



Justification Document for the Selection of a CoRAP Substance

Substance Name (public name):	4-methylanisole
EC Number:	203-253-7
CAS Number:	104-93-8
Authority:	Health & Safety Authority, Ireland
Date:	17/03/2021

Cover Note

This document has been prepared by the evaluating Member State given in the CoRAP update

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1 IDENTITY OF THE SUBSTANCE

1.1 Other identifiers of the substance

Table: Other Substance identifiers

EC name (public):	4-methylanisole
IUPAC name (public):	1-methoxy-4-methylbenzene
Index number in Annex VI of the CLP Regulation:	Not applicable
Molecular formula:	C ₈ H ₁₀ O
Molecular weight or molecular weight range:	122.1644
Synonyms:	Anisole, p-methyl- (8CI) Benzene, 1-methoxy-4-methyl- (9CI) Methyl p-methylphenyl ether Methyl p-tolyl ether p-Cresol methyl ether p-Methoxytoluene p-Methylanisol p-Methylanisole p-Tolyl methyl ether

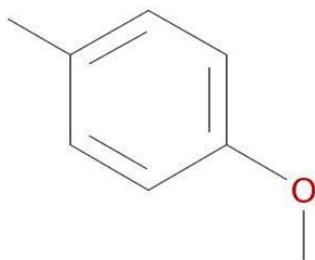
Type of substance

Mono-constituent

Multi-constituent

UVCB

Structural formula:



1.2 Similar substances/grouping possibilities

Not Applicable.

2 OVERVIEW OF OTHER PROCESSES / EU LEGISLATION

Table: Completed or ongoing processes

RMOA	<input type="checkbox"/> Risk Management Option Analysis (RMOA)	
REACH Processes	Evaluation	<input type="checkbox"/> Compliance check
		<input type="checkbox"/> Testing proposal
		<input checked="" type="checkbox"/> CoRAP and Substance Evaluation
	Authorisation	<input type="checkbox"/> Candidate List
		<input type="checkbox"/> Annex XIV
Restriction	<input type="checkbox"/> Annex XVII ¹	
CLH	<input type="checkbox"/> Annex VI (CLP) (see section 3.1)	
Processes under other EU legislation	<input type="checkbox"/> Plant Protection Products Regulation Regulation (EC) No 1107/2009	
	<input type="checkbox"/> Biocidal Product Regulation Regulation (EU) 528/2012 and amendments	
Previous legislation	<input type="checkbox"/> Dangerous substances Directive 67/548/EEC (NONS)	
	<input type="checkbox"/> Existing Substances Regulation 793/93/EEC (RAR/RRS)	
(UNEP) Stockholm convention (POPs Protocol)	<input type="checkbox"/> Assessment	
	<input type="checkbox"/> In relevant Annex	
Other processes/ EU legislation	<input type="checkbox"/> Other (provide further details below)	
Further details	4-Methylanisole was included on the CoRAP for evaluation in 2012, with initial concerns related to developmental toxicity (reproductive toxicity) and exposure. The Competent Authority of Ireland was appointed to carry out the evaluation. Based on the evaluation of the available data, the evaluating MSCA concluded there was a need to request further information to clarify the concerns relating to developmental toxicity and worker long-term exposure	

¹ Please specify the relevant entry.

	<p>(systemic, inhalation) and therefore prepared a draft decision to request further information.</p> <p>During the registrants commenting period on the draft decision, the only registrant of the substance informed the evaluating MSCA of their intention to cease manufacture of the substance in accordance with Article 50(3) of REACH. The registration was subsequently revoked and as there were no other registrants of the substance at that time, the substance evaluation decision making process related to the draft decision was terminated and no further information was requested. Therefore, the concern relating to developmental toxicity remained unverified.</p> <p>As of July 2020, there are 7 active full registration dossiers and 1 individual intermediate registration dossier for this substance, which were submitted since the termination of the original substance evaluation. These registrations include industrial, professional and consumer uses. No additional toxicological data is reported in these registrations and therefore the concern from the original substance evaluation in 2012 relating to developmental toxicity remains unverified.</p>
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3 HAZARD INFORMATION (INCLUDING CLASSIFICATION)

3.1 Classification

3.1.1 Harmonised Classification in Annex VI of the CLP

A harmonised classification is not available for this substance.

