

Provisional Peer Reviewed Toxicity Values for
Endrin aldehyde
(CASRN 7421-93-4)

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Acronyms and Abbreviations

bw	body weight
cc	cubic centimeters
CD	Caesarean Delivered
CERCLA	Comprehensive Environmental Response, Compensation and Liability Act of 1980
CNS	central nervous system
cu.m	cubic meter
DWEL	Drinking Water Equivalent Level
FEL	frank-effect level
FIFRA	Federal Insecticide, Fungicide, and Rodenticide Act
g	grams
GI	gastrointestinal
HEC	human equivalent concentration
Hgb	hemoglobin
i.m.	intramuscular
i.p.	intraperitoneal
IRIS	Integrated Risk Information System
IUR	inhalation unit risk
i.v.	intravenous
kg	kilogram
L	liter
LEL	lowest-effect level
LOAEL	lowest-observed-adverse-effect level
LOAEL(ADJ)	LOAEL adjusted to continuous exposure duration
LOAEL(HEC)	LOAEL adjusted for dosimetric differences across species to a human
m	meter
MCL	maximum contaminant level
MCLG	maximum contaminant level goal
MF	modifying factor
mg	milligram
mg/kg	milligrams per kilogram
mg/L	milligrams per liter
MRL	minimal risk level
MTD	maximum tolerated dose
MTL	median threshold limit
NAAQS	National Ambient Air Quality Standards
NOAEL	no-observed-adverse-effect level
NOAEL(ADJ)	NOAEL adjusted to continuous exposure duration
NOAEL(HEC)	NOAEL adjusted for dosimetric differences across species to a human
NOEL	no-observed-effect level
OSF	oral slope factor
p-IUR	provisional inhalation unit risk
p-OSF	provisional oral slope factor
p-RfC	provisional inhalation reference concentration

p-RfD	provisional oral reference dose
PBPK	physiologically based pharmacokinetic
ppb	parts per billion
ppm	parts per million
PPRTV	Provisional Peer Reviewed Toxicity Value
RBC	red blood cell(s)
RCRA	Resource Conservation and Recovery Act
RDDR	Regional deposited dose ratio (for the indicated lung region)
REL	relative exposure level
RfC	inhalation reference concentration
RfD	oral reference dose
RGDR	Regional gas dose ratio (for the indicated lung region)
s.c.	subcutaneous
SCE	sister chromatid exchange
SDWA	Safe Drinking Water Act
sq.cm.	square centimeters
TSCA	Toxic Substances Control Act
UF	uncertainty factor
µg	microgram
µmol	micromoles
VOC	volatile organic compound

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Background

On December 5, 2003, the U.S. Environmental Protection Agency's (EPA's) Office of Superfund Remediation and Technology Innovation (OSRTI) revised its hierarchy of human health toxicity values for Superfund risk assessments, establishing the following three tiers as the new hierarchy:

1. EPA's Integrated Risk Information System (IRIS).
2. Provisional Peer-Reviewed Toxicity Values (PPRTV) used in EPA's Superfund Program.
3. Other (peer-reviewed) toxicity values, including:
 - ▶ Minimal Risk Levels produced by the Agency for Toxic Substances and Disease Registry (ATSDR),
 - ▶ California Environmental Protection Agency (CalEPA) values, and
 - ▶ EPA Health Effects Assessment Summary Table (HEAST) values.

A PPRTV is defined as a toxicity value derived for use in the Superfund Program when such a value is not available in EPA's Integrated Risk Information System (IRIS). PPRTVs are developed according to a Standard Operating Procedure (SOP) and are derived after a review of the relevant scientific literature using the same methods, sources of data, and Agency guidance for value derivation generally used by the EPA IRIS Program. All provisional toxicity values receive internal review by two EPA scientists and external peer review by three independently selected scientific experts. PPRTVs differ from IRIS values in that PPRTVs do not receive the multi-program consensus review provided for IRIS values. This is because IRIS values are generally intended to be used in all EPA programs, while PPRTVs are developed specifically for the Superfund Program.

