



Committee for Risk Assessment

RAC

Annex 1

Background document

to the Opinion proposing harmonised classification
and labelling at EU level of

1,1',1''-nitrilotripropan-2-ol (TIPA)

ECHA/RAC/CLH-O-0000002510-87-01/A1

EC Number: 204-528-4

CAS Number: 122-20-3

The background document is a compilation of information considered relevant by the dossier submitter or by RAC for the proposed classification. It includes the proposal of the dossier submitter and the conclusion of RAC. It is based on the official CLH report submitted to public consultation. RAC has not changed the text of this CLH report but inserted text which is specifically marked as 'RAC evaluation'. Only the RAC text reflects the view of RAC.

Adopted

8 March 2012

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Part A.

1 PROPOSAL FOR HARMONISED CLASSIFICATION AND LABELLING

1.1 Substance

Table 1: Substance identity

Substance name:	1,1',1''-nitriлотрипропан-2-ол
EC number:	204-528-4
CAS number:	122-20-3
Annex VI Index number:	603-097-00-3
Degree of purity:	See IUCLID file and confidential Annex
Impurities:	See IUCLID file and confidential Annex

1.2 Harmonised classification and labelling proposal

Table 2: The current Annex VI entry and the proposed harmonised classification

	CLP Regulation	Directive 67/548/EEC (Dangerous Substances Directive; DSD)
Current entry in Annex VI, CLP Regulation	Eye Irrit. 2 (H 319) Aquatic Chronic 3 (H 412)	Xi; R36, R52/53
Current proposal for consideration by RAC	Eye Irrit. 2 (H 319) Not classified for Aquatic Chronic 3	Xi; R36 Not classified for R52/53
Resulting harmonised classification (future entry in Annex VI, CLP Regulation)	Eye Irrit. 2 (H 319)	Xi; R36

1.3 Proposed harmonised classification and labelling based on CLP Regulation and/or DSD criteria

Table 3: Proposed classification according to the CLP Regulation

CLP Annex I ref	Hazard class	Proposed classification	Proposed SCLs and/or M-factors	Current classification ¹⁾	Reason for no classification ²⁾
2.1.	Explosives				Conclusive but not sufficient for classification
2.2.	Flammable gases				Conclusive but not sufficient for classification
2.3.	Flammable aerosols				Conclusive but not sufficient for classification
2.4.	Oxidising gases				Conclusive but not sufficient for classification
2.5.	Gases under pressure				Conclusive but not sufficient for classification
2.6.	Flammable liquids				Conclusive but not sufficient for classification
2.7.	Flammable solids				Conclusive but not sufficient for classification
2.8.	Self-reactive substances and mixtures				Conclusive but not sufficient for classification
2.9.	Pyrophoric liquids				Conclusive but not sufficient for classification
2.10.	Pyrophoric solids				Conclusive but not sufficient for classification
2.11.	Self-heating substances and mixtures				Conclusive but not sufficient for classification
2.12.	Substances and mixtures which in contact with water emit flammable gases				Conclusive but not sufficient for classification
2.13.	Oxidising liquids				Conclusive but not sufficient for classification
2.14.	Oxidising solids				Conclusive but not sufficient for classification
2.15.	Organic peroxides				Conclusive but not sufficient for classification
2.16.	Substance and mixtures corrosive to metals				Conclusive but not sufficient for classification
3.1.	Acute toxicity - oral				Conclusive but not sufficient for classification
	Acute toxicity - dermal				Conclusive but not sufficient for classification
	Acute toxicity - inhalation				Conclusive but not sufficient for classification
3.2.	Skin corrosion / irritation				Conclusive but not sufficient for classification

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CLP Annex I ref	Hazard class	Proposed classification	Proposed SCLs and/or M-factors	Current classification ¹⁾	Reason for no classification ²⁾
3.3.	Serious eye damage / eye irritation	Eye irritant 2	none	Eye irritant 2	
3.4.	Respiratory sensitisation				Conclusive but not sufficient for classification
3.4.	Skin sensitisation				Conclusive but not sufficient for classification
3.5.	Germ cell mutagenicity				Conclusive but not sufficient for classification
3.6.	Carcinogenicity				Conclusive but not sufficient for classification
3.7.	- Reproductive toxicity - Reproductive toxicity – Effects on or via lactation				Conclusive but not sufficient for classification Conclusive but not sufficient for classification
3.8.	Specific target organ toxicity –single exposure				Conclusive but not sufficient for classification
3.9.	Specific target organ toxicity – repeated exposure				Conclusive but not sufficient for classification
3.10.	Aspiration hazard				Conclusive but not sufficient for classification
4.1.	Hazardous to the aquatic environment	Not classified	none	Aquatic Chronic 3	Conclusive but not sufficient for classification
5.1.	Hazardous to the ozone layer				Conclusive but not sufficient for classification

¹⁾ Including specific concentration limits (SCLs) and M-factors

²⁾ Data lacking, inconclusive, or conclusive but not sufficient for classification

Labelling:

Signal word:

Hazard statements:

Precautionary statements:

Warning

H319: causes serious eye irritation

2 BACKGROUND TO THE CLH PROPOSAL

2.1 History of the previous classification and labelling

The current environmental classification results from no ready biodegradability and E_bC_{50} (algae) < 100 mg/L. According to the CLP Regulation the classification shall be based on E_rC_{50} (algae). The E_rC_{50} (algae) of 1,1',1''-nitriлотрипропан-2-ол is greater than 100 mg/L, therefore the substance should not be classified for environment.

2.2 Short summary of the scientific justification for the CLH proposal

Data from registration dossiers were taken as a basis for this CLH proposal.

Based on the available/presented data the classification/labelling with R 52/53 (aquatic chronic 3) is deemed to be not justified. The endpoints derived from acute aquatic toxicity studies are > 100 mg/L at each trophic level. Hence, the chemical is considered to be acutely not harmful to aquatic organisms including fish, aquatic invertebrates and algae. In addition, the water solubility of the substance is high (830 g/L). The experimentally determined BCF was < 1. Therefore the test substance does not fall under the criteria for Aquatic Chronic 3 (R52/53).

In 1998 the ready biodegradability of the test substance was assessed in an OECD 301 F study performed for DOW Elanco. Biodegradation was not observed during the test period. In a DOC Die Away-Test according to OECD 301 A (Hüls AG, 1997) a biodegradation degree of 18% was measured after 28 d of exposure indicating that the chemical is not readily biodegradable. In a MITI test (1992) resembling the test guideline OECD 302 C the absence of inherent biodegradation was demonstrated. Further, an OECD 302 B BASF-study from 1981 demonstrated a low potential for elimination from water. In terms of the CLP criteria the test substance has to be considered as not rapidly degradable.

