012255

RPA 202248 Acute Oral Toxicity (81-1)

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DATA EVALUATION REPORT

STUDY TYPE:	Acute Oral Toxicity/Rats OPPTS 870.1100 [81-1]
EPA ID NUMBERS:	DP BARCODE: D227954 P. C. CODE: 123000 SUBMISSION NO.: S508413 MRID NUMBER: 44044701
TEST MATERIAL: CHEMICAL NAME:	RPA 202248 2-Cyano-3-cyclopropyl-1-(2-methylsulfonyl-4-trifluoromethylphenyl)-propan-1,3-dione
STUDY NUMBER:	SA 96140
TESTING FACILITY:	Rhone-Poulenc Agrochimie, Sophia Antipolis, France
SPONSOR:	Rhone-Poulenc Agrochimie, Lyon, France
TITLE OF REPORT:	RPA 202248. Oral LD_{50} in the Rat
AUTHOR:	P. Katchadourian
REPORT ISSUED:	May 22, 1996

EXECUTIVE SUMMARY: In an acute oral toxicity study (MRID # 44044701), groups of five male and five female Sprague-Dawley rats received single oral administration of RPA 202248 in 0.5% aqueous methylcellulose at dose levels of 2000, 2710, 3690 and 5000 mg/kg. Two males and one female at 3690 and one male at 5000 mg/kg died by Day 3. The clinical signs of toxicity observed in both sexes, within few hours of dosing, included piloerection at all dose levels and hunched posture in males at 2710 mg/kg as well as in both sexes at 3690 and 5000 mg/kg/day. Reduced motor activity was noted in both sexes at 3690 and 5000 mg/kg/day. **Based on the estimated acute oral LD_{50} of >5,000 mg/kg in both sexes, RPA 202248 is placed in Toxicity Category IV.**

The study is classified as **Acceptable** and **satisfies** the requirements (81-1) for an acute oral toxicity study in rats.

I. MATERIALS

A. Test Material

Name: RPA 202248
Chemical Name: 2-Cyano-3-cyclopropyl-1-(2-methylsulfonyl-4-trifluoromethylphenyl)propan-1,3-dione
Purity: 99.9%
Batch Number: SR0712
Description: White powder
Storage Conditions: At room temperature protected from light

The dosing formulation was prepared by suspending the test substance in 0.5% methylcellulose in distilled water to produce required dosing concentrations (w/v).

B. Test Animals

Species: Sprague-Dawley rats (ICO OFA SD) (IOPS Caw)
Source: Iffa-Credo, L'Arbresle, France
Age: 8 weeks at initiation of dosing
Weight: Males - 237 to 281 g; Females - 173 to 191 g when dosed
Housing: Individually in stainless steel cage
Environmental Conditions: Temperature: 20–24°C
Relative Humidity: 40-70%
Photoperiod: 12 hours light/dark
Air Changes: 10 to 15/hour
Food and Water: Pelleted Certified Rodent Diet A04C (from Usine d'Alimentation Rationnelle, France) and tap water *ad libitum*
Acclimation Period: 14 days

II. METHODS

Animals were fasted overnight prior to dosing. Each animal received a single dose of the test solution. Five male and five female rats/group were dosed with test solution (20 ml volume/kg b.w.) at 2000, 2710, 3690 or 5000 mg/kg via gavage. The animals were observed for mortality and clinical signs of toxicity on two occasions during Day 1 and at least once daily for the remainder of the 14-day observation period. Body weights were recorded ten days prior to dosing, on the day of dosing and at 8 and 15 days post-dosing. At the end of the observation period, all animals were sacrificed and necropsied.

III. RESULTS

At 2000 and 2710 mg/kg, none of the animals died during the study. No clinical signs or macroscopic postmortem findings were observed at 2000 mg/kg. Mortalities were noted at 3690 mg/kg on Day 2 (one female and two males) and at 5000 mg/kg/day on Day 3 (one male); treatment-related clinical signs of toxicity were observed within few hours of dosing. Piloerection was noted in animals at all dose levels and hunched posture was reported in males at 2710 mg/kg as well as in both sexes at 3690 and 5000 mg/kg/day. In addition, males and females at 3690 and 5000 mg/kg exhibited reduced motor activity. The surviving animals recovered on Day 5. Body weights were unaffected. There were no lesions observed on gross necropsy of animals sacrificed at study termination. The estimated acute oral LD_{50} was >5000 mg/kg for males and females.

IV. COMPLIANCE

The following compliance documents were submitted: 1) signed statement by the sponsor indicating that the study was conducted in accordance with GLP Regulations; 2) signed Quality Assurance statement by the testing facility; 3) signed statement by the sponsor claiming no data confidentiality. Any deviations from the protocol were appropriately reported.

V. CONCLUSIONS

The acute oral LD_{50} for RPA 202248 in male and female rats was >5,000 mg/kg. Toxicity Category IV.

The study is classified as **Acceptable** and **satisfies** the requirements (81-1) for an acute oral toxicity study in rats.

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One-Liner for RPA 202248

Study Type:	Guideline: OPP §81-1 Acute Oral Toxicity Species: Rat
Test Material:	RPA 202248
Chemical Name:	2-Cyano-3-cyclopropyl-1-(2-methylsulfonyl-4-trifluoromethylphenyl)-propan-1,3-dione
EPA MRID No.:	44044701
Testing Facility:	Rhone-Poulenc Agrochimie, Sophia Antipolis, France
Study Number:	SA 96140
Report Issued:	May 22, 1996

Executive Summary: In an acute oral toxicity study (MRID # 43904810), groups of five male and five female Sprague-Dawley rats were orally administered RPA 202248 in 0.5% aqueous methylcellulose (20 ml/kg b.w.) at dose levels of 2000, 2710, 3690 and 5000 mg/kg. Two males and one female at 3690 and one male at 5000 mg/kg died by Day 3. The clinical signs of toxicity observed in both sexes, within few hours of dosing, included piloerection at all dose levels and hunched posture in males at 2710 mg/kg as well as in both sexes at 3690 and 5000 mg/kg/day. Reduced motor activity was noted in both sexes at 3690 and 5000 mg/kg/day. **Based on the estimated acute oral LD_{50} of >5,000 mg/kg in both sexes, RPA 202248 is placed in Toxicity Category IV.**

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