



UNITED STATES ENVIRONMENTAL PROTECTION AGENCY
WASHINGTON, D.C. 20460

OFFICE OF PREVENTION,
PESTICIDES AND TOXIC SUBSTANCES

13/MAY/2001

MEMORANDUM

Subject: EPA Reg. No: 352-ARN
DP Barcode: D272547
Case No: 070000
PC Code: 128845, 128887, 122010

From: Masih Hashim, Toxicologist
Technical Review Branch
Registration Division (7505C) *MHS SCR*

To: Vickie Walter, PM Team 25
Herbicide Branch
Registration Division (7505C)

Applicant: E.I du Pont De Nemours and Company
Walker's Mill, Barley Mill Plaza
Wilmington, DE 19880-0038

FORMULATION FROM LABEL:

<u>Active Ingredient(s):</u>	<u>% by wt.</u>
Thifensulfuron methyl	37.5
Tribenuron	18.7
Metsulfuron-methyl	15.0
<u>Inert Ingredients</u>	<u>28.8</u>
Total:	100.0

①

BACKGROUND: DuPont De Nemours and Company has submitted a set of toxicity studies to support the registration of its Product DuPont Canvas px Herbicide, File Symbol 352-ARN. The studies were conducted at the Haskell Laboratory of DuPont in Newark, DE. The company requested a waiver for three studies as discussed in the next paragraph.

RECOMMENDATIONS: Each of the toxicity studies is in accordance with the Sub Division F guideline. The Company requested a waiver for the acute inhalation, skin sensitization and skin irritation* studies (letter dated 12-20-2000, MRID 452939-00) so that the cited study 352-586 may be used for a similarity action. However, Tribenuron methyl is the additional herbicide in the proposed product will not qualify the two products for a similarity action. TRB would require the three referenced studies to fill in the data gap. If there are any questions, please let us know.

a-(meant to be acute dermal which is missing)

acute oral toxicity	IV	acceptable MRID 4308010-04
acute dermal toxicity	-	data gap
acute inhalation	-	data gap
primary eye irritation	IV	acceptable MRID 452939-06
primary dermal irritation	IV	acceptable MRID 452939-05
dermal sensitization	-	data gap

LABELING: Based on the acute toxicity studies, the following precautionary labeling is recommended. However, it will be complete only after the required studies (discussed in the last paragraph) are submitted by the Registrant.

Date: 05/15/01 LABEL REVIEW SYSTEM

ID #: 000352-00610 DuPont Canvas px Herbicide

SIGNAL WORD: CAUTION

PRECAUTIONARY STATEMENTS:

Wear long-sleeved shirt and long pants, socks and shoes..

STATEMENT OF PRACTICAL TREATMENT (SOPT):

DATA REVIEW FOR ACUTE ORAL TOXICITY STUDY (870.1100)

Product Manager: 25

Reviewer: M. Hashim

Citation: Carol Finlay, study director to conduct an acute oral toxicity study in rats with Thifensulfuron methyl, Tribenuron and Metsulfuron- methyl in a formulation for Du Pont Canvas px Herbicide. Testing Lab; Haskell Laboratory of Du Pont, Laboratory Report No. DuPpont 5410 dated 12-15-2000, MRID 452939-04. Sponsor: DuPont. Rats-weight; 185-229 g, source: Charles River, Stoneridge, NY.

Summary: *In an acute oral toxicity study using Du Pont Canvas px Herbicide was administered at a single limit dose of 5000 mg/kg to 5 SD rats per sex. Animals were observed for clinical signs and mortality for a period of 14 days.*

Oral LD50 was found to be 5000 mg/kg in both sexes, and the test article is classified as Tox Category IV.

There were no deaths on the study. The clinical signs were: one animal with lethargy and another 4 animals with white discharge from the genitals. There was some weight loss in a couple of rats. Gross pathology was unremarkable.

This study is in compliance with Sub Division guideline and is Acceptable.

DATA REVIEW FOR PRIMARY EYE IRRITATION STUDY (870.2400)

Product Manager: 25

Reviewer: M. Hashim

Citation: Carol Finlay, Study Director. Thifensulfuron methyl, Tribenuron and Metsulfuron- methyl in a formulation for Du Pont Canvas px Herbicide to conduct a Primary Eye irritation study in rabbits. Testing Lab; Haskell Laboratory of Du Pont Company, Newark, DE, Laboratory Report No. DuPont 5412. Study director carol Finlay, dated 12-15-2000, MRID 452939-06. Sponsor: DuPont. Study animal, NZW rabbits, supplier Hare Marland, Hewitt, NJ.

Summary: *In a primary eye irritation study 0.1 ml of the test article was placed into the conjunctival sac of the right eye of six young adult NZW rabbits. The other eye served as the control. Animals were observed for ocular irritation for 7 days.*

Eye irritation included corneal opacity and conjunctivitis which subsided within 7 days. The average score was 1.5. This study is classified as Tox Category III.

GLP statement accompanies the study. This study is Acceptable in accordance with the Sub Division F guidelines.

Following summary table illustrates the eye irritation response:

Lesion	1 hr	24 hrs	48 hr	72 hr	7 days
corneal opacity	2/6	0/6	0/6	0/6	0/6
iritis	0/6	0/6	0/6	0/6	0/6
conjunctivitis	0/6	4/6	4/6	1/6	0/6

DATA REVIEW FOR PRIMARY DERMAL IRRITATION STUDY (870.2500)

Product Manager: 25

Reviewer: M. Hashim

Citation: Carol Finlay, Study Director, in a formulation for Du Pont Canvas px Herbicide to conduct a dermal irritation study in NZW rabbits. Testing Lab; Haskell Laboratory of Du Pont Company, Newark, DE, Laboratory Report No. DuPont 5294. Report dated 12-15-2000, MRID 452939-05. Sponsor: DuPont. Species Rabbit, NZW, source Hare Marland, Hewitt, NJ.

Summary: *Six rabbits were used in the study. One half gram of the test article, moistened in water as a paste, was applied to the site of an intact skin of each rabbit. The test site was covered with a semi occlusive dressing for 4 hrs. Then the sites were evaluated at 1, 24, 48 and 72 hrs. (except one animal was evaluated for 6 days).*

There was no dermal irritation in 4 rabbits. One rabbit showed erythema after 24 hours, and the other one for 72 hours. Skin irritation subsided by the 6th day of observation. An over all average irritation score was calculated to be 0.4. This classifies the test compound as Category IV.

The study meets GLP compliance and is Acceptable in accordance with the Sub- Division F guidelines.

ACUTE TOX ONE-LINERS

1. DP BARCODE: D272547
2. PC CODE: 128845, 128887, 122010
3. CURRENT DATE: May 9, 2001
4. TEST MATERIAL: DuPont Canvas px Herbicide

Study/Species/Lab Study #/Date	MRID	Results	Tox. Cat.	Core Grade
Acute oral toxicity/ rat/ Haskell Lab/DuPont 5410/ 12-15-00	452939-04	LD ₅₀ > 5000 mg/kg	IV	A
Primary eye irritation/rabbit/Haskell Lab / DuPont 5412/ 12-15-00	452939-06	mild irritant	IV	A
Primary dermal irritation/rabbit/ DuPont 5294/ 12-15-00	452939-05	mild irritant	IV	A

Core Grade Key: A =Acceptable, S = Supplementary, U = Unacceptable, V = Self Validated