According to CLP criteria the test substance is not harmful to fish as was demonstrated in a BASF AG study performed in 1987. The 96 -h LC_{50} value calculated as geometrical mean was 3158 mg/L (nominal test item concentrations). This result is supported by an acute toxicity test conducted according to EU Method C.1 (Acute Toxicity for Fish) performed by Hüls (1997). In this limit-test no mortality was observed at 1000 mg/L (LC_{50} (96 h) >1000 mg/L, nominal confirmed by concentration control analysis).

A BASF AG study conducted in 1987 according to the test method presented in directive 79/831/EEC, Annex V, part C indicated that the test substance is according to CLP criteria also most probably not acutely harmful to aquatic invertebrates. The EC_{50} based on mobility of *D. magna* was determined to be > 500 mg/L (based on nominal concentrations). These results are supported by an acute toxicity test according to EU Method C.2 (Acute Toxicity for *Daphnia*) performed by Hüls (1997). The EC_{50} (48 h) was 857 mg/L (nominal, confirmed by concentration control analytics).

Finally, according to CLP criteria the test substance is most probably not acutely harmful to algae as demonstrated in a study sponsored by Sasol Germany GmbH in 1997. In a test according to EU method C.3 an E_rC_{50} of 710 mg/L was determined. These results are supported by a BASF study

conducted in 1990. The E_rC_{50} , recalculated from fluorescence data, after 72 hours of exposure was determined to be > 100 mg/L.

The summarised results above combined with the high water solubility and the low bioconcentration factor demonstrate that the classification Aquatic Chronic 3 is not justified.

2.3 Current harmonised classification and labelling

2.3.1 Current classification and labelling in Annex VI, Table 3.1 in the CLP Regulation

Eye Irrit. 2
Aquatic Chronic 3

2.3.2 Current classification and labelling in Annex VI, Table 3.2 in the CLP Regulation

Xi; R36
R52/53

2.4 Current self-classification and labelling

2.4.1 Current self-classification and labelling based on the CLP Regulation criteria

Proposal: Eye Irrit. 2

2.4.2 Current self-classification and labelling based on DSD criteria

Proposal: Xi; R36

3 JUSTIFICATION THAT ACTION IS NEEDED AT COMMUNITY LEVEL

It is proposed that the substance is no more to be classified as aquatic chronic 3 (R52/53) based on the available test data presented in chapter 5. Harmonized classification and labelling for 1,1',1''-nitrilotripropan-2-ol is considered a Community-wide action under Article 42 and it is recommended that the classification proposal is considered for inclusion on Annex VI to Regulation (EC) No 1272/2008, table 3.1 and table 3.2.

In 1998 the ready biodegradability of the test substance was assessed in an OECD 301 F study performed for DOW Elanco. Biodegradation was not observed during the test period. In a DOC Die Away-Test according to OECD 301 A (Hüls AG, 1997) a biodegradation degree of 18% was measured after 28 d of exposure indicating that the chemical is not readily biodegradable. In a MITI test (1992) resembling the test guideline OECD 302 C the absence of inherent biodegradation was demonstrated. Further, an OECD 302 B BASF-study from 1981 demonstrated a low potential for elimination from water. In terms of the CLP criteria the test substance has to be considered as not rapidly degradable.

According to CLP criteria the test substance is not harmful to fish as was demonstrated in a BASF AG study performed in 1987. The 96 -h LC₅₀ value calculated as geometrical mean was 3158 mg/L (nominal test item concentrations). This result is supported by an acute toxicity test conducted according to EU Method C.1 (Acute Toxicity for Fish) performed by Hüls (1997). In this limit-test no mortality was observed at 1000 mg/L (LC₅₀ (96 h) >1000 mg/L, nominal confirmed by concentration control analysis).

A BASF AG study conducted in 1987 according to the test method presented in directive 79/831/EEC, Annex V, part C indicated that the test substance is according to CLP criteria also most probably not acutely harmful to aquatic invertebrates. The EC₅₀ based on mobility of *D. magna* was determined to be > 500 mg/L (based on nominal concentrations). These results are supported by an acute toxicity test according to EU Method C.2 (Acute Toxicity for *Daphnia*) performed by Hüls (1997). The EC₅₀ (48 h) was 857 mg/L (nominal, confirmed by concentration control analytics).

Finally, according to CLP criteria the test substance is most probably not acutely harmful to algae as demonstrated in a study sponsored by Sasol Germany GmbH in 1997. In a test according to EU method C.3 an E_rC₅₀ of 710 mg/L was determined. These results are supported by a BASF study conducted in 1990. The E_rC₅₀, recalculated from fluorescence data, after 72 hours of exposure was determined to be > 100 mg/L.

The summarised results above combined with the high water solubility and the low bioconcentration factor demonstrate that the classification Aquatic Chronic 3 is not justified.

Part B.

SCIENTIFIC EVALUATION OF THE DATA

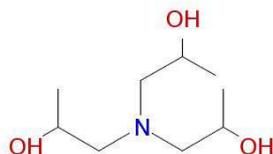
1 IDENTITY OF THE SUBSTANCE

1.1 Name and other identifiers of the substance

Table 5: Substance identity

EC number:	204-528-4
EC name:	1,1',1''-nitrilotripropan-2-ol; Triisopropanolamine
CAS number (EC inventory):	122-20-3
CAS number:	122-20-3
CAS name:	2-Propanol, 1,1',1''-nitrilotris-
IUPAC name:	1,1',1''-nitrilotripropan-2-ol
CLP Annex VI Index number:	603-097-00-3
Molecular formula:	C ₉ H ₂₁ NO ₃
Molecular weight range:	191.27

Structural formula:



1.2 Composition of the substance (as manufactured)

Table 6: Constituents (non-confidential information)

Constituent	Typical concentration	Concentration range	Remarks
1,1',1''-nitriлотрипропан-2-ол			

Current Annex VI entry: Eye Irrit. 2 (H 319), Aquatic Chronic 3 (H 412)

Table 7: Impurities (non-confidential information)

Impurity	Typical concentration	Concentration range	Remarks

Current Annex VI entry:

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Table 8: Additives (non-confidential information)

Additive	Function	Typical concentration	Concentration range	Remarks

Current Annex VI entry:

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1.2.1 Composition of test material

Physico-chemical and toxicological studies:

not relevant for this dossier. However, information on the test material used in the different studies is given in the IUCLID 5 dossier.

Eco-toxicological studies:

Information on the test material used is given in chapter 5 of this dossier and is reported in the IUCLID 5 dossier.

