7	CAS No.: 91-22-5	Substance: Quinoline						
Chemical	Chemical Substances Control Law Reference No.: 5-794							
PRTR Law	v Cabinet Order No.: 1-81							
Molecular	Formula: C ₉ H ₇ N	Structural Formula:						
Molecular	Weight: 129.16							

1. General information

The aqueous solubility of this substance is 6.33×10^3 mg/1000 g (20°C), the partition coefficient (1-octanol/water) (log K_{ow}) is 2.03, and the vapor pressure is 0.083 mmHg (=11 Pa) (25°C). Biodegradability (aerobic degradation) is characterized by a BOD degradation rate of 0.2%, and bioaccumulation is judged to be non-existent or low. Furthermore, the substance does not have any hydrolyzable groups.

This substance is designated as a Class 1 Designated Chemical Substance under the Law Concerning Reporting, etc. of Releases to the Environment of Specific Chemical Substances and Promoting Improvements in Their Management (PRTR Law). The main use of this substance is as a raw material for 8-hydroxyquinoline. Whereas 70–80% of the 8-hydroxyquinoline thus produced is estimated to be used as a raw material for an agricultural chemical (the bactericide copper 8-hydroxyquinoline), 5–10% is estimated to be used as a raw material for pharmaceuticals, analytical reagents, or chelating agents for removing metals. The production quantity in fiscal 2010 was approximately 900 t (estimated). The production and import category under the PRTR Law is more than 100 t.

2. Exposure assessment

Total release to the environment in fiscal 2010 under the PRTR Law was approximately 0.13 t, and all releases were reported. All reported releases were to the atmosphere. In addition, approximately 12 t was transferred to waste materials, and 0.024 t was transferred to sewage. The main source of reported releases was the chemical industry. A multi-media model used to predict the proportions distributed to individual media in the environment indicated that in regions where the largest quantities were estimated to have been released to the overall environment or to the atmosphere in particular, the predicted proportion distributed to the soil was 88.9%.

The maximum expected concentration of exposure to humans via inhalation, based on general environmental atmospheric data, was around $0.0069 \,\mu g/m^3$. The mean annual value for atmospheric concentration in fiscal 2010 was calculated by using a plume-puff model on the basis of reported releases to the atmosphere according to the PRTR Law; this model predicted a maximum level of $0.021 \,\mu g/m^3$. The maximum expected oral exposure was estimated to be around $0.00032 \,\mu g/kg/day$ on the basis of calculations from data for public freshwater bodies.

The predicted environmental concentration (PEC), which indicates exposure to aquatic organisms, was reported to be around $0.0081 \,\mu g/L$ for public freshwater bodies and $0.0067 \,\mu g/L$ for seawater.

3.Initial assessment of health risk

This substance may cause irritation to eyes and skin. Inhalation exposure to the substance may cause coughing and sore throat, while oral exposure may cause sore throat. Redness of skin and the redness and pain of eyes may occur if they accidentally come into direct contact with the substance.

With regard to the substance's non-carcinogenic health risk, as some studies reported that its repeated administration to rats had caused tumor in all of the low dose groups, it would not be logical to assume the lowest dose as its LOAEL for assessment of its non-carcinogenic health risk. Therefore, the 'non-toxic level*' of

the substance could not be identified on the basis of its non-carcinogenic effects.

Although sufficient information was not available to evaluate the carcinogenic potential of the substance to human, most of the information suggests its possible carcinogenic effects. Therefore, assessment was conducted with assumption that there would be no threshold for its carcinogenicity, although it was still uncertain whether the substance was carcinogenic to human.

As for oral exposure to the substance, with assumption that there would be no threshold for its carcinogenicity, a slope factor of 3 $(mg/kg/day)^{-1}$ (for hemangioendothelioma or hemangiosarcoma) obtained from experiments on rats was used. As for inhalation exposure to the substance, its unit risk could not be identified.

As for oral exposure to the substance, its maximum exposure was estimated to be approximately 0.00032 μ g/kg/day when its intakes through freshwater from public water bodies were assumed. For its carcinogenic potential, the excess cancer incidence rate for the predicted maximum exposure was calculated to be 9.6×10^{-7} with the above slope factor. In addition, when its intakes through fish and freshwater from public water bodies were assumed, its oral exposure would be below 0.0061 μ g/kg/day (for 60 to 69 years of age) and below 0.0044 μ g/kg/day (for all age groups). The excess incidence rates for exposures below 0.0061 μ g/kg/day and below 0.0044 μ g/kg/day would be below 1.8×10^{-5} and below 1.3×10^{-5} , respectively, with the above slope factor. Therefore, collection of further information would be required to assess health risk from its oral exposure.

