**FOREWORD** 

**INTRODUCTION** 

# **1-Aminoanthraquinone**

# CAS N°: 82-45-1

# **SIDS Initial Assessment Report**

# For

# SIAM 4

## Tokyo, Japan, 20-22 May 1996

1. Chemical Name: 1-Aminoanthraquinone

Japan

- **2. CAS Number:** 82-45-1
- 3. Sponsor Country:

National SIDS Contact Point in Sponsor Country: Mr. Yasuhisa Kawamura, Ministry of Foreign Affairs, Japan

#### 4. Shared Partnership with:

- 5. Roles/Responsibilities of the Partners:
- Name of industry sponsor /consortium
- Process used

#### 6. Sponsorship History

 How was the chemical or category brought into the OECD HPV Chemicals Programme ?
 As a high priority chemical for initial assessment, 1-aminoanthraquinone was selected in the framework of the HPV Programme.

SIDS Dossier and Testing Plan were reviewed at a SIDS Review Meeting in 1993, where the following SIDS Testing Plan was agreed.

No testing ()

Testing(X) Physical-Chemical Properties Vapour pressure Partition coefficient Water solubility Environmental fate/Biodegradation Biodegradation Bioaccumulation Photodegradation Stability in water Ecotoxicity

> Acute toxicity to fish Acute toxicity to daphnids

Toxicity to algae Chronic toxicity to daphnids

Toxicity

Acute dermal toxixity Repeated dose toxicity Reproductive toxicity Gene mutation Chromosomal aberration Genetic toxicity in vivo

Original report already circulated in August 1995, and the report was revised according to the comments from member countries.

- 7. Review Process Prior to the SIAM:
- 8. Quality check process:
- 9. Date of Submission: April 30, 1996
- 10. Date of last Update:
- 11. Comments:

## SIDS INITIAL ASSESSMENT PROFILE

CAS No.	82-45-1
Chemical Name	9,10-Anthracenedione, 1-amino-
Structural Formula	O NH <sub>2</sub>

#### CONCLUSIONS AND RECOMMENDATIONS

A potential hazard to the environment due to toxicity to algae is identified, but exposure is low in the sponsor country.

Unless further information on exposure in other Member countries presents evidence to the contrary, it is currently considered of low potential risk and low priority for further work.

# SHORT SUMMARY WHICH SUPPORTS THE REASONS FOR THE CONCLUSIONS AND RECOMMENDATIONS

Production volume of 1-aminoanthraquinone in Japan is ca. 1,000 - 2,000 tonnes/year in 1990-1993. This chemical is used as intermediates for dyes and pharmaceuticals in closed systems in Japan. This chemical is stable in neutral, acidic or alkaline solutions, and is considered as "not readily biodegradable". Direct photodegradation is expected as this chemical absorbs UV light with half-life of about one week.

The potential environmental distribution of the chemical obtained from a generic fugacity model (Mackey level III) showed that the chemical would be distributed mainly into water and soil. Predicted environmental concentration ( $PEC_{local}$ ) of this chemical was estimated as  $1.7 \times 10^{-4}$  mg/l from Japanese local exposure scenario.

For the environment, various NOEC and LC<sub>50</sub> values were gained from test results; LC<sub>50</sub> => 1000 mg/l (acute fish); EC<sub>50</sub> => 1000 mg/l (acute daphnia); EC<sub>50</sub> = 0.25 mg/l (acute algae); NOEC = 0.10 mg/l (acute algae); NOEC = 0.32 mg/l (long-term daphnia reproduction). From the lowest toxicity data to algae, acute-NOEC of *Algae* (0.1 mg/l) was adopted for the calculation of PNEC. The assessment factor of 100 was used to both acute and chronic toxicity data to determine PNEC according to the OECD Provisional Guidance for Initial Assessment of Aquatic Effects. Thus, PNEC of the chemical is 0.001 mg/l in the present report. The PEC is lower than the PNEC, therefore environmental risk is presumably low.

As 1-aminoanthraquinone is produced in a closed system, exposure during synthesis may be excluded. The product is filled into barrels under the local exhaust ventilation. Inhalation at work place is considered to be main exposure route while skin contact plays a minor role. However workers wear personal protective equipment (e.g. safety glasses, dust respirator, rubber gloves) during the filling process. Therefore, the exposure at work place is considered to be negligible at present situation. In addition, this chemical is not contained in consumer products, because it is an intermediate in industrial use.

Although the chemical showed positive results only in *S. typhimurium* TA 1537 with metabolic activation, negative results were obtained by other bacterial strains and chromosomal aberration tests *in vitro* and *in vivo*. In a combined repeat dose and reproductive/developmental toxicity screening test, several toxicological findings in kidney and spleen were observed at the lowest dose (eosinophilic droplet/body [kidney], nephropathy [spleen]). The parental animals exhibited no effects on reproductive parameters such as fertility index. However, nursing behaviour disappeared in all of the treatment female groups. Viability of pups on day 4 after birth was decreased in all treatment groups. Therefore, NOEL was less than 40 mg/kg/day both for repeated dose toxicity and reproductive

toxicity.

As for indirect exposure via environment, PEC was estimated as  $1.7 \times 10^{-4}$  mg/l from local exposure scenario. For human health, although NOEL is estimated as less than 40 mg/kg/day for both repeated dose and reproductive toxicity, the margin of safety is large. Therefore, health risk through the environment, in general, is considered to be presumably low due to its use pattern and exposure situation.

In conclusion, no further testing is needed at present considering its toxicity and exposure levels.

#### NATURE OF FURTHER WORK RECOMMENDED

CAS NO	): 82-45-1	SPECIES	PROTOCOL	RESULTS
PHYSIC	CAL-CHEMICAL			
2.1	Melting Point			256 – 258 °C
2.2	<b>Boiling Point</b>			> 300 °C
2.3	Density			No data available
2.4	Vapour Pressure		OECD TG 104	1.2 x 10 <sup>-4</sup> Pa at 100 °C
2.5	Partition Coefficient (Log Pow)		OECD TG 107	3.74 at 25 °C
2.6 A.	Water Solubility		OECD TG 105	32 mg/L at 25 °C
B.	рН			No data available.
	рКа			Na data available.
2.12	Oxidation: Reduction Potential			No data available.
ENVIR AND PA	ONMENTAL FATE ATHWAY			
3.1.1	Photodegradation		Estimation	$T_{1/2} = 1.4 \text{ x } 10^{-2} \text{ y}$ (direct photolysis in water)
3.1.2	Stability in Water		OECD TG 111	Stable at pH 4.0, 7.0 and 9.0.
3.2	Monitoring Data			Not detected from surface water and sediment in Japan
3.3	Transport and Distribution		Calculated (Fugacity Level III)	100% released to water,           In Air         0.04%           In Water         62.57%           In Soil         21.34%           In Sediment         16.06%
3.5	Biodegradation		OECD TG 301C	Not readily biodegradable: 0 - 1 % (BOD) in 28 days, 1 - 3% (HPLC) in 28 days
3.6	Bioaccumulation	Carp	OECD TG 305C	BCF: 50 – 150
ЕСОТС	OXICOLOGY			
4.1	Acute/Prolonged Toxicity to Fish	Oryzias latipes	OECD TG 203	$\begin{split} & LC_{50}(72hr):>1,\!000~mg/L\\ & LC_{50}(96hr):>1,\!000~mg/L \end{split}$
4.2	Acute Toxicity to Aquatic Invertebrates ( <i>Daphnia</i> )	Daphnia magna	OECD TG 202	EC <sub>50</sub> (48hr): > 1,000 mg/l
4.3	Toxicity to Aquatic Plants e.g. Algae	Selenastrum capricornutum	OECD TG 201	(biomass method) EC <sub>50</sub> (72hr): 0.25 mg/l NOEC: 0.1 mg/l
4.5.2	Chronic Toxicity to Aquatic Invertebrates ( <i>Daphnia</i> )	Daphnia magna	OECD TG 202	$EC_{50}$ (21d, Mortality): 0.62 mg/l $EC_{50}$ (21d, Reproduction): 0.56 mg/l NOEC (21d, Repro): 0.32 mg/l
4.6.1	Toxicity to Soil Dwelling Organisms			No data available.
4.6.2	Toxicity to Terrestrial Plants			No data available.