1.3 Physico-chemical properties

Table 9: Summary of physico - chemical properties

Property	Value	Reference	Comment (e.g. measured or estimated)
State of the substance at 20°C and 101,3 kPa	solid	Lewis (1997)	Visual inspection
Melting/freezing point	45°C	Lide (1998)	Measured
Boiling point	301°C (1013 hPa)	BASF (1972)	Measured
Density	1.0 g/cm ³ (20°C)	Lide (1998)	Measured
Vapour pressure	0.000000008 hPa (20°C)	BASF (1972)	Measured
Surface tension	Not surface active; based on chemical structure, no surface activity is to be expected	Expert judgment	Expert judgement
Water solubility	830 g/l (20°C)	Davis (1997), IPCS (2006)	Measured
Partition coefficient n-octanol/water	-0.015 (23°C)	BASF (1987a)	Measured
Flash point	174°C (1013 hPa) c.c.	BASF (1978)	Measured, DIN 51758

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Property	Value	Reference	Comment (e.g. measured or estimated)
Flammability upon ignition (solids)	Combustible when exposed to heat or flame Because of the low melting point, the substance is used in a liquid form therefore the flammability is deduced from flash point and boiling point.	Lewis (2004), Sax`s, 11th ed.	Measured
Flammability in contact with water	Not conducted (Testing can be waived) ¹⁾	BAM-II.2 (2010)	Expert judgement
Pyrophoric properties	Not conducted (Testing can be waived) ²⁾	BAM-II.2 (2010)	Expert judgement
Explosion limits in air	(LEL/LFL) = 0.8 vol% (UEL/UFL) = 5.8 vol%	IPCS (2006)	Measured
Dust explosion hazard	Dust explosion possible if in powder or granular form, mixed with air. Combustion and explosion characteristic of dust are not available.	IPCS (2006)	Literature value
Explosive properties	Not conducted (Testing can be waived) ³⁾	BAM-II.2 (2010)	Expert judgement
Self-ignition temperature	285°C	BASF (1978)	Measured, DIN 51794
Oxidising properties	Not conducted (Testing can be waived) ⁴⁾	BAM-II.2 (2010)	Expert judgement
Stability in organic solvents and identity of relevant degradation products	Not applicable; the stability of the substance is not considered as critical	Expert judgment	Expert judgement
Dissociation constant	7.86 (25°C)	Schwabe (1959)	Measured
Viscosity	100 mPa s (60°C)	Flick (1998)	Measured

¹⁾ Testing can be waived based on a consideration of the chemical structure in accordance with Annex I, section 2.12.4.1 of the CLP Regulation: The classification procedure needs not to be applied because the organic substance does not contain metals or metalloids.

²⁾ Testing can be waived in accordance with Annex I, section 2.10.4.1 of the CLP Regulation: The classification procedure needs not to be applied because the organic substance is known to be stable in contact with air at room temperature for prolonged periods of time (days).

³⁾ Testing can be waived based on a consideration of the chemical structure in accordance with REACH Column 2 of Annex VII, section 7.11: The classification procedure needs not to be applied because there are no chemical groups present in the molecule which are associated with explosive properties.

⁴⁾ Testing can be waived based on a consideration of the chemical structure in accordance with REACH Column 2 of Annex VII, section 7.13: The classification procedure needs not to be applied because the organic substance contains oxygen, which is chemically bonded only to carbon.

2 MANUFACTURE AND USES

2.1 Manufacture

Reaction of ammonia and propylenoxide at elevated temperature and pressure and further distillation.

2.2 Identified uses

In industrial settings 1,1',1''-nitrilotripropan-2-ol is used, besides manufacture and formulation, as an intermediate, as a processing aid for paper, textile and leather and as gas treatment. It is further used in metal working fluids and as an additive in fuel. Industrial uses are also the use in wood protection and as additive in plastic.

Professional uses include uses as additive in concrete and cement, as processing aid for paper, textile and leather as well as the use in metal working fluids. It is further used in coatings and adhesives, detergents and cleaners, as laboratory chemical and in fuels.

Consumers use the substance in concrete and cement, as well as in fuels. Further it is used in detergents and cleaners, wood protection formulations and in personal care products.

3 CLASSIFICATION FOR PHYSICO-CHEMICAL PROPERTIES

Chapter 3 is not relevant for this dossier.

4 HUMAN HEALTH HAZARD ASSESSMENT

Chapter 4 is not relevant for this dossier.

5 ENVIRONMENTAL HAZARD ASSESSMENT

5.1 Degradation

Table 21: Summary of relevant information on degradation

Method	Results	Remarks	Reference
Gas chromatography to study the stability of triisopropanolamine in an aqueous milieu Reliability 2: Peer reviewed data	Triisopropanolamine proved to be stable in water	Test item: Triisopropanolamine, purity: n.d.	Toropkov. 1980
OECD Guideline 301 F (Ready Biodegradability: Manometric Respirometry Test) Reliability 1: GLP guideline study	0% BOD/ThOD (28 d)	Test item: Triisopropanolamine, purity: 95%	Dow, 1998
OECD guideline 301 A (Ready Biodegradability: DOC Die Away Test) Reliability 2: Well documented study according to OECD guideline	15% DOC removal (28 d)	Test item: Triisopropanolamine	Huels AG, 1997
OECD Guideline 302 C (Inherent Biodegradability: Modified MITI Test (II)) Reliability 2: Guideline study with acceptable restrictions	3.4% BOD/ThOD (28 d)	Test item: tris(2-hydroxypropyl)amine [synonym: triisopropanolamine], purity: n.d.	MITI, 1992
According to OECD Guideline 302 B (Inherent biodegradability: Zahn-Wellens/EMPA Test) Reliability 2: Guideline study with acceptable restrictions	<10% DOC (28d)	Inoculum: activated sludge, industrial; Test item: Triisopropanolamine, purity: n.d.	BASF, 1981
EPA Subdivision N Pesticide Guideline 162-4 (Aerobic Aquatic Metabolism) Reliability 1: GLP guideline study	39% ¹⁴ CO ₂ , radiochem. meas. (30d); 64% radiochem. meas. (64d); half-life: 14.3 days	Test item: Triisopropanolamine, purity: 99.5%	Krieger, 1995
EPA Subdivision N Pesticide Guideline 162-3 (Anaerobic Aquatic Metabolism) Reliability 1: GLP guideline study	<1% ¹⁴ CO ₂ , radiochem. meas. (6m)	Test item: Triisopropanolamine, purity: 99.5%	Cleveland, 1995a
EPA 162-1: Aerobic soil metabolism study Reliability 1: GLP guideline study	66-72% ¹⁴ CO ₂ , radiochem. meas. (20d)	Test item: Triisopropanolamine, purity: n.a. purity (radiolab.): 95+x%	Cleveland, 1995b

5.1.1 Stability

Toropkov (1980) used gas chromatography to study the stability of triisopropanolamine in an aqueous milieu. No details of the tested concentration range, temperature range or pH range were provided. According to Toropkov, triisopropanolamine proved to be stable in water. At environmental pH conditions hydrolysis is not expected to be a relevant degradation process due to the absence of hydrolysable groups (Kollig et al. 1993, Boethling and Mackay 2000).

