

The American College of Occupational and Environmental Medicine (ACOEM) is an association of physicians and other health care providers specializing in the field of occupational and environmental medicine. Contact: ACOEM, 55 West Seegers Road, Arlington Heights, IL 60005 • Phone: 847-228-6850 • FAX: 847-228-1856.

CHLOROETHANE ix

## **CONTRIBUTORS**

#### CHEMICAL MANAGER(S)/AUTHOR(S):

G. Daniel Todd, Ph.D. ATSDR, Division of Toxicology, Atlanta, GA

Carol Eisenmann, Ph.D. Sciences International, Inc., Alexandria, VA

Kara B. Altshuler, Ph.D. Sciences International, Inc., Alexandria, VA

#### THE PROFILE HAS UNDERGONE THE FOLLOWING ATSDR INTERNAL REVIEWS:

- 1 . Health Effects Review. The Health Effects Review Committee examines the health effects chapter of each profile for consistency and accuracy in interpreting health effects and classifying end points.
- 2 . Minimal Risk Level Review. The Minimal Risk Level Workgroup considers issues relevant to substance-specific minimal risk levels (MRLs), reviews the health effects database of each profile, and makes recommendations for derivation of MRLs.
- 3 . Data Needs Review. The Research Implementation Branch reviews data needs sections to assure consistency across profiles and adherence to instructions in the Guidance.

CHLOROETHANE xi

#### **PEER REVIEW**

A peer review panel was assembled for chloroethane. The panel consisted of the following members:

- 1. Dr. Martin Alexander, Cornell University, Ithaca, NY.
- 2. Dr. Syed GhiasUddin, Toxicologist, Indiana Department of Environmental Management, Indianapolis, IN.
- 3. Dr. Shane Que Hee, UCLA School of Public Health, Los Angeles, CA.

These experts collectively have knowledge of chloroethane's physical and chemical properties, toxicokinetics, key health end points, mechanisms of action, human and animal exposure, and quantification of risk to humans. All reviewers were selected in conformity with the conditions for peer review specified in Section 104(1)(13) of the Comprehensive Environmental Response, Compensation, and Liability Act, as amended.

Scientists from the Agency for Toxic Substances and Disease Registry (ATSDR) have reviewed the peer reviewers' comments and determined which comments will be included in the profile. A listing of the peer reviewers' comments not incorporated in the profile, with a brief explanation of the rationale for their exclusion, exists as part of the administrative record for this compound. A list of databases reviewed and a list of unpublished documents cited are also included in the administrative record.

The citation of the peer review panel should not be understood to imply its approval of the profile's final content. The responsibility for the content of this profile lies with the ATSDR.

# **CONTENTS**

FOREW	ORD			v
QUICK I	REFERE	NCE FOR I	HEALTH CARE PROVIDERS	. vii
CONTRI	BUTORS	S		ix
PEER RE	EVIEW .			xi
LIST OF	FIGURE	ES		xvii
LIST OF	TABLES	5		. xix
1. PUBL	IC HEAI	LTH STAT	EMENT	1
1.1			OETHANE?	
1.2			TO CHLOROETHANE WHEN IT ENTERS THE ENVIRONMENT?	
1.3			E EXPOSED TO CHLOROETHANE?	
1.4			ROETHANE ENTER AND LEAVE MY BODY?	
1.5	HOW C	AN CHLO	ROETHANE AFFECT MY HEALTH?	4
1.6	HOW C	AN CHLO	ROETHANE AFFECT CHILDREN?	6
1.7	HOW C	AN FAMII	LIES REDUCE THE RISK OF EXPOSURE TO CHLOROETHANE?	7
1.8	IS THEF	RE A MED	ICAL TEST TO DETERMINE WHETHER I HAVE BEEN EXPOSED	
			ANE?	8
1.9			ENDATIONS HAS THE FEDERAL GOVERNMENT MADE TO	
			N HEALTH?	
1.10	WHERE	E CAN I GI	ET MORE INFORMATION?	9
2. HEAI	TH EFF	ECTS		. 11
2.1			•••••	
2.2			HEALTH EFFECTS BY ROUTE OF EXPOSURE	
	2.2.1	Inhalation	Exposure	. 13
		2.2.1.1	Death	
		2.2.1.2	Systemic Effects	. 29
		2.2.1.3	Immunological and Lymphoreticular Effects	. 38
		2.2.1.4	Neurological Effects	
		2.2.1.5	Reproductive Effects	. 42
		2.2.1.6	Developmental Effects	. 43
		2.2.1.7	Genotoxic Effects	. 44
		2.2.1.8	Cancer	. 44
	2.2.2	Oral Expo	sure	. 45
		2.2.2.1	Death	. 45
		2.2.2.2	Systemic Effects	
		2.2.2.3	Immunological and Lymphoreticular Effects	
		2.2.2.4	Neurological Effects	
		2.2.2.5	Reproductive Effects	. 46

