TOXICOLOGICAL PROFILE FOR BROMOFORM AND DIBROMOCHLOROMETHANE

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

i

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 Bromoform and Dibromochloromethane, Draft for Public Comment was released in September 2003. This edition supersedes any previously released draft or final profile.

Toxicological profiles are revised and republished as necessary. 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
Mailstop F-32
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.

Julie Louise Gerberding M.D

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 7, 2003 (68 FR 63098). For prior versions of the list of substances, see *Federal Register* notices dated 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); February 28, 1994 (59 FR 9486); April 29, 1996 (61 FR 18744); November 17, 1997 (62 FR 61332); October 21, 1999(64 FR 56792) and October 25, 2001 (66 FR 54014). Section 104(i)(3) of CERCLA, as amended, directs the Administrator of ATSDR to prepare a toxicological profile for each substance on the list.

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: Relevance to Public Health**: The Relevance to Public Health Section evaluates, interprets, and assesses the significance of toxicity data to human health.
- **Chapter 3: Health Effects**: Specific health effects of a given hazardous compound are reported by type of health effect (death, systemic, immunologic, reproductive), by route of exposure, 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 3.7 Children's Susceptibility

Section 6.6 Exposures of Children

Other Sections of Interest:

Section 3.8 Biomarkers of Exposure and Effect Section 3.11 Methods for Reducing Toxic Effects

ATSDR Information Center

Phone: 1-888-42-ATSDR or (404) 498-0110 **Fax:** (770) 488-4178

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 Center for 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@AOEC.ORG • Web Page: http://www.aoec.org/.

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, 25 Northwest Point Boulevard, Suite 700, Elk Grove Village, IL 60007-1030 • Phone: 847-818-1800 • FAX: 847-818-9266.

CONTRIBUTORS

CHEMICAL MANAGER(S)/AUTHOR(S):

John Risher, Ph.D. Dennis Jones, D.V.M. ATSDR, Division of Toxicology, Atlanta, Georgia

Michael H. Lumpkin, Ph.D.
Lisa Ingerman, Ph.D., DABT
Daniel Plewak, M.S.
Lori Moilanen, Ph.D., DABT
Dolores Beblo, Ph.D.
Jennifer Walters, M.S., M.P.A.
Syracuse Research Corporation, North Syracuse, New York

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.
- 4. Green Border Review. Green Border review assures the consistency with ATSDR policy.

PEER REVIEW

A peer review panel was assembled for bromoform and dibromochloromethane. The panel consisted of the following members:

- 1. Bruce Jarnot, Ph.D., DABT, Senior Toxicologist, American Petroleum Institute, Washington DC;
- 2. Kannan Krishnan, Ph.D., Professor, Human Toxicology Research Group, University of Montreal Montreal Canada; and
- 3. Clint Skinner, Ph.D., Consultant, Skinner Associates, Creston California

These experts collectively have knowledge of bromoform and dibromochloromethane'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(I)(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.

