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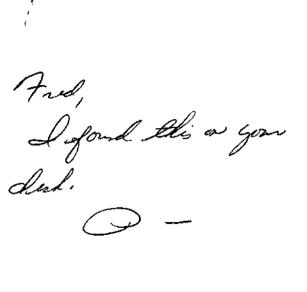
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The Toxicity of Some Organic Herbicides To Cattle, Sheep, and Chickens

Production Research Report No. 106

Agricultural Research Service
UNITED STATES DEPARTMENT OF AGRICULTURE

CONTENTS

	Page
Interpretation of hazard	2
Experimental animals	2
Experimental materials and dosages	3
Results	3
Chlorophenoxy compounds	8
Amide compounds	7
Phenyl ures compounds	
Thiocarbamate compounds	11
Trissine compounds	13
Benzolo sold compounds	18
Miscellaneous compounds	19
Comment	23
Summary and conclusions	25
Titanatura altad	98



This publication reports research involving pesticides. It does not contain recommendations for their use, nor does it imply that the uses discussed here have been registered. All uses of pesticides must be registered by appropriate State and Federal agencies before they can be recommended.

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Washington, D.C.

Issued May 1969

The Toxicity of Some Organic Herbicides To Cattle, Sheep, and Chickens

By J. S. PALMER and R. D. RADELEFF, veterinary medical officers Animal Disease and Parasite Research Division, Agricultural Research Service 1.2

The Animal Disease and Parasite Research Division, Agricultural Research Service, has conducted studies on the toxicity of various organic herbicides to livestock for many years. Initial studies at Kerrville, Tex., for the most part involved long-term experiments with the more commonly applied chlorinated phenoxy acid compounds (12, 14). Further studies with cattle and sheep have involved these and other organic herbicides. Results from some of these studies have been reported (6, 7, 8, 9, 10, 13).

Complementing all these studies have been additional reports that substantiate our own conclusions that a number of organic herbicides provide a comfortable margin of safety for cattle, sheep, and chickens when these herbicides are used according to directions (1, 2, 3, 4, 11, 15, 16,

19, 20). The studies at Kerrville were intensified, beginand chickens were added as experimental subjects. Usually these were short-term studies and were limited to 10, or fewer, consecutive daily doses. This would be a type of exposure that might be expected when these test animals consume relatively large quantities of the organic herbicides, either by carelessness or accident, for a short time. In some instances, however, sheep were exposed to as many as 481 consecutive daily doses.

The 29 organic herbicides chosen for our studies

were registered by the Pesticides Regulation Division of the Agricultural Research Service, at the time the studies were conducted. They were selected to allow as much diversification as possible, and correlated in considerable degree to the extent they were being used in agriculture. The herbicides in this report have been classified according to chemical similarity as follows: chlorophenoxy, amide, phenyl urea, thiocarbamate, triazine, benzoic acid, and miscellaneous compounds.

In this report poisoning was considered to have occurred when any observable sign of abnormal function or behavior was observed. In the mildest form of poisoning with most of the compounds, the signs were not spectacular. In many instances anorexia (lack or loss of appetite for feed-either partial or complete) was the only sign displayed. In these instances either the anorexia limited feed intake or the herbicide interfered with proper utilization of feed, or a combination of both could have affected the test animals. When such factor(s) resulted in either a 5-percent or more weight loss in cattle and sheep or a 5-percent or more decreased weight gain in chickens, it was considered significant. A few of the test animals died without previous signs of poisoning.

Cattle and sheep were routinely studied without parallel controls. Special tests that have been reported previously indicate that animals treated at less than toxic doses gained weight comparable with that of control animals (9, 21). The behavior and weight changes of cattle and sheep in these and other current studies and of animals in previous studies formed the basis for judging effect. In some measure the assessment of effect upon weight was therefore subjective rather than

objective in cattle and sheep.

Chickens were studied with parallel controls and fed to make extensive weight gains. Because of the large numbers available, the assessment of effect upon weight gains could be objective.

Italic numbers in parentheses refer to Literature Cited, p. 26.

J. S. Palmer is at the Toxicological Investigations Laboratory, Kerrville, Tex. 78028, and R. D. Radeleff is at the Southwestern Veterinary Toxicology and Live-stock Insects Research Laboratory, College Station, Tex.

a Acknowledgment is made to personnel of the Crops Research and the Pesticides Regulation Divisions for their revisions and comments.

Also McNamara, B. P. Summary of toxicity data on phenoxyacetates. 1967. [Personal correspondence.]

Weight gains or losses in chickens cannot be directly compared with the same changes in cattle and sheep because the latter were more mature and were being fed to maintain weight or to gain slightly. Chickens were being fed to gain approximately 40 percent of their initial weight in the test period.

