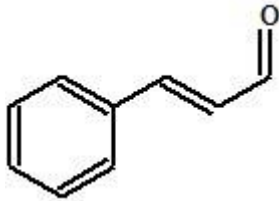


Cinnamic aldehyde

CAS N°:	104-55-2	Empirical formula:	C ₉ H ₈ O
Structure:			
Synonyms:	Cinnamal Cinnamaldehyde Phenylacrolein 3-Phenyl-2-propenal 3-Phenyl-2-propen-1-al 'Cassia aldehyde'		

History:	Initial reviews:	March 1978, April 2004, May 2006, May 2007, June 2008		
	Current revision date:	June 2013		
	Implementation date:	For new submissions*:	August 10, 2013	
		For existing fragrance compounds*:	August 10, 2014	
	Next review date	2018		

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

RECOMMENDATION:

RESTRICTED

RESTRICTIONS:

Limits in the finished product:			
Category 1 See Note box (1)	0.02 %	Category 7	0.04 %
Category 2	0.02 %	Category 8	0.05 %
Category 3	0.05 %	Category 9	0.05 %
Category 4	0.05 %	Category 10	0.05 %
Category 5	0.05 %	Category 11	Not Restricted (2)
Category 6	0.4 %		
Note box:			
<p>For this material, for pragmatic reasons, restrictive levels allowed by the QRA for certain categories but actually being higher than those already in place before applying the QRA, will temporarily not be implemented until the end of a 5 year monitoring phase. At the end of the 5 years the position will be reevaluated again.</p> <p>(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 - www.iofi.org)</p> <p>(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.</p> <p>The Standard on cinnamic aldehyde covers and replaces the former existing Standards on cassia oil, cinnamon bark oil as well as cinnamic aldehyde – methyl anthranilate schiff base. The existing Standards for cinnamic aldehyde, cassia oil, cinnamic aldehyde – methyl anthranilate schiff base and cinnamon bark oil are no longer valid.</p>			

Cinnamic aldehyde

Fragrance material specifications: N/A

CONTRIBUTION FROM OTHER SOURCES:

See **Annex I** and **Annex II**

CRITICAL EFFECT: **SENSITIZATION**

RIFM SUMMARIES:

LLNA weighted mean EC3 values ($\mu\text{g}/\text{cm}^2$) [no. studies]	Potency Classification Based on Animal Data ²	Human Data			WoE NESIL ³ ($\mu\text{g}/\text{cm}^2$)
		NOEL – HRIPT (induction) ($\mu\text{g}/\text{cm}^2$)	NOEL – HMT (induction) ($\mu\text{g}/\text{cm}^2$)	LOEL ¹ (induction) ($\mu\text{g}/\text{cm}^2$)	
262 [23]	Moderate	591 ⁴	NA	775	590

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 cinnamic aldehyde and, based on the weight of evidence, established the No Expected Sensitization Induction Level (NESIL) as 590 $\mu\text{g}/\text{cm}^2$. They recommend the limits for the 11 different product categories, which are the acceptable use levels of cinnamic aldehyde 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:

Basketter, D.A., Wright, Z.M., Warbrick, E.V. Dearman, R.J., Kimber, I., Ryan, C.A., Gerberick, G.F., White, I.R., 2001. Human potency predictions for aldehydes using the local lymph node assay. *Contact Dermatitis*, 45(2), 89-94.

Basketter, D.A., Wright, Z., Gilmour, N.J., Ryan, C.A., Gerberick, G.F., Robinson, M.K., Dearman, R.J., Kimber, I., 2002. Prediction of human sensitization potency using local lymph node assay EC3 values. *The Toxicologist*, 66(1-S), 240.

Elahi, E.N., Wright, Z., Hotchkiss, S.A.M., Basketter, D.A., Smith Pease, C.K., 2002. Protein binding and metabolic inhibition reveals clues on the mechanisms surrounding relative potency of sensitizing cinnamic compounds. *Toxicology*, 178, 52.

Elahi, E.N., Wright, Z., Hinselwood, D., Hotchkiss, S.A.M., Basketter D.A., Smith Pease, C.K., 2004. Protein binding and metabolism influence the relative skin sensitization potential of cinnamic compounds. *Chemical Research in Toxicology*, 17(3), 301-310.

RIFM (Research Institute for Fragrance Materials, Inc.), 1973. Repeated Insult Patch Test on Cinnamaldehyde. RIFM report number 12509, January 23. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2002a. Repeated Insult Patch Test on Cinnamaldehyde. RIFM report number 41692, August 27. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2002b. Repeated Insult Patch Test on Cinnamaldehyde. RIFM report number 41693, August 27. (RIFM, Woodcliff Lake, NJ, USA).

Cinnamic aldehyde

RIFM (Research Institute for Fragrance Materials, Inc.), 2003a. Local Lymph Node Assay on Cinnamaldehyde. RIFM report number 42032, February 18. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2003b. Local Lymph Node Assay on Cinnamaldehyde. RIFM report number 42033, February 18. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2003c. Local Lymph Node Assay on Cinnamaldehyde. RIFM report number 42040, February 18. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2003d. Local Lymph Node Assay on Cinnamaldehyde. RIFM report number 42034, February 18. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2003e. Local Lymph Node Assay on Cinnamaldehyde. RIFM report number 42036, February 18. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2003f. Local Lymph Node Assay on Cinnamaldehyde. RIFM report number 42035, February 18. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2003g. Local Lymph Node Assay on Cinnamaldehyde. RIFM report number 42037, February 18. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2003h. Local Lymph Node Assay on Cinnamaldehyde. RIFM report number 42038, February 18. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2003i. Local Lymph Node Assay on Cinnamaldehyde. RIFM report number 42039, February 18. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2003j. Local Lymph Node Assay on Cinnamaldehyde. RIFM report number 42041, February 18. (RIFM, Woodcliff Lake, NJ, USA).

RIFM (Research Institute for Fragrance Materials, Inc.), 2004. Repeated Insult Patch Test on Cinnamaldehyde. RIFM report number 47158, April 22a. (RIFM, Woodcliff Lake, NJ, USA).

Smith, C.K., Hotchkiss, S.A.M., 2001. Allergic Contact Dermatitis. Chemical and Metabolic Mechanisms. *Taylor and Francis*, London.

Wright, Z.M., Basketter, D.A., Blaikie, L., Cooper, K.J., Warbrick, E.V., Dearman, R.J., Kimber, I., 2001. Vehicle effects on skin sensitization potency of four chemicals assessment using the local lymph node assay. *Journal International Journal of Cosmetic Science*, 23(2), 75-83.