Chemical Name: Diphenyl cresyl phosphate
Trade Name(s): Kronitex CDP
CAS No: 26444-49-5
Lab Study ID No: CIBA-GEIGY 49/79/15. L. 15 Mar 1979

FINAL REPORT ACTION ITEM CHECK-OFF LIST

- ☐ Reviewed for possible:
  - ☐ FIFRA 6 (a) (2) and/or
  - ☐ TSCA Section 8 (e) reporting
- ☐ Copy of FIFRA 6 (a) (2) and/or TSCA Section 8 (e) letter to the following Agency(ies), if applicable:
  - ☐ EPA-FIFRA
  - ☐ EPA-TSCA
  - ☐ California [FIFRA 6 (a) (2)s]
  - ☐ Other States [FIFRA 6 (a) (2)s]: _____
- ☐ Confidentiality Statement page addressed, signed, and dated in FIFRA reports
- ☐ GLP Compliance page signed and dated in FIFRA reports
- ☐ Flagging Statement page addressed, signed and dated in FIFRA reports
- ☐ Copy of report submitted to the Agency(ies) in conjunction and/or support of one or more of the following:
  - ☐ TSCA Consent Order/Agreement
  - ☐ FIFRA Registration or Re-registration
  - ☐ California Registration
  - ☐ EU Notification
  - ☐ Japanese MITI Notification
  - ☐ Japanese MAFF Notification
  - ☐ Canadian (DSL) Notification
  - ☐ FIFRA 6 (a) (2) Submission
  - ☐ TSCA 8 (e) Submission
  - ☐ TSCA 8 (d) Data-Call-In
  - ☐ PMN Submission
  - ☐ Other: _____
- ☒ All information regarding the chemical and the study report entered into the IUCLID Toxicity Data Base
- ☒ Study Report reviewed for MSDS information Already on MSDS
- ☒ Copy of Cover & Summary report pages to Business Unit MSDS information center (domestic and international)
- ☐ Active Study file merged with final report in Regulatory Affairs file room

On MSDS

TOX K

GPP ✓
PC

(85)

CIBA-GEIGY Pharmaceuticals Division
STAMFORD LODGE UK
PHARMA TOXICOLOGY

49/79/S.L.
CONFIDENTIAL

p-CRESYL DIPHENYL PHOSPHATE

N.B.

LIC 124/47

Delayed Neurotoxicity in Domestic Hens (ED_{50})

(2,500 ppm oCDP — could be enough to account for observed effect — comment, pending results on dilution(s) of oCDP in other TAPs

23/3/79

AJD

15th March, 1979.

Toxicology Department,
Geigy Pharmaceuticals,
Stamford Lodge,
Wilmslow

A. J. Duke
Trafford Park

Toxicology tests required and species of animals to be used

Chicks neurotoxicity single doses of 200 mg/kg ± 2 g/kg. Please report any positive clinical observations as soon as noted.

Chemical name and formula

LIC 124/47

By GLC - 97.4 % p-cresyl diphenyl phosphate
0.3 % triphenyl phosphate
2.3 % bis-(p-cresyl) phenyl phosphate

(< 0.01 % o-cresyl diphenyl phosphate)

Intended use

Reference sample, main component of triaryl phosphate products

Other relevant information i.e. possible hazards, toxicity of similar chemicals if known, etc.

None known: Possible neurotoxicity speculated by analogy with p-ethyl phenyl phosphates

Lab. Notebook No. or Plant batch No.

V. i.

To whom report is to be sent

A. J. Duke
Dr. ~~H. Fletcher~~
~~Heald Green~~ Trafford Park

Date chemical to be sent to Stamford Lodge and amount

16/8/78
ca. 300 g.

Date: 16/8/78

Signed: Agnes J Duke

ALL REQUESTS FOR TOXICOLOGY STUDIES SHOULD BE SENT TO HEAD OF TOXICOLOGY DEPARTMENT AT STAMFORD LODGE

Test Substance	:	LIC 124/47
Description of Study	:	Delayed Neurotoxicity in Domestic Hens (ED_{50})
Study Sponsor	:	CIBA-GEIGY (U.K.) Ltd., Tenax Road, Trafford Park. Industrial Chemicals Division
Testing Facility	:	CIBA-GEIGY Pharmaceuticals Division Stamford Lodge Wilmslow Cheshire UK Pharma Toxicology
Test No.	:	7154 78
Experimental No.	:	78C03
Starting Date	:	21.8.78
Finishing Date	:	29.1.79
Location of Raw Data	:	As for testing facility
Study Director	:	..... 15.3.79 (W. A. Bradley, BVSc, MRCVS)
Responsible for conduct of animals studies	:	..... 15.3.79 (K. O'Brien, AIAT)
Head of Toxicology, CIBA-GEIGY Pharmaceuticals Division Stamford Lodge Wilmslow Cheshire UK	:	..... 15.3.79 (W. A. Bradley, BVSc, MRCVS)

ASSESSMENT

The median oral effective dose (ED_{50}) for delayed neurotoxicity in hens with LIC 124/47 was 1.70g/kg body weight (95% confidence limits 1.18 and 2.31g/kg).