INTERPRETATION OF HAZARD

To relate the toxic desages found for cattle, sheep, and chickens to the application rates recommended for each herbicide (17, 18, 19), we calculated the probable amounts that could be consumed daily from recently sprayed fields or pastures. In these calculations, we considered neither the influence of environmental factors such as soil type, temperature, and rainfall, nor the decomposition rates of the herbicides being

studied (δ)

"The U.S.D.A. Summary of Registered Agricultural Pesticide Chemical Uses" was utilized for the application rates (17). An arbitrary, although realistic, yield of 0.1 pound of air-dry forage per square foot of area was selected, which is equivalent to approximately 2 tons per acre. This would represent a high-quality, improved pasture. The reader must, of course, make adjustments for local conditions. A sparse cover of vegetation would allow more of the herbicide to reach the ground and be unavailable to animals, whereas a more lush vegetative cover would tend to hold more of the material available. In the latter case, however, less of the total forage of the area would be consumed in any one day.

Further assumptions were: (I) that an animal would consume, as forage, 3 percent of its body weight each day; and (2) that all the chemical formulation applied would adhere to the vegetation. Although this latter is never actually the case, this assumption gives the maximum exposure

to be expected.

An application of I pound of chemical to 1 acre of land provides 10.4 milligrams for each square foot. We may simplify the whole calculation to a single statement that I pound actual of herbicide per acre provides a 7-milligram per kilogram (mg./ kg.) dosage to the animal under the conditions here assumed to exist. Each 2.2 pounds of animal weight equals 1 kilogram or 1,000 grams. In turn, 1 pound equals 454 grams. The equivalent of 1,000 mg./kg. is 454 milligram per pound (mg./lb.).

EXPERIMENTAL ANIMALS

The cattle and sheep treated were obtained directly from local farmers and ranchers or through local auctions and were commercial stock. Cattle were of mixed breeding and sex, approximately 9 to 16 months old, and were classified as yearlings. Sheep were ewes and wethers of predominantly Delaine breeding. The cattle and sheep were maintained in open pens and were provided with rations of grain concentrates and hay. Mineral supplement and water were allowed as free choices.

The chickens were White Leghorns purchased from a commercial hatchery as day-old cockerels. One trial, however, included 31 chickens of Cornish-White Rock breeding. All chickens were held in brooder or grower pens for 6 weeks, first on a commercial starting feed and later on a growing mash; then the trials were started. Before each experiment was started, each of the birds was weighed and legbanded, then placed with 4 others in isolated cages. All 5 were treated at the same dosage level of the organic herbicide. An additional 1 or 2 birds in each cage, also legbanded and weighed, served as controls.

EXPERIMENTAL MATERIALS AND DOSAGES

The various organic herbicides utilized were usually commercially available formulations; however, exceptions were the use of: 2,4,5-T equivalent—pure acid or propylene glycol butyl ether ester—in 2 sheep in which residual effects were also studied (table 3); technical diphenamid in all trials (table 6); and, because of the unavailability of the commercial formulation, technical polychlorobicyclopentadiene isomers in a minority of cattle, sheep, and chickens more recently treated (table 25). Dosages were calculated on a mg./kg. basis for the active ingredient(s) of each chemical formulation to the body weight of the test animals. They were periodically recalculated in chronic studies as the weight of the animals changed.

Cattle and sheep were dosed with water-diluted formulation by use of a syringe, followed by a water rinse, or by use of a balling gun with the formulations in appropriate-sized gelatin capsules.

formulations in appropriate-sized gelatin capsules.

Chickens were dosed by water-diluted formulation by use of pipettes or by formulations in gelatin capsules. The treated chickens and the controls in

each cage were handled similarly; the controls were given untreated water or empty capsules.

The method used to select the number of doses administered and the dosage rates for each test animal involved several variables. The initial dosage rate was usually based on trial and error. When a toxic dosage was found, additional dosages above and below this rate were applied to other cattle, sheep, and chickens. Where a step-bystep increase of dosages indicated increased toxicity, repetition of individual dosages was not considered essential.

After each trial was completed, usually lasting 10 days or less, each surviving animal was weighed; the weight change was expressed as the percentage of initial, or preexposure, weight. Animals that died or were sacrificed during the course of the dosing or afterwards were examined by necropsy. Those sacrificed were moribund and death was considered imminent. Specimens for histopathological examination were collected for later atudy.

