

river channel structure database, estimating the concentration in rivers by taking into consideration only dilution gave a maximum value of 0.00014 µg/L (however, this excludes releases from factories that considered to calculate releases based on minimum determination limits).

3. Initial assessment of health risk

This substance causes effects on the liver, kidneys and central nervous system, and may result in unconsciousness. Inhalation exposure to the substance causes dizziness, drowsiness, headache, nausea and vomiting. Oral exposure causes abdominal pain and diarrhea in addition to the same symptoms as inhalation. The substance is irritating to the eyes, and causes redness and pain. Contact with the skin causes redness and pain. The substance on the skin may be absorbed to cause dizziness and drowsiness.

As sufficient information on the carcinogenicity of the substance was not available, the initial assessment was conducted on the basis of information on the non-carcinogenic effects. However, the carcinogenicity was taken into consideration for this risk assessment, because there is sufficient evidence in experimental animals for the carcinogenicity of this substance.

The lowest reliable NOAEL based on the non-carcinogenic effects of 5 ppm was determined from toxicity tests in rats and mice. However, it would be appropriate to use the LOAEL of 5 ppm based on the carcinogenic effects for the conservative assessment, considering the significantly increased incidence of hepatocellular adenomas in female rats exposed to 5 ppm of the substance. The LOAEL of 5 ppm for inhalation exposure was adjusted according to exposure conditions to obtain 0.89 ppm (5.6 mg/m³) and subsequently divided by a factor of 10 to account for uncertainty in using a LOAEL. The calculated value of 0.56 mg/m³ was identified as the ‘non-toxic level*’ of the substance for inhalation exposure.

With regard to inhalation exposure, the predicted maximum exposure concentration in ambient air was 0.85 µg/m³, approximately. The MOE (Margin of Exposure) would be 13, when calculated from the predicted maximum exposure concentration and the ‘non-toxic level*’ of 0.56 mg/m³, and subsequently divided by a factor of 10 to account for extrapolation from animals to humans and by another factor of 5 to take into consideration the carcinogenicity in animals. In addition, the maximum concentration (annual mean) in ambient air near the operators releasing large amount of the substance was estimated to be 1.1 µg/m³ approximately, based on the releases reported in FY 2014 under the PRTR Law. The MOE would be 10, when calculated from this concentration. In indoor air, the maximum concentration reported in 2004 was 1.9 µg/m³. The MOE would be 6, when calculated from this concentration. Therefore, collection of information on the health risk of this substance via inhalation in ambient air would be required, and collection of further information would also be required to assess the health risk via inhalation in indoor air.

| Exposure Path | Toxicity | | | Exposure assessment | | Result of risk assessment | | | Judgment |
|---------------|---|--------|-----------------------------------|--------------------------|---|---------------------------|-----|-----|----------|
| | Criteria for risk assessment | Animal | Criteria for diagnoses (endpoint) | Exposure medium | Predicted maximum exposure dose and concentration | | | | |
| Oral | ‘Non-toxic level*’ (—) mg/kg/day | (—) | (—) | Drinking water | (—) µg/kg/day | MOE | (—) | (—) | (—) |
| | | | | Public Freshwater bodies | (—) µg/kg/day | MOE | (—) | (—) | |
| Inhalation | ‘Non-toxic level*’ 0.56 mg/m ³ | Mice | Hepatocellular adenomas | Ambient air | 0.85 µg/m ³ | MOE | 13 | ▲ | ▲ |
| | | | | Indoor air | — µg/m ³ | MOE | — | × | (▲) |

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 EC₅₀ of 246 µg/L for growth inhibition in the green algae *Chlamydomonas reinhardtii*, a 48-h EC₅₀ of 8,090 µg/L for immobilization in the crustacean *Daphnia magna*, and a 96-h LC₅₀ of 7,610 µg/L for the fish species *Oryzias latipes* (medaka). Accordingly, based on these acute toxicity values and an assessment factor of 100, a predicted no effect concentration (PNEC) of 2.4 µg/L was obtained.

With regard to chronic toxicity, the following reliable data were obtained: a 72-h NOEC of 121 µg/L for growth inhibition in the green algae *P. subcapitata*, and a 21-d NOEC of 494 µg/L for reproductive inhibition in the crustacean *D. magna*. Accordingly, based on these chronic toxicity values and an assessment factor of 100, a PNEC of 1.2 µg/L was obtained.

The value of 1.2 µg/L obtained from the chronic toxicity to the algae was used as the PNEC for this substance.

The PEC/PNEC ratio is less than 0.3 for freshwater bodies and less than 0.4 for seawater. Accordingly, efforts to collect data are needed. The lower detection limit for this substance in freshwater is 2 µg/L according to one data source; accordingly, efforts are needed to improve the sensitivity of environmental concentration measurements and understand PRTR data trends. Further, when releases to public freshwater bodies in fiscal 2014 reported according to the PRTR Law were divided by the ordinary water discharge of the national river channel structure database, estimating the concentration in rivers by taking into consideration only dilution gave a maximum value of 0.00014 µg/L.

| Hazard Assessment (Basis for PNEC) | | | Assessment Coefficient | Predicted no effect concentration PNEC (µg/L) | Exposure Assessment | | PEC/PNEC ratio | Judgment based on PEC/PNEC ratio | Assessment result |
|------------------------------------|---------------|------------------------|------------------------|---|---------------------|--|----------------|----------------------------------|-------------------|
| Species | Acute/chronic | Endpoint | | | Water body | Predicted environmental concentration PEC (µg/L) | | | |
| Green algae | Chronic | NOEC growth inhibition | 100 | 1.2 | Freshwater | 0.35 | 0.3 | ▲ | ▲ |
| | | | | | Seawater | <0.5 | <0.4 | | |

5. Conclusions

| | Conclusions | | Judgment |
|-----------------|-----------------------------------|--|----------|
| Health risk | Oral exposure | It was not the object of evaluation. | (—) |
| | Inhalation exposure (atmosphere) | Requiring information collection. | ▲ |
| | Inhalation exposure (room air) | Collection of further information would be required. | (▲) |
| Ecological risk | Requiring information collection. | | ▲ |

[Risk judgments] ○: No need for further work ▲: Requiring information collection
 ■: Candidates for further work ×: Impossibility of risk characterization
 (○) : Although risk to human health could not be confirmed, collection of further information would not be required.

(▲) : Further information collection would be required for risk characterization.

(—) : It was not the object of evaluation.