CHLOROETHANE xiv

		2.2.2.6	Developmental Effects	. 46
		2.2.2.7	Genotoxic Effects	. 46
		2.2.2.8	Cancer	. 46
	2.2.3	Dermal E	xposure	. 46
		2.2.3.1	Death	. 46
		2.2.3.2	Systemic Effects	. 46
		2.2.3.3	Immunological and Lymphoreticular Effects	. 47
		2.2.3.4	Neurological Effects	. 48
		2.2.3.5	Reproductive Effects	. 48
		2.2.3.6	Developmental Effects	. 48
		2.2.3.7	Genotoxic Effects	. 48
		2.2.3.8	Cancer	. 48
2.3	TOXICO	OKINETIC	'S	. 48
	2.3.1	Absorptio	n	. 49
		2.3.1.1	Inhalation Exposure	
		2.3.1.2	Oral Exposure	
		2.3.1.3	Dermal Exposure	
	2.3.2	Distributi	on	
		2.3.2.1	Inhalation Exposure	
		2.3.2.2	Oral Exposure	
		2.3.2.3	Dermal Exposure	
	2.3.3		.m	
	2.3.4		on and Excretion	
		2.3.4.1	Inhalation Exposure	
		2.3.4.2	Oral Exposure	
		2.3.4.3	Dermal Exposure	
	2.3.5		ically Based Pharmacokinetic (PBPK)/Pharmacodynamic (PD) Models	
2.4			F ACTION	
	2.4.1		kinetic Mechanisms	
	2.4.2		ms of Toxicity	
	2.4.3		o-Human Extrapolations	
2.5			PUBLIC HEALTH	
2.6			SCEPTIBILITY	
2.7			F EXPOSURE AND EFFECT	
2.7	2.7.1		rs Used to Identify or Quantify Exposure to Chloroethane	
	2.7.2		rs Used to Characterize Effects Caused by Chloroethane	
2.8			WITH OTHER CHEMICALS	
2.9			THAT ARE UNUSUALLY SUSCEPTIBLE	
2.10			REDUCING TOXIC EFFECTS	
2.10	2.10.1		Peak Absorption Following Exposure	
	2.10.1	_	Body Burden	
	2.10.2	U	g with the Mechanism of Action for Toxic Effects	
2 11			THE DATABASE	
∠.11	•			
	2.11.1	_	Information on Health Effects of Chloroethane	
	2.11.2		tion of Data Needs	
	2.11.3	Ungoing :	Studies	. 89

3.	CHE		ND PHYSICAL INFORMATION	
	3.1		CAL IDENTITY	
	3.2	PHYSIC	CAL AND CHEMICAL PROPERTIES	91
4.	PROI	OUCTION	N, IMPORT/EXPORT, USE, AND DISPOSAL	95
	4.1		CTION	
	4.2		Γ/EXPORT	
	4.3			
	4.4		AL	
5.	POTE	ENTIAL F	FOR HUMAN EXPOSURE	99
	5.1	OVERV	IEW	99
	5.2	RELEAS	SES TO THE ENVIRONMENT	101
		5.2.1	Air	101
		5.2.2	Water	103
		5.2.3	Soil	103
	5.3	ENVIRO	ONMENTAL FATE	104
		5.3.1	Transport and Partitioning	104
		5.3.2	Transformation and Degradation	
			5.3.2.1 Air	105
			5.3.2.2 Water	105
			5.3.2.3 Sediment and Soil	106
	5.4	LEVELS	S MONITORED OR ESTIMATED IN THE ENVIRONMENT	107
		5.4.1	Air	107
		5.4.2	Water	107
		5.4.3	Sediment and Soil	108
		5.4.4	Other Environmental Media	
	5.5	GENER	AL POPULATION AND OCCUPATIONAL EXPOSURE	
	5.6	EXPOS	URES OF CHILDREN	109
	5.7		ATIONS WITH POTENTIALLY HIGH EXPOSURES	
	5.8		JACY OF THE DATABASE	
		5.8.1	Identification of Data Needs	
		5.8.2	Ongoing Studies	114
6.	ANA	LYTICAI	L METHODS	
	6.1	BIOLOG	GICAL SAMPLES	115
	6.2	ENVIRO	ONMENTAL SAMPLES	115
	6.3	ADEQU	JACY OF THE DATABASE	121
		6.3.1	Identification of Data Needs	122
		6.3.2	Ongoing Studies	123
7.	REGU	JLATION	NS AND ADVISORIES	125
8.	REFE	ERENCES	S	129
a	GI O	SSARY		145

# APPENDICES

A.	ATSDR MINIMAL RISK LEVELS AND WORKSHEETS	<b>A-</b> 1
B.	USER'S GUIDE	B-1
C.	ACRONYMS, ABBREVIATIONS, AND SYMBOLS	C-:

# **LIST OF FIGURES**

2-1	Levels of Significant Exposure to Chloroethane—Inhalation	25
2-2	Metabolic Pathways of Chloroethane Biotransformation	54
2-3	Conceptual Representation of a Physiologically Based Pharmacokinetic (PBPK) Model for a Hypothetical Chemical Substance	58
2-4	Existing Information on Health Effects of Chloroethane	82
5-1	Frequency of NPL Sites with Chloroethane Contamination	100

	•		
•			

# LIST OF TABLES

2-1	Levels of Significant Exposure to Chloroethane—Innalation
2-2	Genotoxicity of Chloroethane In Vivo
2-3	Genotoxicity of Chloroethane In Vitro
3-1	Chemical Identity of Chloroethane 92
3-2	Physical and Chemical Properties of Chloroethane
4-1	Facilities that Manufacture or Process Chloroethane
5-1	Releases to the Environment from Facilities that Manufacture or Process Chloroethane
6-1	Analytical Methods for Determining Chloroethane in Biological Samples
6-2	Analytical Methods for Determining Chloroethane in Environmental Samples
7-1	Regulations and Guidelines Applicable to Chloroethane