RESULTS

Chlorophenoxy Compounds

2,4-Dichlorophenoxyacetic acid (2,4-D), alkanolamine salts (of the ethanol and isopropanol series)

Cattle and sheep were dosed by drench in all cases except one sheep, which was dosed by capsule (table 1). This sheep lost less weight comparatively (4 percent) at 250 mg./kg. after 10 doses than another (10 percent) at the same dosage by drench. Cattle were poisoned also at the 250 mg./kg., but after only 1. A sheep on a chronic study at 100 mg./kg. lost significant weight after 8. This loss was subsequently regained with no other apparent toxic effects in the following 473 doses. Chickens dosed at 250 or 500 mg./kg. 10 times by capsule had a significant reduced weight gain when compared with the controls.

All five cattle were treated on a 5-day per week regimen to arrive at the total doses received (6). The yearling dosed at 100 mg./kg. 86 times became tympanitic (distended with gas) and was

medicated for this condition. After 2 additional doses, this yearling was removed from the experiment because of rumen atony (lack of normal tone), and an uneventful recovery followed. One yearling treated at 250 mg./kg. one time developed dyspnea (difficult breathing) and other signs of respiratory difficulty with partial anorexia associated with swellings in the mandibular region. As a result of a later experiment discussed on page 5 of this report, we decided that the difficulty was probably caused by the irritation of the diluted formulation on the mucous membrane of the pharynx.

Signs of poisoning in cattle were anorexia, ataxia (failure of muscular coordination), and, in cases of prolonged exposure, ulceration of the oral mucous membranes. Signs in sheep were depression and weight loss.

At necropsy on 2 cattle and 1 sheep, lesions in the animals varied. Hemorrhages on the surface of the epicardium with excessive quantity of peri-

TABLE 1.—Results of multiple dosing of cattle, sheep, and chickens with 2,4-dichlorophenoxyacetic acid (2,4-D), alkanolamine salts (of the ethanol and isopropanol series)1

and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks ²
	Num- ber		
Cattle:	A¢1		
50	112	Drench	NIE.
100			Poisoned after 86 and survived.
200			Poisoned after 25 and died.3
250			Poisoned and survived.
250	20	do	Poisoned after 15 and died.
Sheep:			
100	481	do	11-percent weight loss after 8, otherwise NIE.
250	10	do	10-percent weight loss.
250	10	Causule	4-percent weight loss.
500			Poisoned and died.
Chickens:			
100	10	Pipette	36-percent weight gain.
250			31-percent weight gain.
500		do	Do.
Controls			38-percent weight gain.

 ^{2,4-}Dow®, 65 percent emulsifiable concentrate, Dow
 Chemical Co., Midland, Mich.
 NIE indicates no ill effects apparent.

cardial fluid were observed. Liver and kidneys were congested and friable and there was rumen stasis characterized by bright, undigested feed. The lungs were generally engorged with blood.

Application rates for 2,4-D salts range from 0.07 (1.12 ounces) to 40 pounds per land acre and to 87 pounds actual per water acre on submerged weeds. Application rates above 30 pounds actual per acre are hazardous for cattle, sheep, and chickens.

2.4-Dichlorophenoxyacetic acid (2,4-D), propylene glycol butyl ether ester

Either capsules or drenches were used for dosing cattle and sheep, capsules for chickens (table 2). Doses by capsule given a few animals resulted in an apparently greater toxicity than by drench. A yearling and a sheep were poisoned at 250 mg./kg. by drench after 3 and 2 doses, respectively, with death resulting in only the sheep after 9. Doses by capsule caused poisoning and death at this dosage after 7 in cattle and 5 in sheep. One sheep had no ill effects at 100 mg/kg. by drench after 481. Chickens showed an average reduction in weight gain at 100 mg./kg. after 10.

Signs of toxicity in cattle and sheep were increasing depression and anorexia, followed by prostration until moribund. At necropsy the liver was soft and friable and the gall bladder was distended with bile. The kidneys were congested and friable and there were petechiae on the surface of the epicardium and large vessels. The lymph nodes in some animals were enlarged and hyperemic.

Application rates for 2,4-D esters range from 0.07 to 40 pounds actual per land acre and to 87 pounds actual per water acre on submerged weeds. Application rates above 30 pounds actual per acre are hazardous for cattle, sheep, and chickens.

TABLE 2.—Results of multiple oral dosing of cattle, sheep, and chickens with 2,4-dichlorophenoxyacetic acid (2,4-D), propylene glycol butyl ether ester 1

Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks 2
Cattle:	Num-	v " i 	<u></u>

	Num-		
Cattle:	ber		
100	10	Drench	NIE.
250	10	do	Poisoned after 3 and aurvived, 12-percent
250	7	Capsule	weight loss. Poisoned and died.
Sheep:			
100	481	Drench	NIE.
250	9	do	Poisoned after 2 and died.
250	5	Capsule	Poisoned and died.
Chickens:			
50	10	do	49-percent weight gain.
100	10	do	19-percent weight gain.
250	10	do	19-percent weight gain. Poisoned after 4, 13- percent weight loss.
Controls			percent weight loss. 41-percent weight gain.

