

4-8-82 RCB

Perfetti

File petition

To: Henry Jacoby P M 21Registration No(s):: 359-685 CFR 180-539Pesticide Petition No(s):: 2F 2596 Acc. No 070441 070443Caswell No(s):: 470 AChemical(s): Iprodione (Rovral)RAC(s) - tolerance(s): Request for tolerances of 10 ppm in appricots , nectarines, peaches , plums inc. prunes and cherries.Inert(s) cleared 180.1001: (e) Yes% of ADI occupied: Existing: 0.03 Resulting: 2.58Resulting % increase in TMRC: From 0.0045 to 0.3876Attached (?): ADI printout: YES/~~NO~~; TOX "one-liner": YES/~~NO~~; DER: YES/~~NO~~Existing regulatory actions against registration: NoneRPAR status: Is not in the R P A R listNew Data: None Submitted / Data required on behalf of this action (next page)Data considered in setting the ADI: Rat reproduction , 3 generations , study .NOEL= 500ppmData gaps: The three following study has been classified as supplementary data ; Acute dermal toxicity, second teratology study and mutagenicity other test.The studies classified as supplementary data must be upgraded.Recommendation: Granting this request will not present a hazard since the ADI is not exceededComments: A one year dog feeding study has been initiated and the report will be submitted within a year . (info obtained by phone by The PM)The study 3 generation reproduction -rat has been used to find the ADI instead of the 3 month dog feeding study. It will lead to a better appriaisal of the toxicity potential ofReviewer: Alex Arce Date: the productSection Head: William M Butte

3-31-82

APR 8 1982

1/7

Data required

		Classification
Acute oral -rat	3,700 mg/kg	m
Acute dermal -rabbit	30,000mg/kg male -5,000 mg/kg female	s
Acute inhalation-rat	3.29mg/l/4h	m
Primary eye-skin-rabbit	Not an irritant	m
90 day feeding -dog	NOEL 2,400 ppm - LEL 7,200 ppm	m
Teratology - rat	NOEL 400 mg/kg /day	m
Teratogenicity - rabbit	NOEL 400 mg/kg/day	s
Chronic oncogenicity -rat 24 month feeding	NOEL 1,000 ppm	m
18 months oncogenicity -mouse	NOEL 1,250 ppm	m
Mutagenicity , dominant lethal - mouse	No evidence of mutagenicity at 1,500 or at 6,000 ppm	m
Three generations reproduction study -rat	NOEL 500 ppm	m

Justification for using the 3 generation reproduction study -rat instead of the
3 month dog feeding study, to find the A D I

The NOEL in the 3 months feeding study -(dog) was found at the 2,400 ppm dose level. The LEL was 7,200 ppm and the symptom observed was muscular atony , no other signs of toxicity were reported

The three generation reproduction study (rat) with a NOEL of 500 ppm and with no toxic signs observed in the parent animals, will give a better indication of the potential toxicity of the material . The only adverse effect observed during the 3 generations study was the post natal pupgrowth for group 1V that was reduced in a minor percent as compared with group 1.

File last updated 4/1/82

ACCEPTABLE DAILY INTAKE DATA

RAI, Older NOEL	S.F.	ADI	DPI
mg/kg	pm	mg/kg/day	mg/day (60kg)
25.000	500.00	100	0.2500
			15.0000

Published Tolerances

CROP	Tolerance	Food Factor	mg/day (1.5kg)
Kiwi Fruit (204)	10.000	0.03	0.00450

ADI	THRC	% ADI
15.0000 mg/day (60kg)	0.0045 mg/day (1.5kg)	0.03

Unpublished, Tox Approved PEF EG2087, OG2402

CROP	Tolerance	Food Factor	mg/day (1.5kg)
Almonds(1)	0.050	0.03	0.00002
Apricots(3)	10.000	0.11	0.01688
Cherries(30)	10.000	0.10	0.01533
Nectarines(100)	10.000	0.03	0.00450
Peaches(114)	10.000	0.90	0.13490
Plums, inc prunes(125)	10.000	0.13	0.01993

ADI	THRC	% ADI
15.0000 mg/day (60kg)	0.1950 mg/day (1.5kg)	1.31

Current Action PEF 2F2596

CROP	Tolerance	Food Factor	mg/day (1.5kg)
Apricots(3)	10.000	0.11	0.01688
Cherries(30)	10.000	0.10	0.01533
Nectarines(100)	10.000	0.03	0.00450
Peaches(114)	10.000	0.90	0.13490
Plums, inc prunes(125)	10.000	0.13	0.01993

