

Ground Water Quality Standard for 2-Ethyl-1-Hexanol

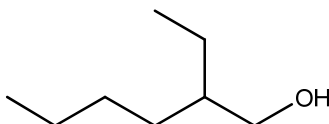
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CASRN# 104-76-7

NJDEP

Summary of Decision: In accordance with the New Jersey Ground Water Quality Standards rules at N.J.A.C. 7:9C-1.7, the Department of Environmental Protection (Department) has developed an interim specific ground water quality criterion of 200 µg/L and PQL of 0.5 µg/L (ppb) for 2-ethyl-1-hexanol. The basis for this criterion and PQL are discussed below. Pursuant to N.J.A.C. 7:9C-1.9(c), **the applicable constituent standard is 200 µg/L.**

2-Ethyl-1-Hexanol
Molecular Formula: C₈H₁₈O
Molecular Structure:



Background: 2-ethyl-1-hexanol is a high production chemical that is widely used yet lacks a complete toxicological database. 2-ethyl-1-hexanol was nominated, but not tested, for carcinogenicity testing by the National Toxicology Program (NTP) because it is a high-volume chemical and a major metabolite of di(2-ethylhexyl) phthalate, which is a known hepatocarcinogen and a known contaminant in blood storage bags (Arneson et al., 1995). 2-ethyl-1-hexanol can be emitted from carpets and some plastics. Occupational or non-occupational standards do not exist for this constituent.

Reference Dose: The long-term F-344 Fischer rat study had a Lowest Observed Adverse Effect Level (LOAEL) of 50 mg/kg/day, adjusted to 35.7 mg/kg/day to account for exposure over 5 days/week, based on a significant dose-related increase in stomach weight. This LOAEL is considered a minimal LOAEL and, as such, calls for an uncertainty factor of 3, rather than the more customary 10, to obtain a No Observed Effect Level (NOEL). The rat study demonstrated statistically significant body weight reductions as early as the first month of dosing, as well as highly statistically significant multiple organ weight changes. As per the U.S. Environmental Protection Agency (USEPA) Guidelines for Carcinogen Risk Assessment (2005), considering the totality of all the evidence available, the Department has classified 2-ethyl-1-hexanol as "Inadequate Evidence to Assess Carcinogen Potential" and has treated it as a non-carcinogen for risk assessment. Based on the adjusted LOAEL of 35.7 mg/kg/day in male rats, the Reference Dose is derived as follows:

Uncertainty factor (UF) adjustment:

UF_{interspecies extrapolation} = 10

UF_{sensitive subpopulations} = 10

$$UF_{\text{conversion from LOAEL to NOEL}} = 3^*$$

$$UF_{\text{database insufficiencies}} = 3^*$$

$$UF = UF_{\text{total}} = 10 \times 10 \times 10 = 1000$$

*Note: Factors of 3 are considered to be 1/2 logs of 10; therefore, the use of 2 factors of 3 is equivalent to one factor of 10 (USEPA, 2002).

$$\begin{aligned} RfD_{\text{oral}} &= \text{NOEL}/UF \\ &= \frac{35.7 \text{ mg/kg/day}}{1000} \end{aligned}$$

$$RfD = 0.0357 \text{ mg/kg/day}$$

Derivation of Ground Water Quality Criterion: The ground water quality criterion was derived pursuant to the formula established at N.J.A.C. 7:9C-1.7(c)4, using 0.0357 mg/kg/day as the Reference Dose (as explained above), and standard default assumptions:

$$\frac{0.0357 \text{ mg/kg/day} \times 70 \text{ kg} \times 0.2}{2 \text{ L/day}} = 0.2499 \text{ mg/L (rounded to 0.2 mg/L)} = \mathbf{200 \mu\text{g/L}}$$

Where:

0.02 mg/kg/day = the derived RfD

70 kg = the assumed weight of an adult human

0.2 = the assumed relative source contribution

2 L/day = the assumed daily volume of water consumed.

Derivation of PQL: The method detection limit (MDL) and the practical quantitation level (PQL) are performance measures used to estimate the limits of performance of analytic chemistry methods for measuring contaminants. The MDL is defined as "the minimum concentration of a substance that can be measured and reported with 99 percent confidence that the analyte concentration is greater than zero" (40 CFR Part 136 Appendix B). USEPA recommends that the MDL be multiplied by a factor of five or 10 to account for the variability and uncertainty that can occur at the MDL. The Department uses a value of five as the median upper boundary of the inter-laboratory MDL distribution from the New Jersey certified laboratory community and multiplies the MDL by five to derive the PQL. Establishing the PQL at a level that is five times the MDL provides a reliable quantitation level that most laboratories can be expected to meet during day-to-day operations.