¹ Esteron 99%, 38 percent emulsifiable concentrate, Dow Chomical Co., Midland, Mich.

^{*} Affected by chemical reaction of the formulation on the pharyngeal mucosa, resulting in mandibular area enlargement accompanied by partial anorexia and dyspnes.

4 Average results of 5 treated chickens.

NIE indicates no ill effects apparent. * Average results of 5 treated chickens.

2.4.5-Trichlorophenoxyacetic acid (2.4.5-T), propylene glycol butyl ether ester

Either capsules or drenches were used for dosing cattle and sheep, capsules for chickens (table 3). One yearling showed signs of poisoning at 250 mg./kg. after 4 doses by drench, whereas another at the same dosage had no ill effect after 10 by capsule. Although one sheep was poisoned at 100 mg./kg. by drench after 3 and died after 8, another had an initial weight loss at the same dosage by capsule but tolerated 367 before being poisoned and fatally affected. Chickens showed a significant

decrease in average weight gain at 250 mg./kg.
Signs of poisoning in cattle and sheep were
increasing anorexia and progressive weight loss, leading to weakness and prostration before death. At necropsy, the liver was swollen and often friable and the kidneys were congested. There were hemorrhages in the heart musculature. A general congestion of the visceral blood vessels was associated with rumen stasis characterized by bright undigested feed. Hyperemia and anlargement of the lymph nodes were noted in some animals. The two dead chickens showed congestion of the kidneys and intestinal mucosa.

Application rates for 2,4,5-T range from 0.5 to 4.5 pounds actual per acre. Such rates of this ester are not hazardous for cattle, sheep, or chickens.

2-(2,4,5-Trichlorophenoxy) propionic acid (silvex), propylene glycol butyl ether ester

Cattle were dosed either orally by drench or capsule or by rumen fistula, sheep were dosed by either drench or capsule, and chickens were dosed by pipette (table 4). Whereas 1 yearling showed no ill effects at 100 mg./kg. after 10 doses by capsule, another was poisoned after 19 by drench and died after 29. A sheep dosed at 50 mg./kg. by drench was poisoned and survived 10, but another dosed at 100 mg./kg. by capsule was poisoned after 9 and died after 11. In one study to determine the effect of silvex on various enzyme systems, 6 sheep were dosed at 100 mg./kg. by drench for 21 days, with no ill effects. The dosage was increased 50 percent to 150 mg./kg. for 10 days; 1 sheep died after 29, another after 31 (21). Chickens had a significant weight reduction after 10 at 100 mg./kg.

Signs of poisoning were only anorexia and weight loss in one sheep at 50 mg./kg., or anorexia with an increasing depressive state until recumbency in other poisoned sheep and cattle. Then a moribund condition developed, fellowed by death.

TABLE 3.—Results of multiple oral dosing of cattle, sheep, and chickens with 2,4,5-trichlorophenoxyacctic acid (2,4,5-T), propylens glycol butyl other ester 1

Animal and desage received (mg./kg.)	Doses	Means of dosing	Results and remarks
	Num-		
Cattle:	ber		
100	10	Drench	NIE.
250	7	do	Poisoned after 4 and died 2 days after last dose.
250	10	Capsule	
Sheep:			
50	10	Drench	Do.
100	869	Capsule	9-percent weight loss after 8, poisoned
100	8	Drench	after 367 and died. Poisoned after 3 and died.
250	7	do	Do.
250	6	Capsule	Poleoned after 4 and died.
250	4	do	Poisoned and sacrificed.
200 20041	-	40	270.
Chickens: *			
100	10	do	36-percent weight gain.
250	îň	do	18-percent weight gain.
600	ĩŏ	do	2 died after 5 and 9, 12-percent weight loss in survivors.
Controls			82-percent weight gain.

^{*}Esteron 245 O.S. ©, 65.3 percent emulsifiable concentrate, Dow Chemical Co., Midland, Mich.

*NIE indicates no ill effects apparent.

2,4,5-T equivalent—pure acid.
2,4,5-T equivalent—pure ester.
4 Average results of 5 treated chickens.

At necropsy, the liver was enlarged and friable and the kidneys were congested. There was an engorged rumen indicating stasis. A small abacess was found in the parotid lymph node in one yearling that developed a swelling in this region related to the chemical reaction associated with drenching. Other lymph nodes of the body were often enlarged and hemorrhagic.