5.1.2 Biodegradation

5.1.2.1 Biodegradation estimation

No data.

5.1.2.2 Screening tests

The ready biodegradability of the test substance was assessed according to OECD 301 F. Biodegradation was not observed during the test period (Dow, 1998). In this test domestic non-adapted activated sludge was exposed to the test substance for 28 days. Additionally to the test item replicates, inhibition controls with benzoate and reference replicates were set up. After connection to the respirometer system, the reaction vessels were purged with ambient air, and the associated headspace volume of each individual reaction vessel was determined by the respirometer system. The reaction vessels were maintained in a dark room at a temperature of 22 ± 1 °C and continuously stirred over the 28-day period. Measurements of gas phase O₂ and CO₂ in the reaction vessels occurred on 4-hour sample intervals throughout the 28-day test period. The inhibition control demonstrated that the test substance was not inhibitory to the activated sludge.

In a DOC die away test performed by Sasol in 1997 the test substance was tested for ready biodegradation using domestic non-adapted activated sludge. At the end of the 28 day exposure period only 18% of the test substance were degraded.

In a MITI test (1992) resembling the test guideline OECD 302 C the absence of inherent biodegradation was demonstrated. The measured BOD after 4 weeks of exposure was 3.4%. 30 mg/L of the test substance were incubated with 100 mg/L MITI inoculum (mixture of sewage, soil and natural water collected from different places in Japan) as recommended by OECD Test guideline 302C (Modified MITI test). Further, an OECD 302 B BASF-study from 1981 demonstrated a low potential for elimination from water. In the test industrial activated sludge at a concentration of 1 g/L dry substance was exposed to 400 mg/L DOC of the test substance for 28 days in well aerated glass vessels. DOC removal at the end of the test was below 10%.

Taking into account all available data the test substance is considered to be not rapidly biodegradable in terms of the CLP criteria.

5.1.2.3 Simulation tests

Aerobic degradation in a water/sediment system and anaerobic degradation in a water/sediment system were conducted for Dow Elanco (Krieger, 1995). A half-life of 14.3 days was determined for the aerobic degradation of the test substance in the water/sediment system, indicating that 1,1',1''-nitriлотripropan-2-ol will not persist in aerobic aqueous compartments. The major identified metabolite was (2-oxopropyl)diisopropanolamine. For this metabolite no further information is available. After 30 days of exposure 39% of the applied radioactivity were found as ¹⁴CO₂ and after

60 days the amount of produced $^{14}\text{CO}_2$ increased to 64%. This demonstrates that TIPA is not rapidly biodegradable but is not persistent in the water compartment.

The anaerobic degradation in a water/sediment system demonstrated that the test substance was not degraded during an observation time of 6 months (Cleveland, 1995a).

In a study conducted for Dow Elanco (Cleveland, 1995b) the degradation of the test substance in two different soils was determined to be between 66 and 72% based on the evolution of $^{14}\text{CO}_2$, indicating that 1,1',1''-nitrilotripropan-2-ol will not persist in soil. The major identified metabolite was 1,1'-iminodiprop-2-ol (CAS 110-97-4), which is also an impurity of the test substance and is considered to be non-toxic to the aquatic environment and readily biodegradable (further information on the metabolite 1,1'-iminodiprop-2-ol may be obtained on: ECHA: Information on Registered Substances: <http://apps.echa.europa.eu/registered/registered-sub.aspx#search>).

5.1.3 Summary and discussion of degradation

Abiotic degradation due to hydrolysis is not expected as was demonstrated by Toropkov (1980). 1,1',1''-nitrilotripropan-2-ol proved to be stable in water. Further, at environmental pH conditions hydrolysis is not expected to be a relevant degradation process due to the absence of hydrolysable groups (Kollig et al. 1993, Boethling and Mackay 2000).

In screening tests 1,1',1''-nitrilotripropan-2-ol was found to be not biodegradable. However, in the water/sediment system the test substance has a half-life of 14.3 days under aerobic conditions. After 30 and 60 days of exposure 39% and 64% of the applied radioactivity were recovered as $^{14}\text{CO}_2$, respectively, indicating that 1,1',1''-nitrilotripropan-2-ol will not persist in aerobic aqueous compartments. In anaerobic media no biodegradation was observed after 6 month of exposure. In natural soil 1,1',1''-nitrilotripropan-2-ol is mineralised to an extent of 66 to 72%. Therefore, 1,1',1''-nitrilotripropan-2-ol is not rapidly or inherently biodegradable in regulatory terms but it does not persist in water/sediment systems due to degradation in surface water and in soil.

Based on the presented data the test substance is considered to be not rapidly biodegradable according to CLP criteria.

5.2 Environmental distribution

Table 22: Summary of relevant information on environmental distribution

Method	Results	Remarks	Reference
SRC PCKOC v2.0 calculation MCI based calculation Reliability 2: Scientifically acceptable method	Adsorption coefficient: log Koc: 1 (Koc estimate from MCI)	Test item: 1,1',1''-nitriлотripropan-2-ol	BASF AG, 2010
SRC PCKOC v2.0 calculation log Kow based calculation Reliability 2: Scientifically acceptable method	Adsorption coefficient: log Koc: 0.0258	Test item: 1,1',1''-nitriлотripropan-2-ol	BASF AG, 2010
Calculation of log Koc for ionized molecule Reliability 2: Scientifically acceptable method	Adsorption coefficient: log Koc: 1.92 (pH 5.0) log Koc: 1.87 (pH 7.0) log Koc: 1.34 (pH 9.0)	Test item: 1,1',1''-nitriлотripropan-2-ol	BASF SE, 2010 Franco A. & Trapp S., 2008
Calculation based on the correction factor recommende by ECHA guidance document R.7, appendix R7.1-2, page 190 to be used for ionisable substances Reliability 2: Scientifically acceptable method	Adsorption coefficient: log Koc: -1.86 (pH 5.0) log Koc: 0.08 (pH 7.0) log Koc: 0.97 (pH 9.0)	Test item: 1,1',1''-nitriлотripropan-2-ol	BASF SE, 2011
SRC HENRYWIN v3.10 calculation Reliability 2: Scientifically acceptable method	Henry's Law constant H: 0.000001 Pa m ³ /mol at 25 °C	Test item: 1,1',1''-nitriлотripropan-2-ol	BASF AG, 2007b
Mackay level I calculation Calculation programme: Level I Model, Version 3.00 Reliability 2: Scientifically acceptable method	Percent distribution in media: Water (%): 100 Soil (%): 0.01 Sediment (%): 0.01	Test item: 1,1',1''-nitriлотripropan-2-ol	BASF AG, 2007c

5.2.1 Adsorption/Desorption

Calculated logKoc-values of 1.0 and 0.0258 are available based on estimates from MCI and log Kow, respectively (BASF SE, KOCWIN v2.00, 2010). This value refers to the uncharged molecule (pKa value: 7.86). The pKa value indicates that the molecule will exist partly as a cation in the environment at neutral to acidic pH conditions. Cations generally adsorb stronger to soils containing organic carbon and clay than their neutral counterparts. Hence, the PCKOC-model may underestimate adsorption to organic carbon since it does not consider the ionic structure of the molecule. Under environmental conditions (pH from 5 to 9) the test substance is partly present in its charged form (as calculated by the formula % ionised = 100/(1+10(pKa - pH)): 7% at a pH of 9, 88% at pH 7, 100% at pH 5). In a calculation conducted according to a publication by Franco & Trapp, 2008 using the parameters pKa = 7.86 and log Pow = -1.22 for the uncharged molecule log Koc values of 1.92, 1.87 and 1.34 were determined for the pH values 5, 7 and 9, respectively.