Because new information becomes available and scientific methods improve over time, PPRTVs are reviewed on a five-year basis and updated into the active database. Once an IRIS value for a specific chemical becomes available for Agency review, the analogous PPRTV for that same chemical is retired. It should also be noted that some PPRTV manuscripts conclude that a PPRTV cannot be derived based on inadequate data.

Disclaimers

Users of this document should first check to see if any IRIS values exist for the chemical of concern before proceeding to use a PPRTV. If no IRIS value is available, staff in the regional Superfund and RCRA program offices are advised to carefully review the information provided in this document to ensure that the PPRTVs used are appropriate for the types of exposures and circumstances at the Superfund site or RCRA facility in question. PPRTVs are periodically updated; therefore, users should ensure that the values contained in the PPRTV are current at the time of use.

It is important to remember that a provisional value alone tells very little about the adverse effects of a chemical or the quality of evidence on which the value is based. Therefore, users are strongly encouraged to read the entire PPRTV manuscript and understand the strengths and limitations of the derived provisional values. PPRTVs are developed by the EPA Office of Research and Development's National Center for Environmental Assessment, Superfund Health Risk Technical Support Center for OSRTI. Other EPA programs or external parties who may choose of their own initiative to use these PPRTVs are advised that Superfund resources will not generally be used to respond to challenges of PPRTVs used in a context outside of the Superfund Program.

Questions Regarding PPRTVs

Questions regarding the contents of the PPRTVs and their appropriate use (e.g., on chemicals not covered, or whether chemicals have pending IRIS toxicity values) may be directed to the EPA Office of Research and Development's National Center for Environmental Assessment, Superfund Health Risk Technical Support Center (513-569-7300), or OSRTI.

INTRODUCTION

Except as noted, literature searches were conducted from the 1960s through December 2007 for studies relevant to the derivation of provisional toxicity values for endrin aldehyde. Databases searched include MEDLINE, TOXLINE (Special), BIOSIS (from August 2000), TSCATS 1/TSCATS 2, CCRIS, DART/ETIC, GENETOX, HSDB, RTECS, and Current Contents (July through December 2007).

This document has passed the STSC quality review and evaluation indicating that the quality is consistent with the SOPs and standards of the STSC and is suitable for use by registered users of the PPRTV system.

No RfD, RfC, or carcinogenicity assessment for endrin aldehyde is available on IRIS (U.S. EPA, 2008), in the Drinking Water Standards and Health Advisories list (U.S. EPA, 2006), or in the HEAST (U.S. EPA, 1997). The CARA list (U.S. EPA, 1991, 1994) does not include any documents for endrin aldehyde either. Documents listed for endrin include a Health Effects Assessment (U.S. EPA, 1987) and a Drinking Water Criteria Document (U.S. EPA, 1992) neither of which contain chronic or subchronic toxicity data for endrin aldehyde. The ATSDR Toxicological Profile for endrin (ATSDR, 1996) includes endrin aldehyde, but it does not contain any chronic or subchronic toxicity data. The chronic toxicity and carcinogenicity of endrin aldehyde were not assessed by IARC (2008) or NTP (2008), and endrin aldehyde is not included in the 11th Report on Carcinogens (NTP, 2005). An Environmental Health Criteria document (WHO, 1992) for endrin was consulted for relevant information. Literature searches were conducted from the 1960s through December 2007 for studies relevant to the derivation of provisional toxicity values for endrin aldehyde. Databases searched include MEDLINE, TOXLINE (Special), BIOSIS, TSCATS/TSCATS 2, CCRIS, DART/ETIC, GENETOX, HSDB, RTECS, and Current Contents.