Silvex administered in drench to cattle often caused an enlargement or swelling in the parotid and, in one instance, in the mandibular region. To determine whether this was the result of a systemic effect of silvex, we made a special study in which 6 yearlings were desed at 50 mg./kg. (See table 4-

Amimal

TABLE 4.—Results of multiple oral dosing of cattle, sheep, and chickens with 8-(2,4,5-trichloro-phenoxy) propionic acid (silvez), propylene glycol butyl ether ester 1

and dosage Doses Means of Results and remarks ?

received (mg./kg.)		dosing	
	Num		
Cattle:	ber		
25	73	Drench	NIE, except for irritation effect after 20.3
50	78	do	
50	19	do	
50	27	do	
50	90	do	NIE
50	56		Poisoned and died.
50	90	do	NIE.
50	90	do	Do.
100	29		Poisoned after 19 and died after 29; irritation effect after 4.9
100	10	Capeule	
Sheep:			
25	10	Drench	Do
50	10	do	Poisoned and survived, 15-percent weight loss,
100	11	Capsule	
100-150 4	20	<u>-</u>	dled.
100-150		do	died.
100-150	31	do	NIE.
100-150 4		do	Do.
100-150 4		do	
100-150 4	81	do	Do.
250	5	do	Poisoned and died.
Chickens:			
50	10	Pipetta	. 48-percent weight gain.
100	10	do	25-percent weight gain. 28-percent weight loss.
250	10	do	28-percent weight loss.
Controls		••••••	41-percent weight gain.

Kuron®, 46.5 percent emulsifiable concentrate, Dew Chemical Co., Midland, Mich.
 NIE indicates no ill effects apparent.

all yearlings dosed at 50 mg./kg. except the first listed.) Three of these were dosed by drench in the customary manner; 1 developed this parotid swelling after 19 and another after 27 doses. Rumen fistulas were prepared in the other 3 yearlings and silvex was introduced directly into the rumen. One of these yearlings died after 56, possibly influenced to some degree by a mild localized infection that developed in the region of the fistula. Three yearlings in this special study showed no ill effects from the dosage. We concluded that the enlargements were caused by the chemical reaction of the diluted herbicide formulation on the mucous membrane and underlying tissue of the pharyngeal region (9).

TABLE 5.—Results of multiple oral dosing of cattle, sheep, and chickens with 2-methyl-4-chlorophen-oxyacetic acid (MCPA), alkanolamine salts (of the ethanol and reopropanol series) 1

and dosage received (mg./kg.)	Dones	Means of doding	Results and remarks *
	Num		
Cattle:	ber		
260		Drenoh	
500	10	do	Poisoned after 8 and survived.
Bheep:			
50	10		NIE.
100	10	Drench	Poisoned after 4 and survived, 17-percent weight loss.
100	38 3	Capsule	Poisoned after 307 and
250	10	do	Poisoned and survived, 6-percent weight
250	. 10	Drench	Poisoned after 6 and survived, 6-percent weight loss.
500	10	do	NIE.
Chickens:			
100	. 10	Capsule	42-percent weight gain.
250	10.	do	31-percent weight
800	10	do	I died after 1 dose; 20-percent weight loss in survivors.
Controls	*****		41-percent weight

MCP Amine®, 69.1 percent emulsifiable concentrate, Dow Chemical Co., Midland, Mich.
NIE indicates no ill effects apparent.

Affected by chemical reaction of the formulation on the pharyngeal nucces, resulting in parotid or mandibular area enlargement accompanied by partial anorexia and dyspnes.

Mild localised infection developed in region of fistula.

^{*} Mild localised intection developed in region of natura.

* 21 does at 100 mg./kg.; then 10 or fewer at 150 mg./kg.

*Average results of 5 treated chickens.

[•] Average results of 5 treated chickens.

Application rates for 2,4,5-TP (silvex) range from 0.13 (61 grams) to 8 pounds actual per acre, usually not more than 5. The higher rates would be no hazard to the three test species.

2-Methyl-4-chlorophenoxyacetic acid (MCPA), alkanolamine salts (of the ethanol and isopropanol series)

Cattle were dosed by drench, sheep by either drench or capsule, chickens by capsule (table 5). A yearling was poisoned at 500 mg./kg. after 8 doses, but the regimen was continued for a total of 10 with no further ill effects observed. Paradoxically, 2 sheep dosed at 250 mg./kg. were poisoned after 6 and 10, with a significant weight loss; whereas another showed no ill effects at 500 mg./kg. after 10. This difference in reaction could very likely be the result of individual tolerance. In a long-term trial I sheep dosed by capsule at

100 mg./kg., was poisoned after 307 and died after 383. In contrast, another sheep was poisoned at 100 mg./kg. after 4 by drench; it survived 10 but had significant weight loss. Chickens had significant reduced weights at 250 mg./kg.

