TOXICOLOGICAL PROFILE FOR TITANIUM TETRACHLORIDE

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

September 1997

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UPDATE STATEMENT

A Toxicological Profile for Titanium Tetrachloride was released on June 1994. 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:

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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.

David Satcher, M.D., Ph.D. 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 Super-fund). Section 211 of SARA also amended Title 10 of the U. S. Code, creating the Defense Environmental Restoration Program. Section 2704(a) of Title 10 of the U. S. Code directs the Secretary of Defense to notify the Secretary of Health and Human Services of not less than 25 of the most commonly found unregulated hazardous substances at defense facilities. Section 2704(b) of Title 10 of the U. S. Code directs the Administrator of the Agency for Toxic Substances and Disease Registry (ATSDR) to prepare a toxicological profile for each substance on the list provided by the Secretary of Defense under subsection (b).

CONTRIBUTORS

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THE PROFILE HAS UNDERGONE THE FOLLOWING ATSDR INTERNAL REVIEWS:

- 1. Green Border Review. Green Border review assures consistency with ATSDR policy.
- 2. 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.
- 3. 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.
- 4. Quality Assurance Review. The Quality Assurance Branch assures that consistency across profiles is maintained, identifies any significant problems in format or content, and establishes that Guidance has been followed.

PEER REVIEW

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

1. Dr. Hugh Farber, Private Consultant, Lake Leelanau, MI;

2. Dr. Arthur Gregory, Private Consultant, Sterling, VA; and

3. Mr. Lyman Skory, Skory Consulting, Midland, MI.

These experts collectively have knowledge of titanium tetrachloride'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. 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
CONTR	IBUTORS		/ii
PEER R	EVIEW		ix
LIST OI	F FIGURES	· · · · · · · · · · · · · · · · · · ·	٢V
LIST OI	TABLES	xv	/ii
1. PUB	LIC HEALTH ST	ATEMENT	1
1.1	WHAT IS TITA	NIUM TETRACHLORIDE?	2
1.2	WHAT HAPPEN	NS TO TITANIUM TETRACHLORIDE WHEN IT ENTERS THE	
	ENVIRONMEN	T?	2
1.3	HOW MIGHT I	BE EXPOSED TO TITANIUM TETRACHLORIDE?	3
1.4	HOW CAN TIT	ANIUM TETRACHLORIDE ENTER AND LEAVE MY BODY?	3
1.5	HOW CAN TIT	ANIUM TETRACHLORIDE AFFECT MY HEALTH?	3
1.6	IS THERE A M	EDICAL TEST TO DETERMINE WHETHER I HAVE BEEN	
	EXPOSED TO 7	ITANIUM TETRACHLORIDE?	5
1.7	WHAT RECOM	MENDATIONS HAS THE FEDERAL GOVERNMENT MADE TO	
	PROTECT HUN	1AN HEALTH?	5
1.8	WHERE CAN I	GET MORE INFORMATION?	6
2 HE/	ATTH FFFFCTS		7
2. 11127			7
2.1		F HEALTH EFFECTS BY ROUTE OF EXPOSURE	
2.2		on Exposure	
	2.2.1		10
	2.2.1.2		11
	2.2.1.3		25
	2.2.1.4		27
	2.2.1.5	0	27
	2.2.1.6		27
	2.2.1.7		28
	2.2.1.8		28
	2.2.2 Oral Ex		30
	2.2.2.1	E - The second se	30
	2.2.2.2		30
	2.2.2.3		30
	2.2.2.4	v v i	30
	2.2.2.5	8	30
	2.2.2.6	L Contraction of the second seco	30
•	2.2.2.7	1	30
	2.2.2.8		30