## FULL SIDS SUMMARY

CAS NO: 82-45-1		SPECIES	PROTOCOL	RESULTS
(4.6.3)	Toxicity to Other Non- Mammalian Terrestrial Species (Including Birds)			No data available
TOXIC	OLOGY			
5.1.1	Acute Oral Toxicity	Rat	OECD TG 401	LD <sub>50</sub> > 5,000 mg/kg
5.1.2	Acute Inhalation Toxicity			No data available.
5.1.3	Acute Dermal Toxicity	Mouse		LD <sub>50</sub> > 2,000 mg/kg
5.4	Repeated Dose Toxicity	Rat	OECD Combined Test	NOAEL < 40 mg/kg/day
5.5	Genetic Toxicity In Vitro			
А.	Bacterial Test (Gene mutation)	Styphimurium E. coli	OECD Guidelines No.471 and 472 and Japanese Guideline	TA1537: Positive in TA1537 with metabolic activation Other bacterial strain: Negative (With And without metabolic activation)
B.	Non-Bacterial In Vitro Test (Chromosomal aberrations)	CHL cells	OECD Guideline No.473 and Japanese Guideline	Negative (With metabolic activation) Negative (Without metabolic activation)
5.6	Genetic Toxicity In Vivo	Mouse	Micronucleus test	Negative
5.8	Toxicity to Reproduction	Rat	OECD Combined Test	NOAEL Parental = < 40 mg/kg/day NOAEL F1 offspring = < 40 mg/kg/day
5.9	Developmental Toxicity/ Teratogenicity			
5.11	Experience with Human Exposure			

# **SIDS Initial Assessment Report**

## **1 IDENTITY**

#### 1.1 Identification of the Substance

CAS Number:	82-45-1
IUPAC Name:	9,10-Anthracenedione, 1-amino-
Molecular Formula:	$C_{14}H_9NO_2$
Structural Formula:	O NH <sub>2</sub>



1-Aminoanthraquinone

## 1.2 Purity/Impurities/Additives

Degree of Purity:	> 97 %
Major Impurities:	Anthraquinone
Essential Additives:	None

#### **1.3** Physico-Chemical properties

	Table 1	Summary	of physico-	chemical	properties
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Property	Value
Melting point	256-258 °C
Boiling point	> 300 °C
Vapour pressure	1.2 x 10 <sup>-4</sup> Pa at 100 °C
Water solubility	32 mg/l
Partition coefficient n-octanol/water (log value)	3.74

## 2 GENERAL INFORMATION ON EXPOSURE

The production level of 1-aminoanthraquinone in Japan was about 1,000 - 2,000 tonnes/year. Most of this amount was sold and handled in Japan. This chemical is used as an intermediate for dyestuff and pharmaceuticals in closed systems. Release into the environment may occur at the production site or specific industrial sites. All disposal wastes are treated by wastewater treatment or incineration. 1-Aminoanthraquinone seems to be released into water and air from its production sites after biological treatment. In a Japanese company, about 1.9 tonnes/year are released into water from the production site. In a Japanese monitoring program by the Environment Agency, this chemical was not detected in the general environment in 1987. No specific local monitoring data of the chemical is available. 1-Aminoanthraquinone is not readily biodegradable (OECD 301C: 0% after 28d). 1-Aminoanthraquinone is not hydrolyzed at pH 4, 7 and 9. Direct photodegradation is

expected because 1-aminoanthraquinone absorbs UV light. The half-life in water is estimated to be about a week.

## 2.1 Environmental Exposure and Fate

## 2.1.1 Estimates of environmental fate, pathway and concentration

### **Global exposure**

The potential environmental distribution of 1-aminoanthraquinone obtained from a generic level III fugacity model is shown in Table 2. The results show that if 1-aminoanthraquinone is released mainly into soil or air, it is likely to distribute into the soil compartment. But if 1-aminoanthraquinone is released mainly into water, it is likely to be transported to both soil and sediment. Due to the low vapour pressure of 1-aminoanthraquinone, it is unlikely to distribute into air.

Compartment	Release: 100% to air	Release: 100% to water	Release: 100% to soil
Air	0.18%	0.04%	0.00%
Water	0.60%	62.57%	0.47%
Soil	99.06%	21.34%	99.41%
Sediment	0.15%	16.06%	0.12%

**Table 2**. Environmental distribution 1-aminoanthraquinone using a generic level III fugacity model.

## Local exposure

According to a Japanese manufacturer, 1,900 kg/year (measured) of 1-aminoanthraquinone are released with  $1.10 \times 10^7$  t/y of effluent into a bay. The local predicted environmental concentration (PEC<sub>local</sub>) is  $1.7 \times 10^{-4}$  mg/l, employing the following calculation model:

Amount of release  $(1.90 \times 10^9 \text{ mg/y})$ 

Volume of effluent  $(1.10 \times 10^{10} \text{ l/y}) \times \text{Dilution factor } (1,000)$ 

## 2.1.2 Photodegradation

A half-life time of  $1.44 \times 10^{-2}$  years is estimated for the direct photodegradation of 1-aminoanthraquinone in water. (MITI, Japan).

## 2.1.3 Stability in Water

The chemical is stable in water at pH 4, 7 and 9 at  $25^{\circ}$ C (OECD TG 111).

## 2.1.4 Biodegradation

If released into water, this substance does not readily biodegrade (MITI (I), corresponding to the OECD 301C: 0 - 1 % after 28 days based on BOD and 1 - 3 % based on HPLC analysis).

## 2.1.5 Bioaccumulation

BCF= 50 - 150 in carp (8 weeks at 25 °C) suggests that the potential for bioconcentration in aquatic organisms is low.

## 2.2 Human Exposure

## 2.2.1 Occupational Exposure

As 1-aminoanthraquinone is produced in a closed system, exposure during synthesis may be excluded. This chemical is used as intermediate for dyestuffs. The product is poured into barrels under local exhaust ventilation. Inhalation uptake is considered to be the main exposure route. Skin contact plays a minor role. Workers wear safety glasses, dust respirators, and protective gloves during the filling process. Therefore, the exposure to workers is estimated to be negligible at the present situation.

## 2.2.2 Consumer Exposure

1-Aminoanthraquinone is not contained in consumer products, because the substance is an intermediate for dyestuffs. No other information on uses is available.