Signs of poisoning of the sheep that died were anorexia, ataxia, muscular spasms, and dyspnea. The poisoned yearling became tympanitic and showed partial anorexia.

At necropsy on the sheep, the spleen was enlarged, lungs were congested, and kidneys were swollen and congested. There were hemorrhages on the surface of the epicardium and the liver was friable and light brown. The meningeal vessels were congested. At necropsy on the one chicken, the spleen was enlarged and there was redness of the intestinal mucosa.

Application rates for MCPA range from 0.13 to 2:10 pounds actual per acre, which would be no hazard to the three test species.

Amide Compounds

Animai

N,N-Dimethyl-2,2-diphenylacetamide (diphenamid)

All test animals were dosed by capsule (table 6). One yearling was poisoned at 250 mg./kg. after 3 doses and a sheep after 7. Chickens had a significant reduced weight gain at the same desage.

The poisoned yearling developed an uncoordinated gait after 3 doses and further dosage was discontinued. The poisoned sheep developed lameness and muscular spasms after the 7th and final dose without forewarning signs. Anorexia, diarrhea, and prostration progressed until a moribund condition brought about the decision to sacrifice the sheep.

At necropsy, the ears of the sheep were con-gested. The subcutaneous tissues and areas delineating the different muscles were filled with yellow gelatinous material. There was rumen stasis characterized by bright undigested feed the liver was enlarged and light brown, the kidneys were congested, the bladder contained brownish-tinged urine, and the adrenals were enlarged.

Application rates for diphenamid for weed control in various crops range from 3 to 6 pounds actual per acre. These rates would not be hazardous for the three test species.

2-Chloro-N.N-diallylacetamide (CDAA)

Cattle and sheep were dosed by either drench or capsule, chickens by capsule (table 7). The toxic effect is evident at 25 mg./kg. in the 3 test species. One yearling was poisoned after I dose by drench, another after 2 by capsule, but each survived 10, with weight loss. One sheep was poisoned after 4 by capsule but survived 10, with weight loss,

TABLE 6.—Results of multiple oral dosing of cattle, sheep, and chickens with N, N-dimethyl-2,2diphenylacetamide (diphenamid) 1

and dosage Doses Means of Results and remarks 2

received (mg./kg.)		doeing	
	Num-		
Cattle:	ber		
100	10	Capsule	NIE.
250		do	Poisoned and survived, 8-percent weight loss.
Bheep:			- F
100	10	do	NIE.
250	7	do	Poisoned and sacrificed.
Chickens:			
100	10	do	44-percent weight gain.
250	10	do	26-percent weight gain.
500	10	do	8-percent weight gain.
Controls			36-percent weight gain.
COLUMNIA			an borooms weren forms

¹ Enide[®], 100 percent technical powder, Upjohn Co., Kalamasoo, Mich.

NIE indicates no ill effects apparent. Average results of 5 treated chickens.

Animal

and dosage

Penaltrad

TABLE 7.—Results of multiple oral doring of cattle, sheep, and chickens with 2-chloro-N.Ndiallylacetamide (CDAA)

Results and remarks?

Doses Means of

dosine

(mg./kg.)	doemg	
	Num-	
Cattle:	<i>ber</i>	
10	10 Drench	
25	10do	Poisoned after 1 and survived, 6-percent weight loss.
25	10 Capsule	Poisoned after 2 and survived, 7-percent weight loss.
50	1 Drench	Poisoned and died.
Sheep: 25	10do	377 E*
25		Poisoned after 4 and
20		survived, 9-percent weight loss.
50	1 Dreneb	Poisoned and died.
50	2do	Do.
50	5 Capsule	died.
100	1do	Poisoned and died.
100	1 Drench	Do.
250	1do	Do.
Chickens; ²	•	
10	10 Capsulo	46-percent weight gain.
25	10do	20-percent weight gain.
25	10do	39-percent weight gain.
50	10do	2 poisoned and died after 9, no weight change in survivors.
100	6do	All died after 1 to 6.
Controls	************	

Randox®, 47.1 percent emulsifiable concentrate, Monsanto Chemical Co., St. Louis, Mo.

whereas another showed no ill effect at the same dosage by drench. Chickens had significant reduced weight gains after 10.

Anorexia was the most prominent sign of poisoning. As additional doses were given to the cattle and sheep, there was increased salivation, and depression and prostration usually followed. Chickens were similarly affected by anorexia.