The environmental pH value influences the sorption behaviour of ionisable substances. Based on the *Guidance on information requirements and chemical safety assessment Chapter R.7a: Endpoint*

specific guidance document, Appendix R.7.1-2 pH correction of partition coefficients for ionisable substances a correction factor to account for this influence may be applied to the values determined for the uncharged molecules. Using this correction factor on the calculated worst case Koc of 10 (MCI-method of PCKOC-model in Episuite), the resulting corrected log Koc was determined to be -1.86, 0.084 and 0.97 for the environmentally relevant pH values of 5, 7 and 9, respectively.

5.2.2 Volatilisation

A Henry law constant of 0.000001 Pa*m³/mol was calculated by SRC HENRYWIN v3.10 for the uncharged molecule (BASF SE, 2007b), indicating that the molecule will not evaporate into the atmosphere from the water surface.

5.2.3 Distribution modelling

Over time, the substance will preferentially distribute into the compartment water (100 %; Mackay Level I) (BASF SE, 2007c).

5.3 Aquatic Bioaccumulation

Table 23: Summary of relevant information on aquatic bioaccumulation

Method	Results	Remarks	Reference
OECD Guideline 305 C (Bioaccumulation: Test for the Degree of Bioconcentration in Fish) Species: <i>Cyprinus carpio</i> Reliability 2: Guideline study with acceptable restrictions	BCF <0.57 (0.25 mg/L); BCF <0.06 (2.5 mg/L)	Test item: tris(2-hydroxypropyl)amine [synonym: triisopropanolamine], purity: n.d.	MITI, 1992

5.3.1 Aquatic bioaccumulation

5.3.1.1 Bioaccumulation estimation

The bioaccumulation of the substance was not estimated, as measured bioaccumulation data from a MITI test according to OECD TG 305 C was available.

5.3.1.2 Measured bioaccumulation data

A MITI test (1992) according to guideline OECD 305 C resulted in bioconcentration factors of < 0.06 and < 0.57 at exposure concentrations of 2.5 mg/L and 0.25 mg/L, respectively. In the presented study carp were continuously exposed to the test chemical for 6 weeks in a flow-through system at a flow rate of 290 – 1150 L/d at 25 °C. The dissolved oxygen levels were kept at 6 – 8 mg/L. Fish were about 10 cm long and had an average body weight of 30 g, the lipid content was 2 - 6%. After termination of the exposure period the content of the test chemical in the whole fish was determined.

The study on the bioaccumulation in aquatic organisms (MITI, 1992) and the low measured log K_{OW} of -0.015 demonstrate that the test substance does not accumulate in aquatic organisms. According to CLP criteria the test substance is not bioaccumulative.

5.3.2 Summary and discussion of aquatic bioaccumulation

A study on the bioaccumulation in aquatic organisms (MITI, 1992) demonstrated that the test substance does not accumulate in aquatic organisms. According to CLP criteria a bioaccumulation factor of ≥ 500 and/or a partition coefficient octanol/water (log K_{ow}) of > 4 is indicative for the potential to bioconcentrate. Compared to the experimentally determined bioconcentration factor of < 1 and the measured log K_{ow} of -0.013 classification triggered by bioconcentration is not justified.

5.4 Aquatic toxicity

Table 24: Summary of relevant information on aquatic toxicity

Method	Results	Remarks	Reference
Fish			
<i>Leuciscus idus</i> - DIN 38412, Part 11 Reliability 2: Non-GLP study in accordance with german national industrial standard test guidelines. No analytical test item concentration verification.	LC50 (96 h): 3158.48 mg/L (geometric mean; nominal)	Test item: Triisopropanolamine, purity: >99% Due to the high water solubility, the test item was directly added to the test medium. The test was performed under static conditions. Test concentrations were 0, 1000, 2150, 4640 and 10000 mg/L. Additionally a neutralised sample of 10000 mg/L was also tested. Neutralisation did not alter the toxicity of the test substance. pH values ranged from 8.0 to 10.0 during the test. Dissolved oxygen concentrations ranged from 8.1 to 8.9 mg/L.	BASF AG, 1987b
<i>Cyprinus carpio</i> - EU Method C.1 (Acute Toxicity for Fish; limit test)	LC50 (96 h): > 1000 mg/L (nominal)	Test item: Triisopropanolamine, purity: >98.6%	Huels AG, 1997a

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Method	Results	Remarks	Reference
Reliability 1: GLP-guideline study with analytical verification of test item concentrations			
<p><i>Pimephales promelas</i> - Standard Methods for the Examination of Water and Wastewater, 13th Edition, 1971, American Public Health Assn., NY, NY 10019.</p> <p>Reliability 2: This study was conducted prior to GLP and test guidelines, but sufficient data is available for interpretation of results</p>	maximum safe level without mortality or observable effects (96 h): > 100 mg/L	Test item: Triisopropanolamine, purity: n.d. Due to the high water solubility, the test item was prepared in stock solutions using distilled water. The test was performed under static exposure conditions.	Dow, 1975
invertebrates			
<p><i>Daphnia magna</i> - Directive 79/831/EEC, Annex V, Part C</p> <p>Reliability 2: Non-GLP study in accordance with european standard test guidelines. No analytical test item concentration verification.</p>	EC50 (48 h): > 500 mg/L (nominal), no immobile daphnids observed	Test item: Triisopropanolamine, purity: n.d. A stock solution with a nominal concentration of 500 mg/l was prepared. The test solutions were fixed by serial dilution of the stock solution. The test was performed under static exposure conditions. Test concentrations were 0, 7.81, 15.6, 31.2, 62.5, 125, 250 and 500 mg/L. During the test the pH value ranged from 7.56 to 9.05. Dissolved oxygen ranged from 8.23 to 8.94 mg/L. The test was performed in 4 replicates per test concentration.	BASF AG, 1987c
<p><i>Daphnia magna</i> - EU Method C.2 (Acute Toxicity for Daphnia)</p> <p>Reliability 1: GLP-guideline study with analytical verification of test item concentrations</p>	EC50 (48 h): 857 mg/L (nominal)	Test item: Triisopropanolamine, purity: >98.6% A stock solution with a nominal concentration of 2.03 g/L was prepared. The test solutions were fixed by serial dilution of the stock solution. The test was performed under static exposure conditions. Test concentrations were 0, 120, 180, 250, 350, 500, 700 and 1000 mg/L. The test was performed in 4 replicates per test concentration.	Huels AG, 1997b