## 2.2.3 Exposure via the Environment

The highest exposure to the general population via the environment would be expected through drinking water processed from surface water. Based on physical chemical properties of 1-aminoanthraquinone, a significant removal during processing is not to be expected. Although reliable  $PEC_{global}$  cannot be estimated, the concentration in drinking water is assumed to be  $1.7 \times 10^{-4}$  mg/l as a worst case.

## **3** HUMAN HEALTH HAZARDS

## 3.1 Effects on Human Health

## 3.1.1 Acute Toxicity

 $LD_{50}s$  from acute oral toxicity studies in rats were reported as > 5,000 mg/kg or >1,600 mg/kg. Also, the  $LD_{50}$  in an acute dermal toxicity study in mice was reported to be > 2,000 mg/kg.

#### **3.1.2** Repeated Dose Toxicity

There is only one key study on repeated dose toxicity of 1-aminoanthraquinone. This chemical was studied for oral toxicity in rats according to the OECD combined repeated dose and reproductive/developmental toxicity test [OECD TG 422]. As the study was well controlled and conducted under GLP, this was considered to be a key study. Male and female SD rats were orally administered (gavage) at doses of 0, 40, 200 and 1,000 mg/kg/day.

Increased spleen weights were observed in males in the 200 mg/kg group and above, as well as females in the 1000 mg/kg group. Also, relative liver weight was increased in males in the 200 mg/kg group and above. Regarding hemato-morphological examination, erythrocyte count, hemoglobin and mean corpuscular hemoglobin was decreased in males in the 200 mg/kg group and above. In clinical chemistry, the potassium concentration in males in the 1000 mg/kg group, and the

chlorine concentration in males in the 200 mg/kg group decreased. In histopathological examination, formation of the eosinophilic droplet and eosinophilic body in kidneys was increased in males in the 40 mg/kg group and above. Nephropathy and dark coloration of the spleen were observed in both males and females in the 40 mg/kg groups.

The NOEL is estimated to be less than 40 mg/kg/day for repeated dose toxicity.

### 3.1.3 Mutagenicity

#### In vitro Studies

A reverse gene mutation assay was conducted in line with Guidelines for Screening Mutagenicity Testing of Chemicals (Japan) and OECD Test Guidelines 471 and 472, using the pre-incubation method. This study was well controlled and regarded as a key study.

Although 1-aminoanthraquinone showed positive results in *S. typhimurium* TA1537 with metabolic activation, negative results were obtained with other bacterial strains at concentrations up to 5 mg/plate with or without a Metabolic activation system (MHW, 1993).

A chromosomal aberration test in line with Guidelines for Screening Mutagenicity Testing of Chemicals (Japan) and OECD Test Guideline 473 was conducted using cultured Chinese Hamster lung (CHL/IU) cells. This study was well controlled and regarded as a key study. The maximum concentration of the chemical was used with no apparent cytotoxic effect in continuous treatment. Neither structural chromosomal aberrations nor polyproidy were recognized up to a maximum concentration of 2.2 mg/ml under conditions of both continuous treatment and short-term treatment with or without an exogeneous metabolic activation system (MHW, 1998).

In vivo Studies

One test result is available on *in vivo* genotoxic effects. A micronucleus test in mice was reported as having negative results. No further information is provided (Bayer AG).

#### 3.1.4 Carcinogenicity

There is some carcinogenicity data, but the data is inadequate.

#### 3.1.5 Toxicity for Reproduction

1-Aminonaphthoquinone was studied for oral toxicity in rats according to the OECD combined repeated dose and reproductive/developmental toxicity test [OECD TG 422] at doses of 0, 40, 200 and 1,000 mg/kg/day. The parental animals exhibited no effects on reproductive parameters including copulation index, fertility index, gestation length, number of corpora lutea or implantation, implantation index, gestation index, delivery index, parturition or maternal behavior. However, nursing behavior disappeared in all of the treatment female groups. Viability of pups on day 4 after birth was decreased in all treatment groups. No external or skeletal anomalies related to the test substance administration were detected in any of the offspring. Furthermore, there are no significant differences in the number of offspring or live offspring, sex ratio, live birth index or body weights.

NOEL is estimated to be less than 40 mg/kg/day for reproductive toxicity.

### 3.2 Initial Assessment for Human Health

As 1-aminoanthraquinone is produced in a closed system, exposure during synthesis may be excluded. The product is poured into barrels under local exhaust ventilation. Inhalation in the workplace is considered to be the main exposure route while skin contact plays a minor role. However, workers wear personal protective equipment (e.g. safety glasses, dust respirators, rubber gloves) during the filling process. Therefore, the exposure in the workplace is considered to be negligible at present. In addition, this chemical is not contained in consumer products.

Although the chemical showed positive results only in *S. typhimurium* TA 1537 with metabolic activation, negative results were obtained by other bacterial strains and chromosomal aberration tests *in vitro* and *in vivo*. In a combined repeat dose and reproductive/developmental toxicity test, several toxicological findings in the kidney and spleen were observed at the lowest dose (eosinophilic droplet [kidney], nephropathy [spleen]). The parental animals exhibited no effects on reproductive parameters such as fertility index. However, nursing behaviour disappeared in all of the treatment female groups. Viability of pups on day 4 after birth was decreased in all treatment groups. Therefore, the NOEL was less than 40 mg/kg/day for both repeated dose toxicity and reproductive toxicity.

For human health, the NOEL is estimated to be 40 mg/kg/day for repeated dose and 40 mg/kg/day for reproductive toxicity. As for indirect exposure via the environment, the PEC was estimated to be  $1.7 \times 10^{-4}$  mg/l from a local exposure scenario. The margin of safety is large. Therefore, health risk through the environment, in general, is considered to be presumably low due to its use pattern and exposure situation.

## 4 HAZARDS TO THE ENVIRONMENT

#### 4.1 Aquatic Toxicity

1-Aminoanthraquinone has been tested in a limited number of aquatic species (*Selenastrum capricornutum*, *Daphnia magna* and *Oryzias latipes*), under OECD test guidelines [OECD TG 201, 202, 203, 204 and 211]. Acute and chronic toxicity data to test organisms for 1-aminoanthraquinonw are summarized in Table 3. No other ecotoxicological data are available.

Various NOEC and LC<sub>50</sub> values were gained from the above-mentioned tests; 96h LC<sub>50</sub> = >1,000 mg/l (acute fish); 24h EC<sub>50</sub> = > 1,000 mg/l (acute daphnia); 72h EC<sub>50</sub> = 0.25 mg/l (acute algae); NOEC = 0.1 mg/l (algae); 21d NOEC = 0.32 mg/l (long-term daphnia reproduction). Therefore, the chemical is considered to be strongly toxic to algae and daphnids (long-term) and non-toxic to fish. As the lowest toxicity result, the NOEC for algae (0.1 mg/l) was adopted. An assessment factor of 100 is applied. Thus the PNEC of 1-aminoanthraquinone is 0.001 mg/l. Since the PEC is lower than the PNEC, environmental risk is presumably low.