At necropsy on the yearling and the 6 sheep, the mucosa of the abomasum and small intestine was hemorrhagic. There was usually congestion of the lungs and the respiratory tract mucosa and the

kidneys were often enlarged and congested. Petechiae were noted on the surface of the bladder mucosa. Congestion of the intestinal mucosa was the most prominent lesion in chickens.

Application rates for CDAA range from 4 to 10 pounds actual per acre. These rates would be highly

hazardous for all three test species.

2-Chloro-N,N-diallylacetamide (CDAA) and Trichlorobenzyl chloride (TCBC)

Dosing was done by drench to cattle, by capsule to chickens, and by either method to sheep. One yearling was poisoned at 50 mg./kg. after 2 doses

TABLE 8.—Results of multiple oral doeing eattle, sheep, and chickens with 2-chloro-N,Ndiallylacetamide (CDAA) and trichlorobenzyl chloride (TCBO)

and dosage received (mg./kg.)	Doses	Means of doeing	Results and remarks ¹
Cattle:	Number		
25	, 10	Dreneb	NIE.
50	. 10	do	Poisoned after 2 and survived, 11-percent weight loss.
100	. 1	do	Poisoned and died.
Sheep:			
25 4	. 10	do	
25 • 25 •	. 10	Capsule	Poisoned after 5 and survived, 11-percent weight loss.
50	_		Poisoned after 2 and survived, 15-percent weight loss.
50	. 10	Capsule	.Do.
100		Drench	Poisoned after 1 and died.
100	. 10	Capsule	Poisoned after 2 and survived, 16-percent weight loss.
250	. 1	Drench	Poisoned and died.
Chickens: 4			
37.5	10	Capsule	43-percent weight gain.
75	. 10	do	37-percent weight gain.
100	. 10	do	24-percent weight gain.
100	. 10	do	7-percent weight loss.
250	. 8	do	All died after I to 8.
Controls			40-percent weight gain.

Randox TS, 85.7 percent emulsifiable concentrate, Monsanto Chemical Co., St. Louis, Mo. (28.1 percent CDAA and 57.6 percent TCBC).

NIE indicates no ill effects apparent. Average results of 5 treated chickens.

NIE indicates no ill effects apparent.
25 mg./kg. is equivalent to 8.15 mg./kg. CDAA and
16.85 mg./kg. TCBC. Average results of 5 treated chickens.

but survived 10, with weight loss (table 8). One sheep was poisoned at 25 mg./kg. after 5 by capsule but survived 10 by drench, with weight loss; another sheep showed no ill effects at the same

dosage administered by drench.

The only signs of toxicity were anorexia with loss of weight in poisoned cattle and sheep and reduced average gain or loss of weight in chickens. At necropsy on cattle and sheep, the abomasal and intestinal mucosa was hemorrhagic and the liver and kidneys were congested. In one animal,

the lungs were engarged with blood and there was blood-tinged froth in the respiratory tract. There were hemorrhages throughout the heart of the yearling that died after 1 dose at 100 mg./kg. At necropsy on chickens, kidneys were congested as was the intestinal mucosa.

Application rates for this combination of CDAA and TCBC range from 3.5 to 7 pounds actual per acre of corn. The 3.5-pound rate would be hazardous for sheep, and the 7-pound rate would be hazardous for cattle but not for chickens.

Phenyl Urea Compounds

3-(p-Chlorophenyl)-1,1-dimethylurea (monuron)

Cattle and sheep were dosed by either drench or capsule, chickens by capsule (table 9). One yearling was poisoned but survived 1 dose at 500 mg./kg., which resulted in weight loss during the following 10-day period. Diarrhea resulted in cattle after 1 to 5 doses at 50, 100, and 250 mg./kg. by drench; however, diarrhea was not severe enough to result in significant weight loss or other related signs of poisoning. In contrast, another yearling dosed at 50 mg./kg. by capsule had no diarrhea or other ill effects after 10.

Sheep were poisoned and survived 100-mg./kg. dosages, with significant weight losses; however, the sheep dosed by capsule at this rate tolerated the formulation better than the 2 sheep dosed by drench. Although 2 sheep showed signs of poisoning after 4, only the sheep dosed by drench died after the additional doses. Chickens dosed at 25 mg./kg. had only reduced weight gains; chickens at 100 mg./kg. either died or lost weight after 8 to 10 doses.

The yearling poisoned at 500 mg./kg. became prostrate and was treated for tympanites. After 2 days it was ambulatory with an uncoordinated gait. One sheep dosed at 100 mg./kg. was poisoned and showed signs of excitability and an uncoordinated gait, which preceded prostration and severe weight loss; the 2d sheep dosed at this rate showed depression and anorexia; the 3d showed loss of equilibrium, which preceded prostration. Torticollis (twisted neck and unnatural position of the head) developed in this 3d sheep 2 days before death, which occurred 5 days after the 9th and last dose.