3.1.2 Self classification

- In the registrations:
Flam. Liquid 3 (H226), Acute Tox 4 (H302), Skin Irrit. 2 (H315), Repr. 2 (H361) and Aquatic Chronic 3 (H412).
- The following hazard classes are notified among aggregated self classifications in the C&L Inventory:

Table: Self classification

Index No	International Chemical Identification	EC No	CAS No	Classification		Spec. Conc. Limits, M-factors	Notes
				Hazard Class and Category Code(s)	Hazard statement code(s)		
-	4-Methylanisole	203-253-7	104-93-8	Flam. Liquid 3 Acute Tox. 4 Skin Irrit. 2 Eye Irrit. 2 Acute Tox. 3 Repr. 2 Aquatic Chronic 3	H226 H302 H315 H319 H331 H361 H412	N/A	

3.1.3 Proposal for Harmonised Classification in Annex VI of the CLP

Not relevant.

4 INFORMATION ON (AGGREGATED) TONNAGE AND USES²**4.1 Tonnage and registration status****Table: Tonnage and registration status**

From ECHA dissemination site *		
<input checked="" type="checkbox"/> Full registration(s) (Art. 10)	<input checked="" type="checkbox"/> Intermediate registration(s) (Art. 17 and/or 18)	
Tonnage band (as per dissemination site)		
<input type="checkbox"/> 1 – 10 tpa	<input checked="" type="checkbox"/> 10 – 100 tpa	<input type="checkbox"/> 100 – 1000 tpa
<input type="checkbox"/> 1000 – 10,000 tpa	<input type="checkbox"/> 10,000 – 100,000 tpa	<input type="checkbox"/> 100,000 – 1,000,000 tpa
<input type="checkbox"/> 1,000,000 – 10,000,000 tpa	<input type="checkbox"/> 10,000,000 – 100,000,000 tpa	<input type="checkbox"/> > 100,000,000 tpa
<input type="checkbox"/> <1 >+ tpa (e.g. 10+ ; 100+ ; 10,000+ tpa)		<input type="checkbox"/> Confidential
There are 7 active full registrations, which are part of a joint submission and 1 individual intermediate registration.		

*the total tonnage band has been calculated by excluding the intermediate uses, for details see the Manual for Dissemination and Confidentiality under REACH Regulation (section 2.6.11): https://echa.europa.eu/documents/10162/22308542/manual_dissemination_en.pdf/7e0b87c2-2681-4380-8389-cd655569d9f0

² Date when the dissemination site was accessed – 08/07/2020.

4.2 Overview of uses

The substance is used by consumers, by professional workers (widespread uses), in formulation or re-packing, at industrial sites and in manufacturing.

Table: Uses

Part 1:

<input checked="" type="checkbox"/> Manufacture	<input checked="" type="checkbox"/> Formulation	<input checked="" type="checkbox"/> Industrial use	<input checked="" type="checkbox"/> Professional use	<input checked="" type="checkbox"/> Consumer use	<input type="checkbox"/> Article service life	<input type="checkbox"/> Closed system
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Part 2:

	Use(s)
Uses as intermediate	Intermediate – substance manufactured and used under controlled conditions.
Formulation	Formulation of fragrances and fragrance end products, flavours for tobacco products, polishes and waxes, and washing and cleaning products.
Uses at industrial sites	Industrial use of washing, cleaning and disinfectant products, and metal surface treatment products.
Uses by professional workers	Professional use of polishes and waxes, washing, cleaning and disinfectant products, cosmetics and personal care products.
Consumer Uses	Consumer use of biocides (e.g. disinfectants, pest control products), washing and cleaning products, air care products, polishes and waxes, cosmetics and personal care products, perfumes and fragrances and tobacco products.
Article service life	-

5. JUSTIFICATION FOR THE SELECTION OF THE CANDIDATE CoRAP SUBSTANCE

5.1. Legal basis for the proposal

- Article 44(2)
 Article 45(5)

5.2. Selection criteria met (why the substance qualifies for being in CoRAP)

- Fulfils criteria as CMR/ Suspected CMR
 Fulfils criteria as Sensitiser/ Suspected sensitiser
 Fulfils criteria as potential endocrine disrupter
 Fulfils criteria as PBT/vPvB / Suspected PBT/vPvB
 Fulfils criteria high (aggregated) tonnage (*tpa* > 1000)
 Fulfils exposure criteria
 Fulfils MS's (national) priorities