Species	Endpoint <sup>*1</sup>	Conc. (mg/L)	Reference
Selenastrum capricornutum (algae)	Biomass: EC <sub>50</sub> (72h) Biomass: NOEC	0.25 mg/L 0.10 mg/L	
Daphnia magna (water flea)	Imm: $EC_{50}(48h)$ Imm: $EC_{50}(21d)$ Rep: $EC_{50}(21d)$ Rep: NOEC	> 1,000 mg/L 0.62 mg/L 0.56 mg/L 0.32 mg/L	EA Japan. (1992)
Oryzias latipes (fish, Medaka)	Mor: LC <sub>50</sub> (24h) Mor: LC <sub>50</sub> (72h) Mor:LC <sub>50</sub> (96h)	> 1,000 mg/L > 1,000 mg/L > 1,000 mg/L	

**Table 3.** Acute and chronic toxicity data of 1,4-diethylbenzene to aquatic organisms.

Notes: <sup>\*1</sup> Mor; mortality, Rep; reproduction.

#### 4.2 Initial Assessment for the Environment

The production volume of 1-aminoanthraquinone in Japan is ca. 1,000 - 2,000 tonnes/year in 1990 - 1993. This chemical is used as an intermediate for dyes and pharmaceuticals in closed systems in Japan. This chemical is stable in neutral, acidic or alkaline solutions, and is considered to be "not readily biodegradable". Direct photodegradation is expected as this chemical absorbs UV light with a half-life in water of about one week.

The potential environmental distribution of the chemical obtained from a generic fugacity model (Mackey level III) showed the chemical will be distributed mainly to water and soil. The predicted environmental concentration ( $PEC_{local}$ ) for this chemical was estimated to be 1.7 x 10<sup>-4</sup> mg/l from a Japanese local exposure scenario.

For the environment, various NOEC and LC<sub>50</sub> values were gained from test results; 96h LC<sub>50</sub> = > 1000 mg/l (acute fish); 24h EC<sub>50</sub> = > 1000 mg/l (acute daphnia); 72h EC<sub>50</sub> = 0.25 mg/l (acute algae); NOEC = 0.10 mg/l (algae); 21d NOEC = 0.32 mg/l (long-term daphnia reproduction). As the lowest toxicity result, the NOEC for algae (0.1 mg/l) was adopted.. An assessment factor of 100 is used to determine a PNEC according to the OECD Provisional Guidance for Initial Assessment of Aquatic Effects. Thus, the PNEC of the chemical is 0.001 mg/l in the present report. Because the PEC is lower than the PNEC, environmental risk is presumably low.

## **5 RECOMMENDATIONS**

A potential hazard to the environment due to toxicity to algae is identified, but exposure is low in the sponsor country.

Unless further information on exposure in other Member countries presents evidence to the contrary, it is currently considered of low potential risk and low priority for further work.

#### **6 REFERENCES**

EA, Japan (1987) Environment Monitoring of Chemicals - Environmental Survey Report of F.Y. 1986 and 1987 (Office of Health Studies, Environmental Health Department, EA, Japan)

EA, Japan (1994) "Investigation of the Ecotoxicological Effects of OECD High Production Volume Chemicals", Office of Health Studies, Environmental Health Department, Environment Agency, Japan (HPV/SIDS Test conducted by EA, Japan)

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Laham S. et al., Toxicol. Appl. Pharmacol. 8, 346 (1966)

Loeser E., Bayer AG data, short report, 11. 8. 1978

Lyman, W.J, W. F. Reehl and D. H. Rosenblatt (1981) "Handbook of Chemical Property Estimation Method", McGraw Hill Book Co.

MHW, Japan (1994a) Unpublished Report on Combined Repeat Dose and Reproductive/ Developmental Toxicity Screening Test of 1-aminoanthraquinone. (HPV/SIDS Test conducted by MHW, Japan)

MHW, Japan (1994b) Unpublished Report on Mutagenicity Test of 1-aminoanthraquinone. (HPV/SIDS Test conducted by MHW, Japan)

MITI, Japan (1994a): Unpublished data

MITI, Japan (1994b) Unpublished Report (HPV/SIDS Test conducted by MITI, Japan. Test was performed in Chemicals Inspection and Testing Institute, Japan)

# **SIDS DOSSIER**

# 9,10-Anthracenedione, 1-amino-

# CAS No. 82-45-1

Sponsor Country: Japan

1.01 A.	CAS No.	82-45-1
1.01 C.	CHEMICAL NAME ( OECD Name)	9,10-Anthracenedione, 1-amino-
1.01 D.	CAS DESCRIPTOR	Not applicable
1.01 G.	STRUCTURAL FORMULA	O NH <sub>2</sub>
	OTHER CHEMICAL IDENTITY INFORMATION	
1.5	QUANTITY	In Japan, 1,000 - 2,000 tonnes/year in 1990 - 1993.
1.7	USE PATTERN	In Japan, Intermediate for dyestuffs and pharmaceuticals Closed system
1.9	SOURCES AND LEVELS OF EXPOSURE	In Japan, 1. Amount released from production site to water is 1.9 tonnes/year All of the waste water is incinerated
ISSUES FOR DISCUSSION (IDENTIFY, IF ANY)		

## SIDS PROFILE

## SIDS SUMMARY

#### 1-Aminoanthraquinone

	CAS NO: 82-45-1		x					50
		Information	OECD Stud	GLP	Other Study	Estimation Method	Acceptable	SIDS Testing Required
	STUDY	Y/N	Y/N	Y/N	Y/N	Y/N	Y/N	Y/N
	PHYSICAL-CHEMICAL DATA							
2.1 2.2 2.3 2.4 2.5 2.6	Melting Point Boiling Point Density Vapour Pressure Partition Coefficient Water Solubility pH and pKa values	Y Y N N N N	N N N	N N N	Y Y Y	N N N	Y Y Y	N N Y Y N
	OTHER P/C STUDIES RECEIVED							
ENV	IRONMENTAL FATE and PATHWAY							
3.1.1 3.1.2 3.2 3.3 3.5 3.6	Photodegradation Stability in water Monitoring data Transport and Distribution Biodegradation Bioaccumulation	N N N N Y	Y	Y	N	N	Y	Y Y N Y N
OTHER ENV FATE STUDIES RECEIVED								
	ECOTOXICITY							
4.1 4.2 4.3 4.5.2 4.6.1 4.6.2 4.6.3	Acute toxicity to Fish Acute toxicity to Daphnia Toxicity to Algae Chronic toxicity to Daphnia Toxicity to Soil dwelling organisms Toxicity to Terrestrial plants Toxicity to Birds	N N N N N						Y Y Y N N N
OTHER ECOTOXICITY STUDIES RECEIVED								
	ΤΟΧΙΟΙΤΥ							
5.1.1 5.1.2 5.1.3 5.4 5.5 5.6 5.8 5.9 5.11	Acute Oral Acute Inhalation Acute Dermal Repeated Dose Genetic Toxicity <i>in vitro</i> . Gene mutation . Chromosomal aberration Genetic Toxicity <i>in vivo</i> Reproduction Toxicity Development / Teratogenicity Human experience	N N N N N N N N N						Y N Y Y Y N Y N
OT	THER TOXICITY STUDIES RECEIVED							

### OECD SIDS 1. GENERAL INFORMATION

#### 1.01 SUBSTANCE INFORMATION

А.	CAS-Number	82-45-1
B.	Name (IUPAC name)	1-Aminoanthraquinone
C.	Name (OECD name)	9,10-Anthracenedione, 1-amino-
D.	CAS Descriptor	Not applicable
E.	EINECS-Number	201-423-5