With regard to inhalation exposure to the substance, its health risk could not be assessed as neither its 'non-toxic level*' nor its unit risk could be identified. If 100 % absorption were assumed, the maximum exposure to the substance would be $0.0021 \,\mu g/kg/day$. This ingestion rate would provide an excess incidence rate of 6.2×10^{-6} with the above slope factor. Therefore, collection of further information on inhalation exposure to the substance would be required to assess health risk from its inhalation.

	Toxicity					Exposure assessment					
Exposure Path			Animal	Criteria for diagnoses (endpoint)	Exposure Predicted maximum exposure dose and concentration		e dose and	Result of risk assessment			Judgment
Oral	'Non-toxic - level*'	mg/kg/day	-	Suppressed body weight increase	Drinking water	-	µg/kg/day	MOE Excess incidence rate	-	×	()
	Slope factor 3	(mg/kg/day) ⁻¹	Rat	Hemangioendothelioma or hemangiosarcoma	Freshwater	0.00032	µg/kg/day	MOE Excess incidence rate	- 9.6 × 10 ⁻⁷	×	
	'Non-toxic - level*'	mg/m ³	-	-	Ambient air	0.0069	$\mu g/m^3$	MOE Excess incidence rate	-	×	()
Inhalation	Unit risk -	$(\mu g/m^3)^{-1}$	-	-	Indoor air	-	$\mu g/m^3$	MOE Excess incidence rate	-	×	×

Non-toxic level *

- When a LOAEL is available, it is divided by 10 to obtain a NOAEL-equivalent level.
- When an adverse effect level for the short-term exposure is available, it is divided by 10 to obtain a level equivalent to an adverse effect level for the long-term exposure.

4.Initial assessment of ecological risk

With regard to acute toxicity, the following reliable data were obtained: a 72-h EC_{50} of 65,900 µg/L for growth inhibition in the green alga *Pseudokirchneriella subcapitata*, a 48-h EC_{50} of 25,000 µg/L for swimming inhibition in the crustacean *Daphnia magna*, a 96-h LC_{50} of 440 µg/L for the fish species *Pimephales promelas* (fathead minnow), and a 96-h LC_{50} of 4,897 µg/L for the midge *Chironomus riparius* species. Accordingly, based

on these acute toxicity values and an assessment factor of 100, a predicted no effect concentration (PNEC) of 4.4 μ g/L was obtained.

With regard to chronic toxicity, the following reliable data were obtained: a 72-h NOEC of 4,800 μ g/L for growth inhibition in the green alga *P. subcapitata*, and a 21-d NOEC of 800 μ g/L for reproductive inhibition in the crustacean *D. magna*. Accordingly, based on these chronic toxicity values and an assessment factor of 100, a predicted no effect concentration (PNEC) of 8 μ g/L was obtained.

The value of 4.4 μ g/L obtained from the acute toxicity to the fish species was used as the PNEC for this substance.

The PEC/PNEC ratio was 0.002 for freshwater bodies and seawater. Accordingly, further work is considered unnecessary at this time.

	Hazard assessment (basis for PNEC)					Exposure assessment			Judgment	
	Species	Acute/ chronic	Endpoint	Assessment factor	Predicted no effect concentration PNEC (µg/L)	Water body	Predicted environmental concentration PEC (µg/L)	PEC/PNEC ratio	based on PEC/PNEC ratio	Assessment result
	Fish	Arresta	LC ₅₀	100		Freshwater	0.0081	0.002		
	(fathead minnow)	Acute	mortality	100	4.4	Seawater	0.0067	0.002		

5. Conclusions

	Conclusions								
	Oral exposure	Collection of further information would be required.							
Health risk	Inhalation	Although risk to human health could not be identified, collection		_					
	exposure	of further information would not be required.	()					
Ecological risk	No need of further work at present.								
[Risk judgmen	[Risk judgments] : No need for further work A: Requiring information collection								
Candidates for further work ×: Impossibility of risk characterization									
(): Though a risk characterization cannot be determined, there would be little necessity									
of collecting information.									
(): Further information collection would be required for risk characterization.									