FEASIBILITY OF DERIVING A PROVISIONAL RfD FOR ENDRIN ALDEHYDE

The data are inadequate to derive a p-RfD for endrin aldehyde. No dose-response information pertinent to any target organs is available in the current database; thus, the database lacks a study that could serve as a suitable basis for the derivation of a p-RfD for endrin aldehyde.

FEASIBILITY OF DERIVING A PROVISIONAL RfC FOR ENDRIN ALDEHYDE

No inhalation toxicity data in humans or animals is identified; thus, no p-RfC could be derived for endrin aldehyde.

PROVISIONAL CARCINOGENICITY ASSESSMENT FOR ENDRIN ALDEHYDE

Because of the lack of carcinogenic data in humans or animals, under the 2005 *Guidelines for Carcinogen Risk Assessment* (U.S. EPA, 2005), this PPRTV document classifies endrin aldehyde as having “*Inadequate Information to Assess Carcinogenic Potential.*”

FEASIBILITY OF DERIVING A PROVISIONAL ORAL SLOPE FACTOR OR INHALATION UNIT RISK FOR ENDRIN ALDEHYDE

Neither a p-OSF nor a p-IUR could be derived for endrin aldehyde because of the lack of suitable oral or inhalation data in both humans and animals.

REFERENCES

ATSDR (Agency for Toxic Substances and Disease Registry). 1996. Toxicological Profile for Endrin. Update. August 1996. ATSDR, U.S. Public Health Service, Atlanta, GA.

IARC (International Agency for Research on Cancer). 2008. Search IARC Monographs. Online. <http://monographs.iarc.fr/ENG/Monographs/allmonos90.php>.

NTP (National Toxicology Program). 2005. 11th Report on Carcinogens. U.S. Department of Health and Human Services, Public Health Service, National Institutes of Health, Research Triangle Park, NC. Available at <http://ntp-server.niehs.nih.gov/>.

NTP (National Toxicology Program). 2008. Management Status Report. Online. <http://ntp.niehs.nih.gov/index.cfm?objectid=78CC7E4C-F1F6-975E-72940974DE301C3F>.

U.S. EPA. 1987. Health Effects Assessment for Endrin. Prepared by the Office of Health and Environmental Assessment, Environmental Criteria and Assessment Office, Cincinnati, OH for the Office of Emergency and Remedial Response, Washington, DC.

U.S. EPA. 1991. Chemical Assessments and Related Activities (CARA). Office of Health and Environmental Assessment, Washington, DC.

U.S. EPA. 1992. Drinking Water Criteria Document for Endrin. Prepared by the Office of Health and Environmental Assessment, Environmental Criteria and Assessment Office, Cincinnati, OH for the Office of Water, Washington, DC. NTIS PB92-173384.

U.S. EPA. 1994. Chemical Assessments and Related Activities (CARA). Office of Health and Environmental Assessment, Washington, DC. December.

U.S. EPA. 1997. Health Effects Assessment Summary Tables. FY-1997 Update. Prepared by the Office of Research and Development, National Center for Environmental Assessment, Cincinnati OH for the Office of Emergency and Remedial Response, Washington, DC. July. EPA/540/R-97/036. NTIS PB97-921199.

U.S. EPA. 2005. Guidelines for Carcinogen Risk Assessment. Risk Assessment Forum, Washington, DC; EPA/630/P-03/001F. Federal Register 70(66):17765--17817. Available online at <http://www.epa.gov/raf>.

U.S. EPA. 2006. 2006 Edition of the Drinking Water Standards and Health Advisories. Office of Water, Washington, DC. EPA 822-R-06-013. Washington, DC. <http://www.epa.gov/waterscience/drinking/standards/dwstandards.pdf>.

U.S. EPA. 2008. Integrated Risk Information System (IRIS). Office of Research and Development, National Center for Environmental Assessment, Washington, DC. Online. <http://www.epa.gov/iris/>.

WHO (World Health Organization). 1992. Endrin. Environmental Health Criteria 130. Geneva, Switzerland.