F. Molecular Formula C<sub>14</sub>H<sub>9</sub>NO<sub>2</sub>

G. Structural Formula



- H. Substance Group Not applicable
- I. Substance Remark None
- J. Molecular Weight 223.20
- 1.02 OECD INFORMATION
- A. Sponsor Country: Japan
- B. Lead Organization:

Name of Lead Organization:

	Ministry of Health and Welfare (MHW)
	Ministry of International Trade and Industry (MITI)
	Environment Agency (EA)
	Ministry of Labor (MOL)
Contact person:	Mr. Yasuhisa Kawamura
	Director
	Second International Organization Bureau
	Ministry of Foreign Affairs
Address:	2-2-1 Kasumigaseki, Chiyoda-ku
	Tokyo 100, Japan
	TEL 81-3-3581-0018
	FAX 81-3-3503-3136

#### C. Name of responder

Name:	Same as above contact person
Address:	

OECD SIDS 1. GENERAL INFORMATION

#### 1.1 GENERAL SUBSTANCE INFORMATION

A. **Type of Substance** element []; inorganic []; natural substance []; organic [X]; organometallic []; petroleum product [] B. **Physical State** gaseous []; liquid []; solid [X] C. > 97 % Purity 1.2 **SYNONYMS** 1-Aminoanthraquinone **IMPURITIES** Anthraquinone 1.3 **ADDITIVES** 1.4 None **QUANTITY** Location 1.5 Production (tonnes) Date 1,000-2,000/year 1990-1993 Japan Reference: MITI, Japan (1994a) 1.6 LABELLING AND CLASSIFICATION None 1.7 **USE PATTERN** General Type of Use: **Category:** A. (1) Industry use Intermediate for dyestuffs (2) Industry use Intermediate for dyes and pharmaceuticals Reference: (1) MITI, Japan (1994a) (2) ECDIN Database **Uses in Consumer Products** B. None 1.8 **OCCUPATIONAL EXPOSURE LIMIT VALUE** None

#### **1.9 SOURCES OF EXPOSURE**

Source:

Reference:

Media of release: Water from a production site Quantities per media: 1.9 tonnes/year MITI, Japan (1994a)

#### 1.10 ADDITIONAL REMARKS

А.	Options for disposal	Incineration
	Reference:	MITI, Japan (1994a)

#### B. Other remarks None

#### 2.1 MELTING POINT

(a) Value: Decomposition: Sublimation: Method: GLP: Reference:	256 - 258 °C Yes [] No [ <b>X</b> ] Ambiguous [] Yes [] No [ <b>X</b> ] Ambiguous [] Yes [] No [] ? [ <b>X</b> ] MITI (1992)
(b) Value: Decomposition: Sublimation: Method: GLP: Reference:	260 °C Yes [ ] No [ <b>X</b> ] Ambiguous [ ] Yes [ ] No [ <b>X</b> ] Ambiguous [ ] Yes [ ] No [ ] ? [ <b>X</b> ] Bayer AG
(c) Value: Decomposition: Sublimation: Method: GLP: Reference: BOILING POINT	251 - 252 °C Yes [] No [ <b>X</b> ] Ambiguous [] Yes [] No [ <b>X</b> ] Ambiguous [] Yes [] No [] ? [ <b>X</b> ] Shibusawa et al. (1977)
(a) Value: Pressure: Decomposition: Method: GLP: Reference:	<pre>&gt; 300 °C Yes [] No [] Ambiguous [] Yes [X] No [] ? [] MITI, Japan (1994b)</pre>
(b) Value: Pressure: Decomposition: Method: GLP: Remarks: Reference:	>300 °C Yes [ ] No [ <b>X</b> ] Ambiguous [ ] Unknown Yes [ ] No [ ] ? [ <b>X</b> ] None Bayer AG

## 2.3 DENSITY (Relative density)

No data available

#### 2.4 VAPOUR PRESSURE

Value:	
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2.2

1.2 x 10<sup>-4</sup> Pa

Temperature:	100°C
Method:	calculated []; measured [X]

OECD Test Guideline 104 Dynamic methodGLP:Yes [X] No [] ? []Reference:MITI, Japan (1994b)

#### 2.5 PARTITION COEFFICIENT log<sub>10</sub>P<sub>ow</sub>

(a)	
Log Pow:	3.74
Temperature:	25 °C
Method:	calculated []; measured [X]
	OECD Test Guideline 107
GLP:	Yes <b>[X]</b> No []? []
Reference:	MITI (1992)
(b)	
Log Pow:	2.1
Temperature:	25 °C
Method:	calculated [X]; measured []
	Leo and Hansch method
GLP:	Yes [] No [X] ? []
Reference:	Bayer AG (1991)

#### 2.6 WATER SOLUBILITY

#### A. Solubility

(a)	
Value:	32 mg/l
Temperature:	25 °C
Description:	Miscible []; Of very high solubility []; Of high solubility []; Soluble []; Slightly soluble []; Of low solubility []; Of very low solubility <b>[X]</b> ; Not soluble []
Method:	OECD Test Guideline 105
GLP:	Yes [X] No [] ? []
Reference:	MITI (1992)
(b)	
Value:	20 mg/l
Temperature:	20 °C
Description:	Miscible []; Of very high solubility []; Of high solubility []; Soluble []; Slightly soluble []; Of low solubility []; Of very low solubility <b>[X</b> ]; Not soluble []
Method:	Unknown
GLP:	Yes [] No [] ? [X]
Reference:	Bayer AG

#### B. pH Value, pKa Value

No data available

#### 2.7 FLASH POINT

No data available

### 2.8 AUTO FLAMMABILITY

No data available

#### 2.9 FLAMMABILITY

No data available

#### 2.10 EXPLOSIVE PROPERTIES

No data available

#### 2.11 OXIDIZING PROPERTIES

No data available

#### 2.12 OXIDATION: REDUCTION POTENTIAL

No data available

#### 2.13 ADDITIONAL DATA

A. Partition co-efficient between soil/sediment and water (Kd)

No data available

B. Other data

None

#### 3.1 STABILITY

#### 3.1.1 PHOTODEGRADATION

Type:	Air []; Water [X]; Soi	l; Other []
Light source:	Sunlight [X]; Xenon la	amp [ ]; Other [ ]
Spectrum of		
substance:	epsilon = $5.46 \times 10^3$ at	300 nm
	epsilon = $6.89 \times 10^3$ at	470 nm
Estimated parameter for calculat	tion:	
	Quantum yield 0.001	
	Concentration	5 x 10 <sup>-5</sup> M
	Depth of water body	500 cm
	Conversion constant	$6.023 \times 10^{20}$
Result:	Degradation rate	7.62 x 10 <sup>-11</sup> mol/l/s
	Half life	$1.44 \times 10^{-2}$ years
Reference:	W. J. Lyman, W. F. Re	eehl and D. H. Rosenblatt (1981)

#### 3.1.2 STABILITY IN WATER

Type:	Abiotic (hydrolysis) <b>[X]</b> ; biotic (sediment) <b>[</b> ]
Result:	Stable at pH 4, 7 and 9 at 25°C
Method:	OECD Test guideline 111
GLP:	Yes <b>[X]</b> No []? []
Test substance:	1-Aminoanthraquinone
Reference:	MITI, Japan (1994b)

