2-Ethoxy-4-methylphenol

CAS N°: 2563-07-7 **Empirical formula:** $C_9H_{12}O_2$

Synonyms: 2-Ethoxy-p-cresol
2-Ethoxy-4-methylphenol
4-Methyl-2-ethoxyphenol
Phenol, 2-ethoxy-4-methyl-
Supravanil, Ultravanil

History:

Initial reviews:	New Standard	
Current revision date:	2008	
Implementation date:	For new submissions*:	August 16, 2008
	For existing fragrance compounds*:	August 16, 2010
Next review date:	2013	

* This date applies to the supply of fragrance compounds (formulas) only, not to the finished products in the marketplace.

Standard: RESTRICTED

RESTRICTIONS:

Limits in the finished product:
For a description of the categories, refer to the QRA Informational Booklet.

Category 1: See Note box(1)	0.01 %	Category 7:	0.02 %
Category 2:	0.01 %	Category 8:	0.2 %
Category 3:	0.03 %	Category 9:	1.2 %
Category 4:	0.1 %	Category 10:	1.9 %
Category 5:	0.1 %	Category 11: Not Restricted (2)	
Category 6: See Note box(1)	0.2 %		

Note box:
(1) IFRA would recommend that any material used to impart perfume or flavour in products intended for human ingestion should consist of ingredients that are in compliance with appropriate regulations for foods and food flavourings in the countries of planned distribution and, where these are lacking, with the recommendations laid down in the Code of Practice of IOFI (International Organisation of the Flavor Industry) (http://www.iofiorg.org/).

(2) Category 11 includes all non-skin contact or incidental skin contact products. Due to the negligible skin contact from these types of products there is no justification for a restriction of the concentration of this fragrance ingredient in the finished product.

Fragrance material specification: N/A

Contribution from other sources: None to consider (see also the note on contributions from other sources in the **Introduction to the IFRA Standards).**

Critical effect: Sensitization

2-Ethoxy-4-methylphenol

RIFM Summaries:

LLNA weighted mean **EC3** values ($\mu g/cm^2$) [No. Studies]	**Potency Classification** Based on Animal Data[1]	**Human Data**			**WoE NESIL[3]** ($\mu g/cm^2$)
		NOEL – HRIPT (induction) ($\mu g/cm^2$)	**NOEL – HMT** (induction) ($\mu g/cm^2$)	**LOEL[2]** (induction) ($\mu g/cm^2$)	
	Moderate	236[4]	2362[4]	NA	230

NOEL = No observed effect level; HRIPT = Human Repeat Insult Patch Test; HMT = Human Maximization Test;

LOEL = lowest observed effect level; NA = Not Available

[1] Based on animal data using classification defined in ECETOC, Technical Report No. 87, 2003

[2] Data derived from HRIPT or HMT

[3] WoE NESIL limited to three significant figures

[4] *MT-NOEL = Maximum Tested No Effect Level. No sensitization was observed in human predictive studies. Doses reported reflect the highest concentration tested, not necessarily the highest achievable NOEL*

REXPAN Rationale / Conclusion:

The RIFM Expert Panel reviewed the critical effect data for 2-Ethoxy-4-methylphenol and, based on the weight of evidence, established the No Expected Sensitization Induction Level (NESIL) as 230 µg/cm². They recommend the limits for the 11 different product categories, which are the acceptable use levels of 2-Ethoxy-4-methylphenol in the various product categories. These were derived from the application of the exposure-based quantitative risk assessment approach for fragrance ingredients, which is detailed in the QRA Expert Group Technical Dossier of June 22, 2006.

References:

RIFM (Research Institute for Fragrance Materials, Inc.), 2003a. Human Repeated Insult Patch Test. RIFM report number 44237, December 10 (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2003b. Human Repeated Insult Patch Test. RIFM report number 44238, December 10 (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2004. Human Repeated Insult Patch Test. RIFM report number 45135, March 30 (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2007. Human Repeated Insult Patch Test. RIFM report number 52894, May 15 (RIFM, Woodcliff Lake, NJ, USA).