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

DISCLA	.IMER	ii
UPDAT	E STATEMENT	iii
	ORD	
	REFERENCE FOR HEALTH CARE PROVIDERS	
CONTR	IBUTORS	ix
	EVIEW	
CONTE	NTS	xiii
	F FIGURES	
LIST O	TABLES	xix
1. PUB	LIC HEALTH STATEMENT	1
1.1	WHAT ARE BROMOFORM AND DIBROMOCHLOROMETHANE?	1
1.2	WHAT HAPPENS TO BROMOFORM AND DIBROMOCHLOROMETHANE WHEN THEY ENTER THE ENVIRONMENT?	Į 3
1.3	HOW MIGHT I BE EXPOSED TO BROMOFORM AND	
1.0	DIBROMOCHLOROMETHANE?	3
1.4	HOW CAN BROMOFORM AND DIBROMOCHLOROMETHANE ENTER AND LEAVE MY BODY?	
1.5	HOW CAN BROMOFORM AND DIBROMOCHLOROMETHANE AFFECT MY HEALTH?	
1.6	HOW CAN BROMOFORM AND DIBROMOCHLOROMETHANE AFFECT CHILDREN?	
1.7	HOW CAN FAMILIES REDUCE THE RISK OF EXPOSURE TO BROMOFORM AN DIBROMOCHLOROMETHANE?	D
1.8	IS THERE A MEDICAL TEST TO DETERMINE WHETHER I HAVE BEEN	
	EXPOSED TO BROMOFORM AND DIBROMOCHLOROMETHANE?	7
1.9	WHAT RECOMMENDATIONS HAS THE FEDERAL GOVERNMENT MADE TO	
	PROTECT HUMAN HEALTH?	
1.10	WHERE CAN I GET MORE INFORMATION?	8
2. RELI	EVANCE TO PUBLIC HEALTH	11
2.1	BACKGROUND AND ENVIRONMENTAL EXPOSURES TO BROMOFORM AND	
	DIBROMOCHLOROMETHANE IN THE UNITED STATES	
2.2	SUMMARY OF HEALTH EFFECTS	
2.3	MINIMAL RISK LEVELS (MRLs)	
3 HEA	LTH EFFECTS	21
3.1	INTRODUCTION	
3.2	DISCUSSION OF HEALTH EFFECTS BY ROUTE OF EXPOSURE	
3.2.		
3	2.1.1 Death	
3	2.1.2 Systemic Effects	
3	2.1.3 Immunological and Lymphoreticular Effects	
3	2.1.4 Neurological Effects	
3	2.1.5 Reproductive Effects	
3	2.1.6 Developmental Effects	24
3	2.1.7 Cancer.	24

3.2.2	Oral Exposure	24
3.2.	2.1 Death	24
3.2.	2.2 Systemic Effects	56
3.2.	2.3 Immunological and Lymphoreticular Effects	63
3.2.	2.4 Neurological Effects	63
3.2.	2.5 Reproductive Effects	65
3.2.	2.6 Developmental Effects	67
3.2.	2.7 Cancer	68
3.2.3	Dermal Exposure	
3.2.		
3.2.	√	
3.2		
3.2.	c	
3.2.	1	
3.2.	1	
3.2.		
	GENOTOXICITY	
	TOXICOKINETICS	
3.4.1	Absorption	
3.4.	r	
3.4. 3.4.	1	
3.4.2	Distribution	
3.4.2		
3.4.	<u>*</u>	
3.4.	1	
3.4.3	Metabolism	
3.4.4	Elimination and Excretion.	
3.4.		
3.4.	•	
3.4.	<u>*</u>	
3.4.5	Physiologically Based Pharmacokinetic (PBPK)/Pharmacodynamic (PD) Models	
3.5	MECHANISMS OF ACTION	
3.5.1	Pharmacokinetic Mechanisms	91
3.5.2	Mechanisms of Toxicity	91
3.5.3	Animal-to-Human Extrapolations	94
	TOXICITIES MEDIATED THROUGH THE NEUROENDOCRINE AXIS	
	CHILDREN'S SUSCEPTIBILITY	
3.8	BIOMARKERS OF EXPOSURE AND EFFECT	98
3.8.1	Biomarkers Used to Identify or Quantify Exposure to Bromoform and	
	Dibromochloromethane	99
3.8.2	Biomarkers Used to Characterize Effects Caused by Bromoform and	
	Dibromochloromethane	
	INTERACTIONS WITH OTHER CHEMICALS	
	POPULATIONS THAT ARE UNUSUALLY SUSCEPTIBLE	
	METHODS FOR REDUCING TOXIC EFFECTS	
3.11.1		
3.11.2		
3.11.3	\mathcal{C}	
3.12 3.12.1	ADEQUACY OF THE DATABASE Existing Information on Health Effects of Bromoform and Dibromochloromethane	
	LAISUUZ HIIOHIIAUOI OII FICAIUI EFICUS OI DIOHIOIOHII AHU DIDIOHIOCHIOIOHICHIAHC	11/2

