(8-3-99) 23

[Trifloxystrobin metabolite] (TXR 013599) 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): NOA-414412 Tech. (Metabolite of CGA-279202)(95 ± 2%)

SYNONYMS:

CITATION: Cantoreggi, S. 1997. NOA-414412 Tech. (Metabolite of CGA-279202). Acute oral toxicity in the rat (Limit test). Toxicology/Experimental Toxicology. Novartis Crop Protection, AG (Formally Ciba-Geigy Limited). 4332 Stein, Switzerland. Test No. 973064. Novartis Nexus Number 737-97. October 20, 1997. MRID 44496621. Unpublished.

SPONSOR: Novartis Crop Protection, Inc.

EXECUTIVE SUMMARY: In an acute oral toxicity study (MRID 44496621),a group of fasted, albino WISTAR Han rats (5/sex) was given a single oral dose of NOA-414412 Tech.(95 ± 2%)in 0.5% carboxymethylcellulose (w/v) in 0.1% (w/v)aqueous polysorbate 80 at a dose of 2,000 mg/kg and observed for 14 days.

Oral LD_{50} Males = >2,000 mg/kg
Females = >2,000 mg/kg
Combined = >2,000 mg/kg

NOA-414412 Tech.is classified as TOXICITY CATEGORY III based on the LD_{50} in both males and females.

There were no treatment-related effects on mortality, body weight or necropsy findings. Treatment-related, but transient clinical signs included piloerection and hunched posture in all males that subsided within one day. Females did not exhibit any treatment-related clinical signs.

This acute oral study is classified as acceptable for a metabolite. This study satisfies the guideline requirement for an acute oral study (81-1) in the rat. This study follows OECD Guideline and OECD principles of GLP.

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: NOA-414412 Tech. (Metabolite of CGA-279202)
Description: solid
Lot/Batch #: MLA-31/4
Purity: 95 ± 2%
CAS #: not available

2. Vehicle and/or positive control: 0.5% (w/v) carboxymethylcellulose in 0.1% (w/v) aqueous polysorbate 80

3. Test animals: Species: rats
Strain: albino WISTAR Han
Age and/or weight at dosing: young adult, 180 - 194 g
Source: BRL Biological Research Laboratories, Ltd, 4414 Fullinsdorf, Switzerland
Acclimation period: 5 days
Diet: NAFAG 890 ad libitum
Water: administered ad libitum

Housing: Groups of 5 same sex were housed in Macrolon cages type 4, with standardized soft wood bedding.

Environmental conditions:
Temperature: 22 ±2°C
Humidity: 55 ±10%
Air changes: 15 per hour
Photoperiod: 12 hours light/ 12 hours dark

B. STUDY DESIGN and METHODS:

1. In life dates - start: 9/23/97 end: 10/9/97

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 NOA-414412 Tech. by gavage in a volume of 10 ml/kg. They were then observed for mortality twice daily, and clinical signs of toxicity daily for 14 days. They were weighed prior to treatment and again on days 7 and 14. Necropsy was performed on animals that were sacrificed on day 14.

TABLE 1. Doses and Mortality/Animals Treated

Dose (mg/kg)	Males	Females	Combined
2,000	0/5	0/5	0/10

3. Statistics: The oral LD_{50} exceeds the limit dose of 2,000 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: >2,000 mg/kg
females: >2,000 mg/kg
combined: >2,000 mg/kg

B. Clinical observations: There were no treatment-related effects on mortality. All animals survived to the scheduled sacrifice. Treatment-related, but transient clinical signs included piloerection and hunched posture in all males that subsided within one day. Females did not exhibit any treatment-related clinical signs.

C. Body Weight: There were no treatment-related effects on body weight.

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

E. Deficiencies: None.

SignOff Date:	8/3/99
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