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Method	Results	Remarks	Reference
algae			
<p><i>Scenedesmus subspicatus</i> (new name: <i>Desmodesmus subspicatus</i>)</p> <p>- EU Method C.3 (Algal Inhibition test)</p> <p>Reliability 1: GLP-guideline study with analytical verification of test item concentrations</p>	<p>EC50 (72 h): 710 mg/L (growth rate) (nominal)</p> <p>EC50 (72 h): 50 mg/L (cell number) (nominal)</p>	<p>Test item: Triisopropanolamine, purity: >98.6%</p> <p>A stock solution with a nominal concentration of 2.03 g/L was prepared. The test solutions were fixed by serial dilution of the stock solution. The test was performed under static exposure conditions. Test was performed in two sets using the following test substance concentrations: set 1: 4, 10, 26, 64, 160, 400 and 1000 mg/L concentrations were 0, 120, 180, 250, 350, 500, 700 and 1000 mg/L. set 2: 0.2, 0.64 and 1.6 mg/L. The test was performed in 5 replicates</p>	Huels AG, 1997c
<p><i>Scenedesmus subspicatus</i> (new name: <i>Desmodesmus subspicatus</i>) (algae)</p> <p>- DIN 38412, Part 9</p> <p>Reliability 2: Non-GLP study in accordance with european standard test guidelines. No analytical test item concentration verification.</p>	<p>EC50 (72 h): > 100 mg/L (growth rate) (nominal)</p> <p>EC50 (72 h): 64.67 mg/L (biomass) (nominal)</p> <p>Values were recalculated from the fluorimetric data according to OECD 201 using ToxRatPro v2.09</p>	<p>Test item: Triisopropanolamine, purity: n.d.</p> <p>The test was performed under static exposure conditions. The test substance concentrations were: 1.56, 3.13, 6.25, 12.5, 25, 50, 100 mg/L. Additionally a neutralised sample of 100 mg/L was tested. During the test the pH value ranged from 7.97 to 9.60. The test was performed in 4 replicates</p>	BASF AG, 1989, ECT Oekotoxikologie GmbH (2008)

5.4.1 Fish

5.4.1.1 Short-term toxicity to fish

The test substance is not harmful to fish as was demonstrated in a BASF AG study from 1987. The 96 -h LC₅₀ value calculated as geometrical mean is 3158 mg/L. This result is supported by an acute toxicity test according to EU Method C.1 (Acute Toxicity for Fish) from Hüls (1997). In this limit-test no mortality was observed at 1000 mg/L (LC₅₀ (96 h) >1000 mg/L, nominal confirmed by concentration control analytics).

5.4.1.2 Long-term toxicity to fish

No data available

5.4.2 Aquatic invertebrates

5.4.2.1 Short-term toxicity to aquatic invertebrates

A BASF AG study conducted in 1987 indicated that the test substance is also most probably not acutely harmful to aquatic invertebrates. The EC₅₀ based on mobility of *D. magna* was determined to be > 500 mg/L. These results are supported by an acute toxicity test according to EU Method C.2 (Acute Toxicity for *Daphnia*) from Hüls (1997). The EC₅₀ (48 h) was 857 mg/L (nominal, confirmed by concentration control analytics). Short-term toxicity to aquatic invertebrates

5.4.2.2 Long-term toxicity to aquatic invertebrates

No data available

5.4.3 Algae and aquatic plants

The test substance is most probably not acutely harmful to algae as demonstrated by a study sponsored by Sasol Germany GmbH in 1997. In a test according to EU method C.3 an E_rC₅₀ of 710 mg/L was determined. These results are supported by a BASF study conducted in 1990. The E_rC₅₀, recalculated from the fluorescence data, after 72 hours of exposure was determined to be > 100 mg/L.

5.4.4 Other aquatic organisms (including sediment)

None.

5.5 Comparison with criteria for environmental hazards (sections 5.1 – 5.4)

1,1',1''-nitrilotripropan-2-ol is not readily, nor easily or inherently biodegradable in regulatory terms but it rapidly dissipates from the environment due to degradation in surface water/sediment and in soil. The endpoints derived from acute aquatic toxicity studies are > 100 mg/L at each trophic level. Hence, the chemical is considered to be acutely not harmful to aquatic organisms including fish, aquatic invertebrates and algae. The experimentally determined BCF was < 1 indicating that the bioaccumulation potential is low.

Abiotic degradation due to hydrolysis is not expected as was demonstrated by Toropkov (1980). 1,1',1''-nitrilotripropan-2-ol proved to be stable in water. Further, at environmental pH conditions hydrolysis is not expected to be a relevant degradation process due to the absence of hydrolysable groups (Kollig et al. 1993, Boethling and Mackay 2000).

In 1998 the ready biodegradability of the test substance was assessed in an OECD 301 F study performed for DOW Elanco. Biodegradation was not observed during the test period. In a DOC Die Away-Test according to OECD 301 A (Hüls AG, 1997) a biodegradation degree of 18% was measured after 28 d indicating that the chemical is not readily biodegradable. In a MITI test (1992) resembling the test guideline OECD 302 C for inherent biodegradability demonstrated the absence of inherent biodegradation. Further, an OECD 302 B BASF-study from 1981 demonstrated a low potential for elimination from water.

In screening tests 1,1',1''-nitrilotripropan-2-ol was found to be not biodegradable. However, in the water/sediment compartment the test substance has a half-life of 14.3 days under aerobic conditions. After 30 and 60 days of exposure 39% and 64% of the applied radioactivity were recovered as CO₂, respectively, indicating that 1,1',1''-nitrilotripropan-2-ol will not persist in aerobic aqueous compartments. In anaerobic media no biodegradation is observed after 6 month of

exposure. In natural soil 1,1',1''-nitrilotripropan-2-ol is mineralised to an extent of 66 to 72%. Therefore, 1,1',1''-nitrilotripropan-2-ol is not rapidly or inherently biodegradable in regulatory terms but it does not persist in the environment due to degradation in surface water and in soil. However, according to CLP criteria this environmental fate can not account for an alteration of the classification.

Based on the calculated Koc values for charged and uncharged molecules at different pH values ranging from a minimum of 0.014 to a maximum of 10 and the Henrys Law Constant of 0.000001 Pa*m³/mol, the test chemical can be considered as not adsorptive to the solid phase of soil and sediment further it does not evaporate into the air from the water surface.