4. CHEMICAL AND PHYSICAL INFORMATION		3.12.2 3.12.3		
4.1 CHEMICAL IDENTITY 117 4.2 PHYSICAL AND CHEMICAL PROPERTIES 117 5. PRODUCTION. IMPORT/EXPORT, USE, AND DISPOSAL 121 5.1 PRODUCTION 121 5.2 IMPORT/EXPORT 122 5.3 USE 122 5.4 DISPOSAL 122 6. POTENTIAL FOR HUMAN EXPOSURE 125 6.1 OVERVIEW 125 6.2 RELEASES TO THE ENVIRONMENT 125 6.2.1 Air 128 6.2.2 Water 130 6.3.3 SOIL 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Vater 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 136 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 <td></td> <td>0.12.0</td> <td>0.180.1180.0000000000000000000000000000</td> <td></td>		0.12.0	0.180.1180.0000000000000000000000000000	
4.2 PHYSICAL AND CHEMICAL PROPERTIES 117 5. PRODUCTION, IMPORT/EXPORT, USE, AND DISPOSAL 121 5.1 PRODUCTION 121 5.2 IMPORTEXPORT 122 5.3 USE 122 5.4 DISPOSAL 122 6. POTENTIAL FOR HUMAN EXPOSURE 125 6.1 OVERVIEW 125 6.2 RELEASES TO THE ENVIRONMENT 125 6.2.1 Air 128 6.2.2 Water 130 6.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transport and Partitioning 134 6.3.2.1 Air 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.1 Air 135 6.3.2.2 Water 136 6.3.2.4 Other Media 137 6.4.1 Air Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 149	4.	CHEM	ICAL AND PHYSICAL INFORMATION	117
5. PRODUCTION, IMPORT/EXPORT, USE, AND DISPOSAL 121 5.1 PRODUCTION 121 5.2 IMPORT/EXPORT 122 5.3 USE 122 5.4 DISPOSAL 122 6. POTENTIAL FOR HUMAN EXPOSURE 125 6.1 OVERVIEW 125 6.2.1 Air 125 6.2.1 Air 128 6.2.2 Water 130 6.3.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transport and Partitioning 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8. ADEQUACY OF THE DATABAS				
5.1 PRODUCTION 121 5.2 IMPORT/EXPORT 122 5.3 USE 122 5.4 DISPOSAL 122 6. POTENTIAL FOR HUMAN EXPOSURE 125 6.1 OVERVIEW 125 6.2 RELEASES TO THE ENVIRONMENT 125 6.2.1 Air 125 6.2.2 Water 130 6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transformation and Partitioning 134 6.3.2.1 Air 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE		4.2	PHYSICAL AND CHEMICAL PROPERTIES	117
5.1 PRODUCTION 121 5.2 IMPORT/EXPORT 122 5.3 USE 122 5.4 DISPOSAL 122 6. POTENTIAL FOR HUMAN EXPOSURE 125 6.1 OVERVIEW 125 6.2 RELEASES TO THE ENVIRONMENT 125 6.2.1 Air 125 6.2.2 Water 130 6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transformation and Partitioning 134 6.3.2.1 Air 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE	5	PRODI	ICTION IMPORT/EXPORT USE AND DISPOSAL	121
5.2 IMPORT/EXPORT 122 5.3 USE. 122 5.4 DISPOSAL 122 6. POTENTIAL FOR HUMAN EXPOSURE 125 6.1 OVERVIEW. 125 6.2 RELEASES TO THE ENVIRONMENT 125 6.2.1 Air 128 6.2.2 Water 130 6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE. 134 6.3.1 Transformation and Partitioning 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air. 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil. 136 6.3.2.4 Other Media. 137 6.4.1 Air 137 6.4.2 Water. 137 6.4.3 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.4 User Control Media. 149 6.4.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES SOF CHILDREN 146 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 </td <td>٠.</td> <td></td> <td></td> <td></td>	٠.			
5.3 USE. 122 5.4 DISPOSAL 122 6. POTENTIAL FOR HUMAN EXPOSURE 125 6.1 OVERVIEW 125 6.2 RELEASES TO THE ENVIRONMENT 125 6.2.1 Air 128 6.2.2 Water 130 6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transformation and Partitioning 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.4.2 Water 139 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL				
5.4 DISPOSAL 122 6. POTENTIAL FOR HUMAN EXPOSURE 125 6.1 OVERVIEW 125 6.2 RELEASES TO THE ENVIRONMENT 125 6.2.1 Air 128 6.2.2 Water 130 6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transport and Partitioning 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157				
6.1 OVERVIEW 125 6.2 RELEASES TO THE ENVIRONMENT 125 6.2.1 Air 128 6.2.2 Water 130 6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transport and Partitioning 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154				
6.1 OVERVIEW 125 6.2 RELEASES TO THE ENVIRONMENT 125 6.2.1 Air 128 6.2.2 Water 130 6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transport and Partitioning 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154	6	DOTEN	TIAL FOR HIMAN EXPOSURE	125