		2.2.3	Dermal E	Exposure	30
			2.2.3.1	Death	31
			2.2.3.2		31
			2.2.3.3	· ·	34
			2.2.3.4	÷ • •	35
			2.2.3.5		35
			2.2.3.6		35
			2.2.3.7	1	35
			2.2.3.7		35
	2.3	TOVIC	2.2.3.8 OKINETIO		35 36
	2.5	2.3.1			30 37
		2.5.1	Absorptio		
			2.3.1.1	1	37
			2.3.1.2	•	37
			2.3.1.3	▲	37
		2.3.2			37
			2.3.2.1	-	37
			2.3.2.2	1	38
			2.3.2.3	*	38
		2.3.3	Metaboli		38
		2.3.4	Eliminati	on and Excretion	38
	2.4	MECHA	ANISMS (DF ACTION	39
	2.5	RELEV	ANCE TO	PUBLIC HEALTH	40
	2.6	BIOMA	RKERS C	OF EXPOSURE AND EFFECT	53
		2.6.1	Biomarke	ers Used to Identify or Quantify Exposure to Titanium Tetrachloride	54
		2.6.2			54
	2.7	INTERA			55
	2.8	POPUL	ATIONS 7	THAT ARE UNUSUALLY SUSCEPTIBLE	55
	2.9				56
		2.9.1			56
		2.9.2	-		56
		2.9.3			57
	2 10				57
	2.10	2.10.1			57
		2.10.1	0		59
					65
		2.10.5	Ongoing	Studies	05
3	CHEI	MICAI		SICAL INFORMATION	67
5.	3.1				67
					67
	3.2	PHISI	AL AND	CHEMICAL PROPERTIES	07
4	הסממ			TEXPORT LICE AND DISDOCAL	771
4.		-			71
	4.1				71
	4.2				71
	4.3		••••••		71
	4.4	DISPOS	SAL		74
5.	POTE				75
	5.1	OVERV	IEW		75

	5.2	5.2.1 Air	76 76
			76
			76
	5.3		79
			79
			79
		5.3.2.1 Air	79
		5.3.2.2 Water	79
		5.3.2.3 Sediment and Soil	79
	5.4	LEVELS MONITORED OR ESTIMATED IN THE ENVIRONMENT	80
		5.4.1 Air	80
		5.4.2 Water	80
			80
			80
	5.5		80
	5.6		81
	5.7		82
	0.1		82
			85
		5.7.2 Ongoing Studies	05
6.	ANA	LYTICAL METHODS	87
	6.1	BIOLOGICAL SAMPLES	87
	6.2	ENVIRONMENTAL SAMPLES	89
	6.3	ADEQUACY OF THE DATABASE	91
		6.3.1 Identification of Data Needs	92
			93
7.	REG	ULATIONS AND ADVISORIES	95
8.	REF	ERENCES	97
9.	GLO	SSARY 1	07

APPENDICES

A.	MINIMAL RISK LEVEL (MRL) WORKSHEETS	A-1
B.	USER'S GUIDE	B-1
C.	ACRONYMS, ABBREVIATIONS, AND SYMBOLS	C-1

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LIST OF FIGURES

2-1	Levels of Significant Exposure to Titanium Tetrachloride - Inhalation	15
2-2	Existing Information on Health Effects of Titanium Tetrachloride	58

• • •

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LIST OF TABLES

2-1	Levels of Significant Exposure to Titanium Tetrachloride - Inhalation	12
2-2	Genotoxicity of Titanium Tetrachloride In Vitro	51
3-1	Chemical Identity of Titanium Tetrachloride	68
3-2	Physical and Chemical Properties of Titanium Tetrachloride	69
4-1	Facilities That Manufacture or Process Titanium Tetrachloride	72
5-1	Releases to the Environment from Facilities That Manufacture or Process Titanium Tetrachloride	77
6-1	Analytical Methods for Determining Titanium and Titanium Dioxide in Biological Materials	88
6-2	Analytical Methods for Determining Titanium and Titanium Dioxide in Environmental Samples	90
7-1	Regulations and Guidelines Applicable to Titanium Tetrachloride	96