ADI	THRC	% ADI
15.0000 mg/day (60kg)	0.3876 mg/day (1.5kg)	2.58

Study/Lab/Study #/Date	Material	EPA Accession No.	Results:		TOX Category	CORE Grade/ Doc. No.
			LD50, LC50, PIS, NOEL, LEL			
Acute oral LD50, mice	TECH	232701	LD50 = 3050 mg/kg (2630-3540)		III	Minimum
Acute oral LD50, mice	TECH	232701	LD50 = 4.0 (3.3-4.8) g/kg - M ✓ 4.4 (3.3-5.9) g/kg - F ✓		III	Minimum
Acute oral LD50, rat	TECH	232701	LD50 = 3700 ± 300 mg/kg - M = 4400 (3200-6100) mg/kg - F		III	Minimum
Acute oral LD50, dog	TECH	232701	Atoxic at 2000 mg/kg		III	Supplementary
Acute dermal LD50, rat	TECH	232701	Atoxic at 2.5 g/kg		III	Supplementary
Acute dermal LD50, rabbit	TECH	232701	Atoxic at 1 g/kg		IV	Supple- mentary
Primary dermal irrita- tion - rabbit	TECH	232701	LD50 > 30,000 mg/kg - M > 5,000 mg/kg - F		IV	Supple- mentary
Primary dermal irrita- tion - rabbit	TECH	232701	Not irritating at 1 g/kg		IV	Minimum
Primary dermal irrita- tion - rabbit	TECH	232701	Not an irritant		IV	Supple- mentary
Primary dermal irrita- tion - rabbit	TECH	232701	Not an irritant (Summary)		IV	Minimum
Primary eye irrita- tion - rabbit	TECH	232701	Not an irritant		IV	Minimum
Acute inhalation LD50, rat	TECH	232701	LC50 = > 3.29 mg/L/4 hr.		IV	Minimum
Sensitization dermal - guinea pig	TECH	232701	Negative		IV	Supple- mentary
28-Day Oral - mice	TECH	232701	NOEL = 1,900 ppm			Supple- mentary

Study/Lab/Study #/Date 90-Day feeding, mice	Material	EPA Accession No.	LD ₅₀ , LC ₅₀ , PIS, NOEL, LEL NOEL = 1,900 ppm	Results:	TOX Category	CORE Grade/ Doc. No.
90-Day feeding, dog	TECH	232701	NOEL = 2,400 ppm LEL = 7,200 ppm			Supplementary
5-Month feeding, rat	TECH	232701	NOEL $\geq$ 1,000 ppm (highest dose tested)			Minimum
24-Month feeding, rat	TECH	097201	NOEL $\geq$ 1,000 ppm (highest dose tested)			Minimum
18-Month feeding - oncogenicity, mice	TECH	097201	NOEL $\geq$ 1,250 ppm (highest dose tested)			Minimum
Teratogenic - rat	TECH	232712	Not carcinogenic. Teratogenic NOEL $\geq$ 400 mg/kg/day (highest level tested) Fetotoxic NOEL = 200 mg/kg/day Fetotoxic LEL = 400 mg/kg/day (decrease in mean number of implantation sites) Maternal NOEL = 200 mg/kg/day Maternal LEL = 400 mg/kg/day (decrease in conception rate and food consumption) Teratogenic NOEL $\geq$ 400 mg/kg/day (highest level tested) Fetotoxic NOEL = undetermined Maternal LEL = undetermined			Minimum
Teratogenic - rabbit	TECH	232712				Supple- mentary 000614

Tox Chem No. 470A

File Last Updated 2-3-81

Current Date 7-11-81

EPA

Accession

No.

Material

Study/Lab/Study #/Date

Results:

LD₅₀, LC₅₀, PIS, NOEL, LEL

TOX

Category

CORE Grade/
Doc. No.Mutagenicity domi-
nant lethal - mice

TECH

232712

No evidence of mutagenicity or
adverse effects on fertility at
1500 or 6000 ppm dose males

Minimum

Mutagenicity - micro-
biologic

TECH

232712

Negative

Supple-
mentary3-Generation repro-
duction - rat

TECH

232712

NOEL = 500 ppm

Minimum

SECTION F

PROPOSED PERMANENT TOLERANCES FOR RESIDUES OF THE FUNGICIDE
IPRODIONE, ITS ISOMER AND METABOLITE IN OR ON THE RAW
AGRICULTURAL COMMODITY STONE FRUITS INCLUDING FRESH AND
PROCESSED FRUITS

Stone Fruits

It is hereby proposed a tolerance for the combined residues of iprodione [3-(3,5-dichlorophenyl)-N-(1-methyl ethyl)-2,4-dioxo-1-imidazolidinecarboxamide], its isomer [3-(1-methyl ethyl)-N-(3,5-dichlorophenyl)-2,4-dioxo-1-imidazolidinecarboxamide], and its metabolite [3-(3,5-dichlorophenyl)-2,4-dioxo-1-imidazolidinecarboxamide] in and/or on:

apricot, cherries (sour and sweet), nectarines,
peaches, plums (fresh prunes) 10 ppm

Animal Products

It is hereby proposed a modified use to restrict grazing livestock in treated orchard and to restrict the feeding of cover crops, therefore, no tolerances for residues in meat, milk, poultry and eggs would be needed. Section 180.6 (a) (3).