6.2 RELEASES TO THE ENVIRONMENT 125 6.2.1 Air 128 6.2.2 Water 130 6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transport and Partitioning 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7.3 ADEQUACY OF THE DATABASE 157	Ο.			
6.2.1 Air 128 6.2.2 Water 130 6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transport and Partitioning 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5.4 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 146 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS				
6.2.2 Water 130 6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transformation and Partitioning 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158<				
6.2.3 Soil 133 6.3 ENVIRONMENTAL FATE 134 6.3.1 Transport and Partitioning 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 A				
6.3 ENVIRONMENTAL FATE. 134 6.3.1 Transport and Partitioning. 134 6.3.2 Transformation and Degradation. 135 6.3.2.1 Air. 135 6.3.2.2 Water. 135 6.3.2.3 Sediment and Soil. 136 6.3.2.4 Other Media. 137 6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT. 137 6.4.1 Air. 137 6.4.2 Water. 139 6.4.3 Sediment and Soil. 145 6.4.4 Other Environmental Media. 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE. 146 6.6 EXPOSURES OF CHILDREN. 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES. 150 6.8 ADEQUACY OF THE DATABASE. 151 6.8.1 Identification of Data Needs. 151 6.8.2 Ongoing Studies. 157 7.1 BIOLOGICAL MATERIALS. 157 7.2 ENVIRONMENTAL SAMPLES. 158 7.3 ADEQUACY OF THE DATABASE. 160				
6.3.1 Transformation and Degradation 134 6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 158 7.3.1 Identification of Data Needs 160 <				
6.3.2 Transformation and Degradation 135 6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171				
6.3.2.1 Air 135 6.3.2.2 Water 135 6.3.2.3 Sediment and Soil 136 6.3.2.4 Other Media 137 6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 146 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9.				
6.3.2.3 Sediment and Soil		6.3.	e	
6.3.2.3 Sediment and Soil		6.3.	2.2 Water	135
6.4 LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT 137 6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8 REGULATIONS AND ADVISORIES 165 9 REFERENCES 171		6.3.	2.3 Sediment and Soil	136
6.4.1 Air 137 6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171		6.3.	2.4 Other Media	137
6.4.2 Water 139 6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171		6.4	LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT	137
6.4.3 Sediment and Soil 145 6.4.4 Other Environmental Media 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171		6.4.1	Air	137
6.4.4 Other Environmental Media. 145 6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171		6.4.2	Water	139
6.5 GENERAL POPULATION AND OCCUPATIONAL EXPOSURE 146 6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171		6.4.3	Sediment and Soil	145
6.6 EXPOSURES OF CHILDREN 149 6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171		6.4.4		
6.7 POPULATIONS WITH POTENTIALLY HIGH EXPOSURES 150 6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171		6.5		
6.8 ADEQUACY OF THE DATABASE 151 6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171		6.6		
6.8.1 Identification of Data Needs 151 6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171				
6.8.2 Ongoing Studies 154 7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171				
7. ANALYTICAL METHODS 157 7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171				
7.1 BIOLOGICAL MATERIALS 157 7.2 ENVIRONMENTAL SAMPLES 158 7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171		6.8.2	Ongoing Studies	154
7.2 ENVIRONMENTAL SAMPLES	7.	ANAL	YTICAL METHODS	157
7.3 ADEQUACY OF THE DATABASE 160 7.3.1 Identification of Data Needs 160 7.3.2 Ongoing Studies 163 8. REGULATIONS AND ADVISORIES 165 9. REFERENCES 171		7.1	BIOLOGICAL MATERIALS	157
7.3.1 Identification of Data Needs		7.2	ENVIRONMENTAL SAMPLES	158
7.3.2 Ongoing Studies		7.3	ADEQUACY OF THE DATABASE	160
8. REGULATIONS AND ADVISORIES		7.3.1	Identification of Data Needs	160
9. REFERENCES		7.3.2	Ongoing Studies	163
	8.	REGUI	LATIONS AND ADVISORIES	165
10. GLOSSARY	9.	REFER	ENCES	171
	10). GLOS	SARY	217