No published method was listed in the [National Environmental Methods Index \(NEMI\)](#) database for this chemical. A Dialog search located a peer reviewed journal article that contained sufficient performance information to generate a PQL. According to this article, Solid Phase Micro Extraction (SPME) headspace/GC/MS has been used extensively over the past seven years to detect purgable organoleptic compounds that impart an undesirable taste and odor to finished drinking water. 2-ethyl-1-hexanol is a purgable organic compound and performance of this method has been observed down to sub parts-per-billion levels. A method detection limit of 0.1 ppb was reported (Furton, 2003).

As explained above, a more conservative detection limit is established using a multiplier of five. $0.1 \text{ ppb} \times 5 = 0.5 \text{ ppb}$. Therefore, the Department has established a PQL of 0.5 ppb for 2-ethyl-1-hexanol.

Conclusion: Based on the information provided above (and cited below), the Department has established an interim specific ground water quality criterion of 200 $\mu\text{g/L}$ and a PQL of 0.5 $\mu\text{g/L}$ (ppb) for 2-ethyl-1-hexanol. Since the ground water quality criterion is higher than the PQL for this constituent, pursuant to N.J.A.C. 7:9C-1.9(c), **the applicable constituent standard for 2-ethyl-1-hexanol is 200 $\mu\text{g/L}$.**

Technical Support Documents: *Interim Specific Ground Water Quality Criterion Recommendation Report for 2-Ethyl-1-Hexanol*, Dr. Thomas Ledoux, NJDEP, May 2006; *Procedure for Describing Process for Development of Analytical Practical Quantitation Levels (PQLs) for 2-Ethyl-1-Hexanol*, R. Lee Lippincott, Ph.D., NJDEP, February 26, 2003.

References:

Arneson, D. W., Kuhn, G. O., and C. W. Jameson. 1995. Analysis of Feed Blends Containing Microencapsulated 2-Ethyl-1-Hexanol: Verification of Homogeneity and Stability. (Abst.) *J. Appl. Toxicol.* Jan-Feb;15(1):1-4.

ATSDR. 2003. Health Consultation, Great Northern Bark Company, Columbia Falls, Flathead County, Montana, Exposure Investigation and Consultation Branch, Division of Health Assessment and Consultation, Agency for Toxic Substances and Disease Registry, Centers for Disease Control and Prevention, Atlanta, Georgia.

Astill, B. C., Deckardt, K., Gembardt, C., Gigell R., Guest, D., Hodgson, J. R., Mellert, W., Murphy, S. R., and T. R. Tyler. 1996. Prechronic Toxicity Studies on 2-Ethylhexanol in F344 Rats and B6C3F1 Mice. *Fund. Appl. Toxicol.* 29:31-39.

Astill, B. D., Gingell, R., Guest, D., Hellwig, J., Hodgson, J. R., Kuettler, K., Mellert, W., Murphy, S. R., Sielken, R. L., and T. R. Tyler. 1996. Oncogenicity Testing of 2-Ethylhexanol in Fischer 344 Rats and B6C3F1 Mice. *Fund. Appl. Toxicol.* 31:2941.

Baker, J. T. 2004. 2-Ethyl-1-Hexanol. Material Safety Data Sheet. Mallinckrodt Baker, Phillipsburg, NJ.

BASF. 1992. Report on the Study of the Oncogenic Potential of 2-Ethylhexanol in Rats After Administration by Gavage (Aqueous Emulsion) for 24 Months. Department of Toxicology of BASF for 2-Ethylhexanol Program Panel, Chemical Manufacturers Association, Washington, DC.

BASF. 1991. Report on the Study of the Oncogenic Potential of 2-Ethylhexanol in Mice after Administration by Gavage (Aqueous Emulsion) for 18 months. Department of Toxicology of BASF for 2-Ethylhexanol Program Panel, Chemical Manufacturers Association, Washington, DC.

Chemopetrol. 2004. 2-Ethylhexanol Material Safety Data Sheet. Chemopetrol, Litvinov - Zaluzil, Czech Republic.