5.3. Initial grounds for concern to be clarified under Substance Evaluation

Hazard based concerns		
CMR <input type="checkbox"/> C <input type="checkbox"/> M <input checked="" type="checkbox"/> R	Suspected CMR ¹ <input type="checkbox"/> C <input type="checkbox"/> M <input type="checkbox"/> R	<input type="checkbox"/> Potential endocrine disruptor
<input type="checkbox"/> Sensitiser	<input type="checkbox"/> Suspected Sensitiser ³	
<input type="checkbox"/> PBT/vPvB	<input type="checkbox"/> Suspected PBT/vPvB ¹	<input type="checkbox"/> Other (please specify below)
Exposure/risk based concerns		
<input type="checkbox"/> Wide dispersive use	<input type="checkbox"/> Consumer use	<input type="checkbox"/> Exposure of sensitive populations
<input type="checkbox"/> Exposure of environment	<input type="checkbox"/> Exposure of workers	<input type="checkbox"/> Cumulative exposure
<input type="checkbox"/> High RCR	<input type="checkbox"/> High (aggregated) tonnage	<input type="checkbox"/> Other (please specify below)

³ CMR/Sensitiser: known carcinogenic and/or mutagenic and/or reprotoxic properties/known sensitising properties (according to CLP harmonized or registrant self-classification or CLP Inventory)

Suspected CMR/Suspected sensitiser: suspected carcinogenic and/or mutagenic and/or reprotoxic properties/suspected sensitising properties (not classified according to CLP harmonized or registrant self-classification)

Suspected PBT: Potentially Persistent, Bioaccumulative and Toxic

There are two reproductive/developmental toxicity screening studies with 4-methylanisole reported in the registration data. In a dermal study in rats, no adverse effects were observed in parental animals or offspring up to the highest dose tested, although there were some uncertainties regarding the dosing regimen and the proportion of the applied dose that was systemically available. In an oral study in rats, adverse effects on pre- and post-natal development in the offspring, including an increase in post-implantation loss, a decrease in live birth index and an increase in the number of stillborn pups, were observed. Based on the results of this study, the registrants self classified 4-methylanisole as Repr. 2 H361.

The registration data reports industrial, professional and consumer uses of 4-methylanisole indicating the potential for exposure to humans. Further assessment of the available reproductive toxicity data is required to determine whether additional data to address the reproductive toxicity endpoint is needed and a more stringent hazard classification is warranted. At the registered tonnage band of 4-methylanisole (10-100 tpa), the data requirements relating to reproductive toxicity are limited to a screening study for reproductive/developmental toxicity (OECD 421/422) (Annex VIII, 8.7.1). Following an assessment of all available data under substance evaluation, further information may be requested to investigate the concern for reproductive toxicity (developmental toxicity), which is not possible to request under dossier evaluation at this tonnage band.

5.4. Preliminary indication of information that may need to be requested to clarify the concern

<input checked="" type="checkbox"/> Information on toxicological properties	<input type="checkbox"/> Information on physico-chemical properties
<input type="checkbox"/> Information on fate and behaviour	<input type="checkbox"/> Information on exposure
<input type="checkbox"/> Information on ecotoxicological properties	<input type="checkbox"/> Information on uses
<input type="checkbox"/> Information ED potential	<input type="checkbox"/> Other (provide further details below)
Following an assessment of all available data, further information to investigate the concern for reproductive toxicity (developmental toxicity) may be requested.	

5.5. Potential follow-up and link to risk management

<input checked="" type="checkbox"/> Harmonised C&L	<input type="checkbox"/> Restriction	<input type="checkbox"/> Authorisation	<input type="checkbox"/> Other (provide further details)
The registrants have self-classified the substance as Repr. 2 H361 based on the available data. Industrial, professional and consumer uses of 4-methylanisole are reported in the registration data. Clarification of the concern for reproductive toxicity under substance evaluation could lead to the conclusion that a more stringent classification for reproductive toxicity may be warranted. Therefore, depending on the outcome of the evaluation, there may be a need to prepare a proposal for harmonised classification and labelling for reproductive toxicity.			