APPENDICES

A. ATSDR MINIMAL RISK LEVELS AND WORKSHEETS	A-1
B. USER'S GUIDE	B-
C. ACRONYMS, ABBREVIATIONS, AND SYMBOLS	
D. INDEX	D-

LIST OF FIGURES

3-1. Levels of Significant Exposure to Bromoform—Oral	35
3-2. Levels of Significant Exposure to Dibromochloromethane—Oral	51
3-3. Proposed Pathway of Trihalomethane Metabolism in Rats	83
3-4. Proposed Pathway of Trihalomethyl-radical-mediated Lipid Peroxidation	85
3-5. Conceptual Representation of a Physiologically Based Pharmacokinetic (PBPK) Mode Hypothetical Chemical Substance	
3-6. Existing Information on Health Effects of Bromoform	105
3-7. Existing Information on Health Effects of Dibromochloromethane	106
6-1. Frequency of NPL Sites with Bromoform Contamination	126
6-2. Frequency of NPL Sites with Dibromochloromethane Contamination	127

LIST OF TABLES

3-1.	Levels of Significant Exposure to Bromoform—Oral	25
3-2.	Levels of Significant Exposure to Dibromochloromethane—Oral	39
3-3.	Genotoxicity of Bromoform In Vivo	71
3-4.	Genotoxicity of Bromoform In Vitro	72
3-5.	Genotoxicity of Dibromochloromethane In Vivo	75
3-6.	Genotoxicity of Dibromochloromethane In Vitro	76
3-7.	Partition Coefficients for Bromoform and Dibromochloromethane	80
4-1.	Chemical Identity of Bromoform and Dibromochloromethane	118
4-2.	Physical and Chemical Properties of Bromoform and Dibromochloromethane	119
5-1.	Facilities that Produce, Process, or Use Bromoform	123
6-1.	Releases to the Environment from Facilities that Produce, Process, or Use Bromoform	129
6-2.	Occurrence of Bromoform and Dibromochloromethane in Finished Drinking Water	140
6-3.	Summary of Typical Human Exposure to Bromoform and Dibromochloromethane	147
6-4.	Ongoing Studies on the Potential for Human Exposure to Bromoform	155
7-1.	Analytical Methods for Determining Bromoform and Dibromochloromethane in Biological Materials	159
7-2.	Analytical Methods for Determining Bromoform and Dibromochloromethane in Environmental Samples	161
8-1.	Regulations and Guidelines Applicable to Bromoform and Dibromochloromethane	167