#### 3.1.3 STABILITY IN SOIL

No data available

#### **3.2 MONITORING DATA (ENVIRONMENT)**

(a)	
Type of Measurement:	Background [], At contaminated Site []; Other [X]
Media:	Surface water
Results:	ND (Detection limits: 0.0002 ug/ml) in 9 areas in Japan
Remarks:	None
Reference:	EA, Japan (1987)
(b)	
Type of Measurement:	Background [], At contaminated Site [], Other [X]
Media:	Sediment

Media:	Sediment
Results:	0.022 ug/g dry (Number of detections/Number of
	samples: 1/21 in 7 areas, Detection limits: 0.02 ug/g
	dry) in Japan
Remarks:	None
Reference:	EA, Japan (1987)

# **3.3 TRANSPORT AND DISTRIBUTION BETWEEN ENVIRONMENTAL COMPARTMENTS INCLUDING ESTIMATED ENVIRONMENTAL**

#### CONCENTRATIONS AND DISTRIBUTION PATHWAYS

### 3.3.1 TRANSPORT

No data available

#### **3.3.2** THEORETICAL DISTRIBUTION (FUGACITY CALCULATION)

The potential environmental distribution of 1-Aminoanthraquinone obtained from a generic level III fugacity model is shown in Table. The results show that if 1-aminoanthraquinone is released mainly to soil or air, it is likely to distribute into soil compartment. But, if 1-Aminoanthraquinone is released mainly to water, it is likely to be transported both to soil and sediment. Due to the low vapour pressure of 1-aminoanthraquinone, it is unlikely to distribute into air.

Environmental distribution 1-Aminoanthraquinone using a generic level III fugacity model.

Compartment	Release: 100% to air	Release: 100% to water	Release: 100% to soil
Air	0.18%	0.04%	0.00%
Water	0.60%	62.57%	0.47%
Soil	99.06%	21.34%	99.41%
Sediment	0.15%	16.06%	0.12%

Reference: EA and MITI, Japan (1994)

## 3.4 IDENTIFICATION OF MAIN MODE OF DEGRADABILITY IN ACTUAL USE

No data available

#### **3.5 BIODEGRADATION**

(-)

(a)	
Туре:	aerobic [X]; anaerobic []
Inoculum:	adapted []; non-adapted [X];
Concentration of the chemical	: 100 mg/l related to Test Substance [X]
Medium:	<pre>water[];water-sediment[];soil[];sewage treatment[]</pre>
	other [Japanese standard activated sludge]
Degradation:	Degree of degradation after 28 days
-	0, 0 and 1 % from BOD
	3, 1 and 2 % from HPLC analysis
Results:	Readily biodeg. []; Inherently biodeg. []; under test
	condition no biodegradation observed [X]
Method:	OECD Test Guideline 301 C
GLP:	Yes <b>[X]</b> No <b>[</b> ] ? <b>[</b> ]
Test substance:	1-Aminoanthraquinone
Reference:	MITI, Japan (1992)
(b)	
Туре:	aerobic []; anaerobic [X]
Inoculum:	adapted []; non-adapted [];

# Concentration of the chemical: related to Test Substance []

	L J
Medium:	<pre>water[];water-sediment[];soil [];sewage treatment []</pre>
	other []
Degradation:	Degree of degradation after 20 days
	0 %
Results:	Readily biodeg. []; Inherently biodeg. []; under test
	condition no biodegradation observed [X]
Method:	OECD Test Guideline 301 D (Closed bottle Test)
GLP:	Yes [] No [] ? [X]
Test substance:	1-Aminoanthraquinone
Reference:	Bayer AG

## 3.6 BOD<sub>5</sub>,COD OR RATIO BOD<sub>5</sub>/COD

## Not applicable

## **3.7 BIOACCUMULATION**

Carp
8 weeks
25 °C
(1) 30 mg/l
(2) 3 mg/l
(1) 50 - 150
(2) 55 - 137
OECD Test Guideline 305 C
calculated []; measured [X] static []; semi-static [];
flow-through []; other []
Yes [X] No [] ? []
1-Aminoanthraquinone
MITI, Japan (1992)

## 3.8 ADDITIONAL REMARKS

A.	Sewage treatment	None
	8	

## **B. Other information** None

## 4.1 ACUTE/PROLONGED TOXICITY TO FISH

(a)	
Type of test:	<pre>static []; semi-static [X]; flow-through []; other [] open-system [X]; closed-system []</pre>
Species:	Oryzias latipes
Exposure period:	96 hr
Results:	$LC_{50}$ (24h) = > 1000 mg/l
	$LC_{50} (48h) = > 1000 \text{ mg/l}$
	$LC_{50}(72h) = > 1000 \text{ mg/l}$
	$LC_{50}(96h) = > 1000 \text{ mg/l}$
	NOEC =
	LOEC =
Analytical monitoring:	Yes [] No [X] ? []
Method:	OECD Test Guideline 203 (1981)
GLP:	Yes [] No [X] ? []
Test substance	1-Aminoanthraquinone purity = 98.8 %
Remarks:	A group of 10 fish were exposed to each of 5
	nominal concentrations (95-1000 mg/l). Stock solution
	was prepared with DMSO(1000 mg/l) Controls with
	and without this vehicle were taken for test
Reference:	EA, Japan (1994)
(b)	
Type of test:	static []; semi-static []; flow-through []; other []
	open-system []; closed-system []
Species:	Leuciscus idus (Goldorfe)
Exposure period:	96 hr
Results:	$LC_0 (48h) = > 1000 \text{ mg/l}$
Analytical monitoring:	Yes [] No [] ? [X]
Method:	Other method
	Bestimmung der akuten Wirkung von Stoffen auf Fische
	Arbeitskreis "Fischtest" im Hauptausschuss
	"Detergenten" (15.10.1973)
Method:	Unknown
GLP:	Yes [] No [] ? [X]
Test substance:	1-Aminoanthraquinone
Remarks:	None
Reference:	Bayer AG

# 4.2 ACUTE TOXICITY TO AQUATIC INVERTEBRATES

## A. Daphnia

Type of test:	<pre>static [X]; semi-static []; flow-through []; other []; open-system [X]; closed-system []</pre>
Species:	Daphnia magna
Exposure period:	24 hr
Results:	$EC_{50} (24h) = > 1000 \text{ mg/l}$
	$EC_{50} (48h) = > 1000 \text{ mg/l}$
	NOEC =

## UNEP PUBLICATIONS

	LOEC =
Analytical monitoring:	Yes [] No [X] ? []
Method:	OECD Test Guideline 202 (1984)
GLP:	Yes [] No [X] ? []
Test substance:	1-Aminoanthraquinone, purity: = 98.8 %
Remarks:	20 daphnids (4 replicates; 5 organisms per replicate) were
	exposed to each of 5 nominal concentrations (100-1000
	mg/l). Stock solution was prepared with DMSO:HCO
	= $9:1(100-1000 \text{ mg/l})$ . Controls with and without this
	vehicle were taken for test.
Reference:	EA, Japan (1994)