A study on the bioaccumulation in aquatic organisms (MITI, 1992) demonstrated that the test substance does not accumulate in aquatic organisms. According to CLP criteria a bioaccumulation factor of ≥ 500 and/or a partition coefficient octanol/water (log Kow) of > 4 is indicative of the potential to bioconcentrate for classification purposes. Compared to the experimentally determined bioconcentration factor of < 1 and the measured log Kow of -0.013 classification triggered by bioconcentration is not justified.

The test substance is not harmful to fish as was demonstrated in a BASF AG study from 1987. The 96 -h LC₅₀ value calculated as geometrical mean is 3158 mg/L. This result is supported by an acute toxicity test according to EU Method C.1 (Acute Toxicity for Fish) from Hüls (1997). In this limit-test no mortality was observed at 1000 mg/L (LC₅₀ (96 h) > 1000 mg/L, nominal confirmed by concentration control analytics).

Based on CLP criteria, the low acute toxicity of the test chemical to fish does not trigger a classification of the test substance.

A BASF AG study conducted in 1987 indicated that the test substance is also most probably not acutely harmful to aquatic invertebrates. The EC₅₀ based on mobility of *D. magna* was determined to be > 500 mg/L. These results are supported by an acute toxicity test according to EU Method C.2 (Acute Toxicity for *Daphnia*) from Hüls (1997). The EC₅₀ (48 h) was 857 mg/L (nominal, confirmed by concentration control analytics).

Based on CLP criteria, the low acute toxicity of the test chemical to aquatic invertebrates does not trigger a classification of the test substance.

Finally, the test substance is most probably not acutely harmful to algae as demonstrated by a study sponsored by Sasol Germany GmbH in 1997. In a test according to EU method C.3 an E_rC₅₀ of 710 mg/L was determined. These results are supported by a BASF study conducted in 1990. The E_rC₅₀, recalculated from the fluorescence data, after 72 hours of exposure was determined to be > 100 mg/L.

Based on CLP criteria, the low toxicity of the test chemical to algae does not trigger a classification of the test substance.

Table 25: CLP criteria compared to the reported results

Endpoint	Results	CLP legislation	Classification
Stability in water	Stable in water	4.1.2.9.2: abiotic degradation of $> 70\%$ under environmental conditions 4.1.2.9.4: ...Hydrolysis can be considered if the hydrolysis products do not fulfil the criteria for classification as hazardous to the aquatic environment	no rapid degradability
OECD Guideline 301 F (Ready Biodegradability: Manometric Respirometry Test)	0% BOD/ThOD (28 d)	4.1.2.9.5.(a)(i): 60% after 28 days	no rapid degradability

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OECD guideline 301 A (Ready Biodegradability: DOC Die Away Test)	18% DOC removal (28 d)	4.1.2.9.5.(a)(ii): 70% after 28 days	no rapid degradability
OECD Guideline 302 C (Inherent Biodegradability: Modified MITI Test (II))	0% BOD/ThOD (14 d) 3.4% BOD/ThOD (28 d)	Guidance on the Application of Regulation (EC) No 1272/2008, p. 406: Inherent-(OECD 302) and sewage treatment simulation (OECD 303) tests are not normally used in this context, due to the high levels of adapted biomass.	no rapid degradability
OECD Guideline 302 B (Inherent biodegradability: Zahn-Wellens/EMPA Test)	< 10% DOC (28d)	Guidance on the Application of Regulation (EC) No 1272/2008, p. 406: Inherent-(OECD 302) and sewage treatment simulation (OECD 303) tests are not normally used in this context, due to the high levels of adapted biomass.	no rapid degradability
EPA Subdivision N Pesticide Guideline 162-4 (Aerobic Aquatic Metabolism)	39% ¹⁴ CO ₂ , radiochem. meas. (30d); 64% radiochem. meas. (64d); half-life: 14.3 days	4.1.2.9.3: ...degradation half-lives [...] can be used in defining rapid degradation provided that ultimate biodegradation of the substance, i.e. full mineralisation, is achieved. 4.1.2.9.3: Primary biodegradation does not normally suffice in the assessment of rapid degradability unless it can be demonstrated that the degradation products do not fulfil the criteria for classification as hazardous to the aquatic environment.	no rapid degradability
EPA Subdivision N Pesticide Guideline 162-3 (Anaerobic Aquatic Metabolism)	< 1% ¹⁴ CO ₂ , radiochem. meas. (6m)	Guidance on the Application of Regulation (EC) No 1272/2008, p. 406: Anaerobic degradation tests (OECD 311/ISO 11734 and analogous tests) do not qualify because of the specificity of the anaerobic compartments.	no rapid degradability
EPA 162-1: Aerobic soil metabolism study	66-72% ¹⁴ CO ₂ , radiochem. meas. (20d)	Guidance on the Application of Regulation (EC) No 1272/2008, p. 459; II.2.3.6.(c): ...the substance is ultimately degraded within 28 days with a half-life < 16 days corresponding to a degradation rate > 0.043 day ⁻¹	rapid degradability
Conclusion: Only the study on the degradation of the test substance in soil demonstrated rapid degradability. However, any other test on degradation presented demonstrates the lack of rapid degradation and hence, the test substance is not considered to rapidly degrade in the environment			
OECD Guideline 305 C (Bioaccumulation: Test for the Degree of Bioconcentration in Fish) Species: <i>Cyprinus carpio</i>	BCF <0.57 (0.25 mg/L); BCF <0.06 (2.5 mg/L)	4.1.2.8.1: A BCF in fish of ≥ 500 is indicative of the potential to bioconcentrate for classification purposes.	not bioaccumulative
The test substance does not fulfil the criteria for bioaccumulation potential			

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<i>Leuciscus idus</i> - DIN 38412, Part 11	LC50 (96 h): 3158.48 mg/L (geometric mean; nominal)	4.1.2.6.; Table 4.1.0	
<i>Cyprinus carpio</i> - EU Method C.1 (Acute Toxicity for Fish; limit test)	LC50 (96 h): > 1000 mg/L (nominal)		
<i>Pimephales promelas</i> - Standard Methods for the Examination of Water and Wastewater, 13th Edition, 1971, American Public Health Assn., NY, NY 10019.	maximum safe level without mortality or observable effects (96 h): > 100 mg/L		
<i>Daphnia magna</i> - Directive 79/831/EEC, Annex V, Part C	EC50 (48 h): > 500 mg/L (nominal), no immobile daphnids observed		
<i>Daphnia magna</i> - EU Method C.2 (Acute Toxicity for Daphnia)	EC50 (48 h): 857 mg/L (nominal)		
<i>Scenedesmus subspicatus</i> (new name: <i>Desmodesmus subspicatus</i>) - EU Method C.3 (Algal Inhibition test)	EC50 (72 h): 710 mg/L (growth rate) (nominal) EC50 (72 h): 50 mg/L (cell number) (nominal)		
<i>Scenedesmus subspicatus</i> (new name: <i>Desmodesmus subspicatus</i>) (algae) - DIN 38412, Part 9	EC50 (72 h): > 100 mg/L (growth rate) (nominal) EC50 (72 h): 64.67 mg/L (biomass) (nominal)		
			Since all relevant available data on the acute toxicity are above the trigger value of 100 mg/L the test substance is not considered to be harmful to aquatic life

5.6 Conclusions on classification and labelling for environmental hazards (sections 5.1 – 5.4)

The summarised results above combined with the high water solubility and the low bioconcentration factor demonstrate that the classification Aquatic Chronic 3 is not justified.

