(TXR 013599) (8-3-99) 24

[Trifloxystrobin] Acute Oral Study (81-1)

EPA Toxicologist: Deborah C. Smegal_____, Date _____
Toxicology Branch II (7509C)
EPA Secondary Reviewer: _____, Date _____
(7509C)

DATA EVALUATION RECORD

STUDY TYPE: Acute Oral Toxicity - Rat
OPPTS 870.1100 [§81-1]

DP BARCODE: 243979 SUBMISSION CODE:S538757
P.C. CODE:129112 TOX. CHEM. NO.:

TEST MATERIAL (PURITY): CGA-279202 Technical(96% a.i.)

SYNONYMS:Trifloxystrobin

CITATION: Glaza, S.M. 1994. Acute oral toxicity study of CGA-279202 Technical in rats. Covance Laboratories Inc. (Formerly Hazleton Wisconsin). 3301 Kinsman Boulevard. Madison, WI. 53704. Laboratory Project ID: HWI 40702444. Novartis Nexus Number 355-94. October 5, 1994. MRID 44496622. Unpublished.

SPONSOR: Novartis Crop Protection, Inc.

EXECUTIVE SUMMARY: In an acute oral toxicity study (MRID 44496622),a group of fasted, albino Crl:CD (SD)BR rats (5/sex) was given a single oral dose of CGA-279202 Technical(96% a.i.)in corn oil at a dose of 5,000 mg/kg and observed for 14 days.

Oral LD_{50} Males = >5,000 mg/kg
Females = >5,000 mg/kg
Combined = >5,000 mg/kg

CGA-279202 Technical is classified as TOXICITY CATEGORY IV based on the LD_{50} in both males and females.

There were no treatment-related effects on mortality or necropsy findings. Clinical signs of toxicity included hypersensitivity to touch, red stained face, excessive salivation, soft or watery stool, and dark stained or wet urogenital area. Symptoms subsided in all animals by day 12 post treatment. Decreased body weight (3-4%) was noted in three of five females during the 7 to 14 day interval, however no effects were noted in males.

This acute oral study is classified as acceptable. This study satisfies the guideline requirement for an acute oral study (81-1) in the rat.

COMPLIANCE: Signed and dated GLP, Quality Assurance, and Data Confidentiality were provided. Flagging statements were not

provided.

I. MATERIALS AND METHODS

A. MATERIALS:

1. Test Material: CGA-279202 Technical
   Description: tan powder
   Lot/Batch #: FL-941274, batch code P.405009
   Purity: 96% a.i.
   CAS #: 141517-21-7

2. Vehicle and/or positive control: corn oil

3. Test animals: Species: rats
   Strain: albino Crl:CD (SD)BR
   Age and/or weight at dosing: young adult, 235-299 g
   Source: Charles River Laboratories, Inc., Portage, Michigan
   Acclimation period: 7 days
   Diet: Laboratory Rodent Diet #5001, PMI Feeds, Inc ad libitum
   Water: administered ad libitum, except for 17 to 20 hours before compound administration
   Housing: Groups of 5 same sex were housed in screen-bottom stainless steel cages.

   Environmental conditions:
   Temperature: 19 - 25°C
   Humidity: 50 ±20%
   Air changes: not described
   Photoperiod: 12 hours light/ 12 hours dark

B. STUDY DESIGN and METHODS:

1. In life dates - not reported

2. Animal assignment and treatment: Animals were assigned to the test groups noted in table 1. Following an overnight fast, rats were given a single dose of CGA-279202 Technical by gavage in a volume of 20 ml/kg. They were then observed for mortality and clinical signs of toxicity approximately 1, 2.5 and 4 hours after treatment. Thereafter, clinical signs were checked daily and mortality was monitored twice daily for 14 days. They were weighed prior to treatment and again on days 7 and 14. Necropsy was performed on the survivors, which were sacrificed on day 14.

TABLE 1. Doses and Mortality/Animals Treated

Dose (mg/kg)	Males	Females	Combined
5,000	0/5	0/5	0/10

3. Statistics: The oral LD_{50} exceeds the limit dose of 5,000 mg/kg, therefore, no statistics were used to calculate this value.

II. **RESULTS AND DISCUSSION:**

A. Mortality is given in table 1.

The calculated acute oral LD_{50}'s are as follows:
males: >5,000 mg/kg
females: >5,000 mg/kg
combined: >5,000 mg/kg

B. Clinical observations: There were no treatment-related effects on mortality. All animals survived to the scheduled sacrifice. Clinical signs of toxicity included hypersensitivity to touch, red stained face, excessive salivation, soft or watery stool, and dark stained or wet urogenital area. Symptoms subsided in all animals by day 12 post treatment.

C. Body Weight: There were no treatment-related effects on body weight in males. Decreased body weight (3-4%) was observed in 3 of 5 females between days 7 and 14, while there was no body weight change in 1 female during this time interval.

D. Necropsy: No abnormalities were observed in post mortem examination of the animals that were sacrificed.

E. Deficiencies: The specific in life dates were not reported. In addition, the animals were dosed at a volume of 20 ml/kg body weight, which exceeds the 10 ml/kg body weight volume.

SignOff Date:	8/3/99
DP Barcode:	D243979
HED DOC Number:	013599
Toxicology Branch:	TOX2