#### B. Other aquatic organisms

#### No studies located

## 4.3 TOXICITY TO AQUATIC PLANTS e.g. Algae

Species:	Selenastrum capricornutum ATCC 22662
End-point:	Biomass [X]; Growth rate []; Other []
Exposure period:	72 hours
Results:	Biomass: $EC_{50} (24h) =$
	$EC_{50}$ (72h) = 0.25 mg/l
	NOEC = $0.10 \text{ mg/l} (p < 0.05)$
	LOEC =
Analytical monitoring:	Yes [] No [X] ? []
Method:	open-system [X]; closed-system []
	OECD Test Guideline 201 (1984)
GLP:	Yes [] No [X] ? []
Test substance:	1-Aminoanthraquinone, purity = $98.8 \%$
Remarks:	The $EC_{50}$ values for biomass were calculated based on
	8 nominal concentrations (0.058-3.2 mg/l). Stock solution
	was prepared with DMSO (100 mg/l). Controls with and
	without this vehicle were taken for the test.
Reference:	EA, Japan (1994)

#### 4.4 TOXICITY TO BACTERIA

No studies located

## 4.5 CHRONIC TOXICITY TO AQUATIC ORGANISMS

## 4.5.1. CHRONIC TOXICITY TO FISH

No studies located

## 4.5.2. CHRONIC TOXICITY TO AQUATIC INVERTEBRATES

(a)	
Type of test:	<pre>static []; semi-static [X]; flow-through []; other [];</pre>
	open-system [X]; closed-system []
Species:	Daphnia magna

**UNEP PUBLICATIONS** 

End-point:	Mortality []; Reproduction rate [X]; Other [X]
Exposure period:	21 day
Results:	
Immobility:	$EC_{50} (48 h) = > 1000 mg/l$
	$EC_{50} (21 \text{ d}) = 0.62 \text{mg/l} (95\% \text{ confidence limits:} 0.49-0.74 \text{ mg/l})$
	NOEC =
	LOEC =
Reproduction:	$EC_{50}$ (21 d) = 0.56mg/l (95% confidence limits:
	0.51-0.62 mg/l)
	NOEC = $0.32 \text{mg/l} (p < 0.05)$
	LOEC = 0.56 mg/l (p < 0.05)
Analytical monitoring:	Yes [] No [X] ? []
Method:	OECD Test Guideline 202 (1984)
GLP:	Yes [] No [X] ? []
Test substance:	1-Aminoanthraquinone, purity $= 98.8\%$
Remarks:	40 daphnids (4 replicates; 10 organisms per replicate)
	were exposed to each of 5 nominal concentrations (100-
	1000 mg/l) or (5.6-56 mg/l) or (0.32-3.2 mg/l). Stock
	solution was prepared with DMSO:HCO-40=9:1(100- $(0, 22, 22, 2, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3,$
	1000 mg/l) or (5.6-56 mg/l) or (0.32-3.2 mg/l).
D.C.	Controls with and without this vehicle were taken for test.
Kelerence:	EA, Japan (1994)
	End-point: Exposure period: Results: Immobility: Reproduction: Analytical monitoring: Method: GLP: Test substance: Remarks: Reference:

#### 4.6 TOXICITY TO TERRESTRIAL ORGANISMS

### 4.6.1 TOXICITY TO SOIL DWELLING ORGANISMS

No data available

#### 4.6.2 TOXICITY TO TERRESTRIAL PLANTS

No data available

# 4.6.3 TOXICITY TO OTHER NON MAMMALIAN TERRESTRIAL SPECIES (INCLUDING AVIAN)

No data available

#### 4.7 BIOLOGICAL EFFECTS MONITORING (INCLUDING BIOMAGNIFICATION)

No studies located

## 4.8 BIOTRANSFORMATION AND KINETICS IN ENVIRONMENTAL SPECIES

No data available

4.9 ADDITIONAL REMARKS

None

## 5.1 ACUTE TOXICITY

#### 5.1.1 ACUTE ORAL TOXICITY

(a)	
Type :	$LD_0$ []; $LD_{100}$ []; $LD_{50}$ [ <b>X</b> ]; $LDL_0$ []; Other []
Species/strain:	Rat
Value :	> 5000 (mg/kg)
Method:	Unknown
GLP:	Yes [] No [] ? <b>[X</b> ]
Test substance:	1-Aminoanthraquinone, purity: Unknown
Remarks:	None
Reference:	Loeser E. (1978)

(b)	
Type :	$LD_0$ []; $LD_{100}$ []; $LD_{50}$ [ <b>X</b> ]; $LDL_0$ []; Other []
Species/strain:	Rat
Value :	> 1600 (mg/kg)
Method:	Unknown
GLP:	Yes [] No [] ? <b>[X</b> ]
Test substance:	1-Aminoanthraquinone, purity: unknown
Remarks:	None
Reference:	Marhold J. (1972)

## 5.1.2 ACUTE INHALATION TOXICITY

No data available

### 5.1.3 ACUTE DERMAL TOXICITY

Type :	$LD_0$ []; $LD_{100}$ []; $LD_{50}$ [ <b>X</b> ]; $LDL_0$ []; Other []
Species/strain:	Mice (ddN strain)
Value:	> 2000 (mg/kg b.w.)
Method:	Fixed dose test
	10 animals/dose, 14 days observation period,
GLP:	Yes [] No [X] ? []
Test substance:	1-Aminoanthraquinone, purity: unknown
Remarks:	No compound related clinical signs were observed
Reference:	Unpublished company data

## 5.1.4 ACUTE TOXICITY, OTHER ROUTES OF ADMINISTRATION

No data available

## 5.2 CORROSIVENESS/IRRITATION

### 5.2.1 SKIN IRRITATION/CORROSION

No data available

## OECD SIDS 5. TOXICITY

# 5.2.2 EYE IRRITATION/CORROSION

Test species/strain:	Rabbit
Test method:	Standard Draize test
GLP:	YES [] NO [] ? [X]
Test result:	500 mg/24h, "Mild" effect
Test substance:	1-Aminoanthraquinone
Remarks:	-
Reference:	Marhold J. (1986)

## 5.3 SKIN SENSITIZATION

No data available

#### 5.4 REPEATED DOSE TOXICITY

Species/strain: Sex: Route of Administration:	Rat (Crj:CD(SD)) Female []; Male []; Male/Female <b>[X]</b> ; No data [] Oral gayage
Exposure period:	Males: 42 days including 14 days before mating Females: from 14 days before mating to day 3 of lactation
Frequency of treatment: Post exposure observation per	7 days/week
Dose:	0.40.200 or 1000 mg/kg (13 animals /group)
Control group	Yes $[\mathbf{X}]$ : No [1: No data []:
condor group.	Concurrent no treatment []: Concurrent vehicle [X]:
	Historical []
NOFI ·	< 40  mg/kg/day
LOFL:	40 mg/kg/day
Results:	Increased spleen weights were observed in males in the 200
Results.	$m\sigma/k\sigma$ group and above as well as in females in the 1000
	mg/kg group and doore us wen us in remates in the root
	than 200 mg/kg male groups. In hemato-morphological
	examination erythrocyte count hemoglobin and mean
	corpuscular hemoglobin were decreased in more than 200
	$m\sigma/k\sigma$ male groups. In clinical chemistry the potassium
	concentration in 1000 mg/kg male group and the chlorine
	concentration in 200 mg/kg male group, and the emotion in concentration in 200 mg/kg male group were decreased In
	historiation in 200 mg/kg mate group were decreased. In
	droplet and agging philic hady in hidray wars increased in more
	then 40 mg/lsg m
	than 40 mg/kg male groups. Nephropathy and dark coloration
	of the spieen were observed in 40 mg/kg both male and remaie
	groups. Extramedullary nematopolesis in spleen were observed
	in males in the 40 mg/kg group and above.
Method:	OECD Combined Repeat dose and reproductive/
CLD	Developmental Screening Toxicity Test (1992)
GLP:	Yes [X] No [] ? []
Test substance:	Commercial, purity: 98.7 %
Reference:	MHW, Japan (1994a)