EXPERIMENTAL PROCEDURE

Hens

A hybrid heavy variety of Rhode Island Red x Light Sussex hens with a mean body weight of 2.44kg were used. All birds had been purchased at 'end of lay' and had been maintained on the premises for at least 6 months and were between 2 - 3 years old. All were clinically healthy at the start of the test.

Husbandry

Twelve birds were placed in one pen 2 weeks prior to the start of the test.

All birds were allowed access to a commercial diet (BOCM SILCOCK, MANDEVILLE BLUE, LAYERS MEAL), limestone grit and water on an ad lib basis.

Compound

A clear liquid labelled LIC 124/47*

Administration of Compound

The compound, as supplied was administered as a single dose by gavage into the crop, at the required dose level.

The hens were divided into 2 series of 6 birds in each (an odd numbered and an even numbered series). Two hens, one from each series, were dosed intially and then depending on the clinical symptoms seen a further 2 hens were dosed 21 days later with either an increased dosage (no clinical symptoms in preceding hen in series) or a reduced dosage (clinical symptoms in preceding hen in the series). This was repeated until 6 hens had been dosed in each series.

*Label not legible

SUMMARY OF RESULTS

Clinical and Pathological Findings

Hen No.	Dose (g/kg)	Day	Symptoms
11	2.0	11	Ataxia.
		15	Severe ataxia.
		21	No change. Hen killed.
21	1.25	21	No symptoms. Hen killed.
31	2.0	21	No symptoms. Hen killed.
41	3.0	14	Slight ataxia.
		21	Ataxia. Hen killed.
51	2.0	13	Slight ataxia.
		21	No change. Hen killed.
61	1.25	21	No symptoms. Hen killed.
12	2.0	11	Ataxia.
		14	Severe ataxia. Hen killed.
22	1.25	21	No symptoms. Hen killed.
32	2.0	10	Slight ataxia.
		21	Ataxia. Hen killed.
42	1.25	21	No symptoms. Hen killed.
52	2.0	12	Ataxia.
		21	No change. Hen killed.
62	1.25	21	No symptoms. Hen killed.

INDIVIDUAL BODY WEIGHTS AT THE TIME OF DOSING

Hen No.	Body Weight (kg)	Hen No.	Body Weight (kg)
11	2.20	12	2.60
21	2.45	22	2.45
31	2.70	32	2.50
41	2.40	42	2.70
51	2.45	52	2.35
61	2.25	62	2.25
Mean	2.41	Mean	2.48

ACUTE TOXICITY

DATE 21·8·78

Exp. No. 78C03 Compound LIC 124/47 Route ORAL Species HEN

Series 1 Sex

E*	Dose g/kg	Log Dose	R*
11	2·0	0·3010	+
21	1·25	0·0969	−
31	2·0	0·3010	−
41	3·0	0·4771	+
51	2·0	0·3010	+
61	1·25	0·0969	−
	2·0	0·3010	
		1·5739	
1		0·2623	

Series 2 Sex

E*	Dose g/kg	Log Dose	R*
12	2·0	0·3010	+
22	1·25	0·0969	−
32	2·0	0·3010	+
42	1·25	0·0969	−
52	2·0	0·3010	+
62	1·25	0·0969	−
	2·0	0·3010	
Sum		1·1937	
Mean		0·1990	

Series 3 Sex

E*	Dose /kg	Log Dose	R*
0			
1			
2			
3			
4			
5			
6			
7			
8			
9			
Sum			
Mean			

Series 4 Sex

E*	Dose /kg	Log Dose	R*
0			
1			
2			
3			
4			
5			
6			
7			
8			
9			
Sum			
Mean			

ies 1+2 Sum of Log Doses 2·7676 Mean Log Dose 0·2306

Mean Dose 1·70g/kg

Series 3+4 Sum of Log Doses _____ Mean Log Dose _____

Mean Dose _____ Mean Dose _____

Series 1+2+3+4 Sum of Log Doses _____ Mean Log Dose _____

LD_{50} 1·70g/kg 95% Confidence Limits 1·18 and 2·31g/kg

*E = Hen No. *R = Result of Experiment + = Animal Ataxic − = Animal Normal