RAC evaluation of hazardous to the aquatic environment

Summary of dossier submitter's proposal

The dossier submitter proposed to remove the current Annex VI classification as hazardous to the aquatic environment, categories Chronic 3 - H412, according to the Regulation (EC) 1272/2008

(CLP), and R52/53, according to Directive 67/548/EEC (DSD).

The current environmental classification resulted from two acute toxicity algae studies, providing E_bC_{50} (algae) < 100 mg/l (based on number of cells and biomass) and from biodegradation screening tests, indicating not rapid biodegradability.

The dossier submitter argues that according to the CLP Regulation the classification should be based on the algal growth rate, and hence the E_rC_{50} should be used instead of the E_bC_{50} .

Since E_rC_{50} (algae) is higher than 100 mg/l in both acute toxicity studies, the dossier submitter considers the substance as not toxic to the aquatic environment. This is consistent with the L(E) C_{50} values for the other tested trophic levels, fish (three species) and crustaceans (one species), which are all also above the trigger value of 100 mg/l.

Under environmental conditions the substance is partly present in its charged form and it is shown to be hydrolytically stable.

Screening tests show that the substance is neither readily nor inherently biodegradable. A simulation tests in a water-sediment system provides a half-life of 14.3 days but for the identified metabolite no information is available on its toxicity and degradability. Another simulation test in soil shows that the substance is mineralised by 66 to 72% in 20 days. Its metabolite is considered to be not toxic and readily biodegradable. Based on all this information, the dossier submitter concludes that the substance is not rapidly degradable.

The water solubility is high (830 g/l) and the experimentally determined BCF is < 0.57, thus indicating no potential to bioaccumulate. Considering also the absence of acute toxicity, the dossier submitter concludes that the substance does not meet the criteria for aquatic Chronic 3 (CLP) and R52/53 (DSD).

After the public consultation, the dossier submitter has resubmitted a new version of the CLH report, which implements the changes (see section “Comments received during public consultation”) proposed by some commenting party. This report is provided as an annex to the RCOM document.

Comments received during public consultation

During the public consultation, comments on hazards to the aquatic environment were received from four Member States.

The comments supported the proposal for removal of the classification. Besides editorial changes, some concern was expressed by a commenting Member State about the table of physico-chemical properties (table 9) since the method used were mostly not indicated and the values for K_{oc} were not present. This comment however does not effect the proposed classification.

For the full set of comments and responses, see the response to comments document (RCOM).

Detailed description on relevant arguments and information received during the public consultation

Not needed.

Outcome of RAC assessment - comparison with criteria and justification

RAC agrees with the proposal of the dossier submitter to remove the classification for aquatic hazard and supports the provided justification and comparison with criteria, summarized in the first heading.

Extended analysis of key information

The key studies provide both the E_rC₅₀ and E_bC₅₀ values as following:

Used algae and test method	Measured endpoint	Test item	Reference
<i>Scenedesmus subspicatus</i> (<i>Desmodesmus subspicatus</i>) EU Method C.3 Algal Inhibition test	EC50 (72 h): 710 mg/L (growth rate) (nominal) EC50 (72 h): 50 mg/L (cell number) (nominal)	Triisopropanolamine, purity: >98.6%	Huels AG, 1997c
<i>Scenedesmus subspicatus</i> (<i>Desmodesmus subspicatus</i>) DIN 38412, Part 9	EC50 (72 h): > 100 mg/L (growth rate) (nominal) EC50 (72 h): 64.67 mg/L (biomass) (nominal)	Triisopropanolamine, purity: n.d.	BASF AG, 1989, ECT Oekotoxikologie GmbH (2008)

The comparison of biomass and growth rate inhibition percentages (as in the key study of Huels AG 1997c and reported in the IUCLID 5 data set) shows that they present clear differences: **52%** of inhibition in biomass appears at **64 mg TIPA/L**, whilst **50%** of inhibition in growth rate occurs between 400 and 1000 mg TIPA/L. The data are **shown in the following table**:

Concentration [mg/L]	Biomass (area under the growth curve)	Inhibition of biomass [%]	Growth rate μ (0–72 h)	Inhibition of growth rate [%]
Control	89.5	-	1.356	-
4	73.5	17.9	1.28	5.6
10	70.0	21.8	1.254	7.5
26	52.5	41.3	1.139	16.0
64	43.0	52.0	1.059	21.9
160	34.5	61.5	0.954	29.6
400	19.5	78.2	0.75	44.7
1000	10.5	88.3	0.568	58.1

This finding is confirmed in BASF AG (1989) and ECT Oekotoxikologie GmbH (2008) supporting studies (reported in the IUCLID data set), summarized in the following table:

Duration	Endpoint	Effect conc. mg/L	Nominal/measured	Basis for effect	Recalculation by
72 h	EC10	19.47	nominal	growth rate	ToxRatPro
72 h	EC20	62.46	nominal	growth rate	ToxRatPro
72 h	EC50	>100	nominal	growth rate	ToxRatPro
72 h	EC10	7.14	nominal	biomass	ToxRatPro
72 h	EC20	15.26	nominal	biomass	ToxRatPro
72 h	EC50	64.67	nominal	biomass	ToxRatPro

Based on this data, RAC supports the proposal of the dossier submitter.

6 OTHER INFORMATION

7 REFERENCES

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BASF AG, 1972: Stoffwertmessungen, testing laboratory: BASF AG, Department of process engineering, report-no.: 172.096.1, report date: 1972-04-06

BASF AG, 1978: Sicherheitstechnische Kenndaten, testing laboratory: BASF AG, TLM/SIK - B 14, report-no.: SIK 78/0966, report date: 1978-10-20

BASF AG, 1981: Standversuch; unpublished data

BASF AG, 1987a: Octanol-Wasser-Verteilungskoeffizient POW von fünf Alkanolaminen bei 23 ° C, testing laboratory: BASF AG, Department of analytics, report-no.: BRU 87.262, report date: 1987-12-18

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