# 5.5 GENETIC TOXICITY IN VITRO

## A. BACTERIAL TEST

(a) Type : System of testing:	Bacterial reverse mutation assay	
Species/strain:	<i>S. typhimurium</i> TA 98, TA 100, TA 1 E. coli WP2 uvrA	1535, TA1537
Concentration: Metabolic activation: Results:	0, 312.5, 625, 1250, 2500, 5000 μg/p. With []; Without []; With and Witho	late out <b>[X]</b> ;No data[ ]
Cytotoxicity conc:	With metabolic activation: $5000 \ \mu g/p$ Without metabolic activation: $5000 \ \mu$	late g/plate
Precipitation conc: Genotoxic effects:		
	S. typhimurium TA 100, TA1535, TA	A98 + 2 -
	With metabolic activation: Without metabolic activation:	[] [] [X] [] [] [X]
	<i>S. typhimurium</i> TA 1537 With metabolic activation: Without metabolic activation:	+ ? - [X] [] [] [] [] [X]
	<i>E. coli</i> WP2 uvrA With metabolic activation: Without metabolic activation:	+ ? - [][][X] [][][X]
Method: GLP: Teat substance:	Japanese Guideline for Screening Mu chemicals Yes <b>[X]</b> No <b>[]</b> ? <b>[]</b> Commercial, purity: 98.7%	tagenicity testing of
Remarks:	Procedure: Plate incorporation method Plates/test: 3 Activation system: Liver S-9 fraction from Phenobarbital and 5,6-Benzoflavone pretreated male SD rats with NADPH-generating system Media:Histidine selective No. replicates: 2	
Reference:	MHW, Japan (1994b)	

#### **B.** NON-BACTERIAL IN VITRO TEST

Type :	Cytogenetics Assay
System of testing:	Chinese hamster lung (CHL/IU) cells
Species/strain:	-S9 (continuous treatment) 0, 0.3, 0.7, 1.3 mg/ml
Concentration:	-S9 (short-term treatment) 0, 0.6, 1.1, 2.2 mg/ml

	+S9 (short-term treatment) 0, 0.6, 1.1, 2.2 mg/ml
Metabolic activation:	With []; Without []; With and Without [X]; No data []
Results:	
	Cytotoxicity conc: With metabolic activation: 2.2 mg/ml Without metabolic activation: 2.2 mg/ml
Precipitation conc:	· ·
Genotoxic effects:	+ ? -
	With metabolic activation: [] [] [X]
	Without metabolic activation: [] [] [X]
Method:	Japanese Guideline for Screening Mutagenicity testing
	of chemicals
GLP:	Yes [X] No []? []
Test substance:	Commercial, purity 98.7 %
Remarks:	Plates/test:2
	Activation system: S-9 fraction from the liver of
	Phenobarbital and 5,6-Benzoflavone induced male SD
	derived rats with NADPH-generating system
	Media: RPMI 1640 medium plus 10% foetal calf serum
	plus phytohaemagglutinin
	No. replicates: 1
Reference:	MHW, Japan (1994b)

## 5.6 GENETIC TOXICITY IN VIVO

Test type:	Micronucleus Test			
Test species/strain:	Mice			
Test method:	i.p. once, 5000 mg/k	g b.w.		
GLP:	Yes [], No [], ? [X]	-		
Test Results:	No indications of a c	lastoge	enic e	effect
Genotoxic effects:		•		
		+	?	-
	Micronucleus test	[]	[]	[X ]
Remarks:	No further information	on are	prov	ided

Remarks:	No further information are provided
Reference:	Bayer AG

## 5.7 CARCINOGENICITY

Species/strain: Method:	Rats once a week, orally, 10 mg/0.5 ml corn oil/rat, 14 months, 20 males and 20 females
GLP:	YES [] NO [X]
Result:	female: 6 adenomas of the mammary gland and other benign tumors (no further information) male: one cell sarcoma of the intestine and neurofibrosarcoma
Test substance:	1-Aminoanthraquinone
Remarks:	Only meeting abstracts (15 lines)
Reference:	Laham S. et al. (1966)

## 5.8 TOXICITY TO REPRODUCTION

Type:	Fertility []; One generation study []; Two generation study []; Other [X]
Species/strain:	Rat Crj:CD(SD)
Sex:	Female []; Male []; Male/Female [X]; No data []
Route of Administration:	Oral, gavage
Exposure period:	Males: 42 days including 14 days before mating Females: from 14 days before mating to day 3 of lactation.
Frequency of treatment:	7 days/week
Postexposure observation per	iod:
Premating exposure period: Duration of the test;	male: 14 days, female: 14 days
Doses:	0, 40, 200, or 1000 mg/kg (10 animals/sex/group)
Control group:	Yes [X]; No []; No data [];
	Concurrent no treatment []; Concurrent vehicle [X];
	Historical []
NOEL Parental :	< 40  mg/kg/day
NOEL F1 Offspring:	< 40 mg/kg/day
NOEL F2 Offspring:	N/A
Results:	The parental animals exhibited no effects on reproductive parameters including copulation index, fertility index, gestation length, number of corpora lutea or implantation, implantation index, gestation index, delivery index, parturition or maternal behavior. However, nursing behavior disappeared in all of the treatment female groups. Viability of pups on day 4 after birth was decreased in all treatment groups. No external or skeletal anomalies related to the test substance administration were detected in any of the offspring. Furthermore, there are no significant differences in the number of offspring or live offspring, sex ratio, live birth index or body weights.
Method: GLP: Test substance: Remarks:	OECD Combined Repeat dose and reproductive/ developmental Screening Toxicity Test (1992) Yes <b>[X]</b> No <b>[]</b> ? <b>[]</b> Purity 98.7 %
Reference:	MHW, Japan (1994b)

## 5.9 DEVELOPMENTAL TOXICITY/ TERATOGENICITY

See 5.8

## 5.10 OTHER RELEVANT INFORMATION

## A. Specific toxicities

No data available

## B. Toxicodynamics, toxicokinetics

No data available

# 5.11 EXPERIENCE WITH HUMAN EXPOSURE

None

EA, Japan (1987) Environment Monitoring of Chemicals - Environmental Survey Report of F.Y. 1986 and 1987 (Office of Health Studies, Environmental Health Department, EA, Japan)

EA, Japan (1994) "Investigation of the Ecotoxicological Effects of OECD High Production Volume Chemicals", Office of Health Studies, Environmental Health Department, Environment Agency, Japan (HPV/SIDS Test conducted by EA, Japan)

EA & MITI, Japan (1994) Unpublished Report on Exposure Estimation (HPV/SIDS Test conducted by EA and MITI, Japan)

ECDIN database (1994)

Laham S. et al., Toxicol. Appl. Pharmacol. 8, 346 (1966)

Loeser E., Bayer AG data, short report, 11. 8. 1978

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