Furton (2003). [SPIE](#) Proceedings. Volume 5071 Sensors, and Command, Control, Communications, and Intelligence (C3I) Technologies for Homeland Defense and Law Enforcement II, Edward M. Carapezza, Editor, September 2003, pp. 183-192

Hardin, B. D., Schuler, R. L., Burg, J. R., Booth, G. M., Hazelden, K. P., MacKenzie, K. M., Piccirillo, V. J., and K. N. Smith. 1987. Evaluation of 60 Chemicals in a Preliminary Developmental Toxicity Test. *Teratog. Carcinog. Mutagen.*, 7: 29-48 as cited by Semino, G., 1998. Saturated Aliphatic Acyclic Branched-Chain Primary Alcohols, Aldehydes, and Acids. WHO Food Additives Series 40, Safety Evaluation of Certain Food Additives and Contaminants, International Programme on Chemical Safety, World Health Organization.

Leffingwell & Associates. 2001. Compilation of Odor and Flavor Thresholds in Water. (<http://www4.wittenberg.edu/academics/chem/LabSafety/odor-in-water.htm> [12-6-05]).

Moody, D. E. and J. K. Reddy. 1978. Hepatic Peroxisome (Microbody) Proliferation in Rats Fed Plasticizers and Related Compounds. *Toxicol. Appl. Pharmacol.* 45:497-504.

N.J.A.C., 1993. Ground Water Quality Standards. New Jersey Administrative Code 7:9-6, January 8, 1993.

NTP. 1991. Developmental Toxicity of 2 Ethylhexanol (CAS NO. 104-76-7) in CD-1 Swiss Mice. National Toxicology Program, NTP Study: TER 90029, NTIS:PB91- 185900. National Toxicology Program Internet Site.

NTP. 2005. 2-Ethylhexanol, Selected Information from the National Library of Medicine Databases:ChemIDPlus and HSDB. National Toxicology Program Internet Site.

NTP. 2006. 2-Ethylhexanol, National Toxicology Program, National Library of Medicine's Hazardous Substance Database.

NUVO. 1999. 2-Ethylhexanol Material Safety Data Sheet. NUVO Australia Pty Ltd. 15 November 1999.

PAN Pesticides Database. 2006. <http://www.pesticideinfo.org/>.

Ritter, R. J., Scott, Jr., W. J., Randall, J. L. and J. M. Ritter. 1987. Teratogenicity of Di (2-Ethylhexyl) Phthalate, 2-Ethylhexanol, 2-Ethylhexanoic Acid, and Valproic Acid, and Potentiation by Caffeine. *Teratology* 35:41-46.

Rhodes, C., Soames, T., Stonard, M.D., Simpson, M.G., Vernal!, A.J. and C. R. Elcombe. 1984. The Absence of Testicular Arophy and In Vivo and In Vitro Effects on Hepatocyte Morphology and Peroxisomal Enzyme Activities in Male Rats Following Administration of Several Alkano. *Toxicol. Lett.* 21, 103-109.

Sanders, P. 2006. Derivation of Henry's Law Constant. NJDEP, DSRT.

Tyl, R. W., Jones-Price, C., Marr, M.C. and C. A. Kimmel. 1988. Developmental Toxicity Evaluation of Dietary Di(2-Ethylhexyl)Phthalate in Fischer 344 Rats and CD-1 Mice. *Fundam. Appl. Toxicol.* 10:395-412.

USEPA. 2002. A Review of the Reference Dose and Reference Concentration Processes, Risk Assessment Forum, Final Report EPA/630/P-02-002F, U. S. Environmental Protection Agency, Washington, D.C.

USEPA. 2005. United States Environmental Protection Agency. Guidelines for Carcinogen Risk Assessment. Risk Assessment Forum, USEPA, Washington, DC. EPA/630.P-03/001F, March 2005.

USEPA. 2006. High Production Volume Information System (HPVIS). Environmental Protection Agency, Washington, D.C. <http://www.epa.gov/hpvis/>.

WHO. 1993. World Health Organization. Ethyl-I-hexanol, 2-, Document Number 786, World Health Organization, Geneva, Food Additives Series 32, prepared by the forty-first meeting of the Joint FAO/WHO Expert Committee on Food Additives (JECFA) based on a draft document prepared by Dr. K. Ekelman, Additive Evaluation Branch, Division of Health Effects Evaluation, Center for Food Safety and Applied Nutrition, Food and Drug Administration, Washington, DC, USA. (<http://www.inchem.org/documents/jecfa/jecmono/v32je04.htm> [November, 2005]) This report contains information from many proprietary unpublished reports voluntarily submitted to the WHO. Some ambiguities were noted in the report. To reconcile these copies of the primary BASF studies were obtained through the U.S. Environmental Protection Agency Docket Center.



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