At necropsy, the lungs of the sheep were engorged with blood with accompanying congestion of the respiratory mucosa. The liver and kidneys were congested and enlarged. The meningeal vessels of the brain were engorged. Chickens

TABLE 9.—Results of multiple oral dosing of oattle, sheep, and chickens with S-(p-chlorophenyl)-1,1-dimethylurea (monuron)¹

Animal and dosage received (mg./kg.)	Doses	Means of doeing	Results and remarks
C I. 413	Num-		
Cattle:	ber		****
25	10		NIE.
50	10		Do. Do.
50	10		
100		do	Do.
250	10	do	_ Do.
500	1	do	Poisoned and survived, 11-percent weight loss.
Sheep:			
25		do	
50		do	Do.
100	4	do	Poisoned and survived, 19-percent weight loss.
100	10	Capsule	Poisoned after 4 and survived, 15-percent weight loss.
100	9	Drench	Poisoned after 4 and died, 20-percent weight loss.
250	2	do	Poisoned and sacrificed
Chickens:			
10	10	Capsule	54-percent weight gain.
25	10	do	33-percent weight gain.
50	10	do	20-percent weight gain.
100	10	do	1 poisoned and died after 8, 1-percent weight loss in survivors.
250 Controls			All died after 6 to 9. 47-percent weight gain.

¹ Telvar[®], 80 percent wettable powder, E. I. DuPont de Nemours and Co., Wibnington, Del.

² NIE indicates no ill effects apparent; however, see

NIE indicates no ill effects apparent; however, se narrative for results to cattle dosed by drench.
 Average results of 5 treated chickens.

and dosage

generally showed congestion of the intestinal

mucosa, with atrophied spleens.

Application rates for monuron generally range from 1.2 to 6.4 pounds actual per acre, although it may be used in water ditches at a rate equivalent to 80 pounds actual per acre. Rates in excess of 3 pounds per acre would be hazardous for chickens. A 6.4-pound rate would not be hazardous for cattle or sheep but the 80-pound rate would be highly hazardous for all three test species.

3-(3,4-Dichlorophenyl)-1-methoxy-1methylurea (linuron)

Sheep were dosed by drench, chickens by capsule, cattle by either method. One yearling was poisoned at 50 mg./kg. by capsule after 10 doses, with a significant weight loss, whereas another yearling had no ill effects after 10 by drench (table 10). Sheep were poisoned at 50 mg./kg. after 1 or 8 with weight loss Chickens mg./kg. after 1 or 8, with weight loss. Chickens dosed at 10 mg./kg. showed a significant reduced

TABLE 10.—Results of multiple oral dosing of cattle, sheep, and chickens with 3-(3,4-dichlorophenyl)-1-methoxy-1-methylurea (linuron) 1

Doses Means of Results and remarks 1

received (mg./kg.)	170560	doeing	resource which remarks.
	Num-		
Cattle:	ber		
25	10	Dronch	NIE.
50	10	do	Do.
50	10	Capsule	Poisoned and survived, 6-percent weight loss.
100	′ 6	Dreuch	Poisoned and died 2 days after last dose.
Sheep:			•
25	10	do	NIE.
50			Poisoned after 1 and survived, 8-percent weight loss.
50	10	do	Poisoned after 8 and survived, 6-percent weight loss.
100	4	do	Polsoned and died.
250		do	
Chickens: 1			
10	10	Capsule	39-percent weight gain.
25	7.2		16-percent weight gain.
50			20-percent weight gain.
100			4 died after 8 or 9, 20-percent weight loss in survivor.
Controls		_	46-percent weight gain.

Lorox®, 50 percent wettable powder, E. I. DuPont de Namours and Co., Wilmington, Del. • NIE indicates no Ill effects apparent.

weight gain. No lesser desage was tried. Four of 5 chickens in the 100-mg./kg. study group died after 8 or 9.

The most general signs of poisoning were ancrexia and depression. Cattle had increased salivation and showed a marked weakness. All 4 poisoned sheep had an uncoordinated gait, and hematuria (discharge of blood in the urine) was observed in 2. Of these 2, one was poisoned at 50 mg./kg. after 8 doses and the other at 100

mg./kg. alter 4 that preceded death.

At necropsy on cattle, there were congestion of the spleen and petechiae on the surface of the epicardium. In sheep there were hemorrhages on the muscle fasciae of the hindquarters and edema between the muscles. There were petechiae on the surface of the epicardium. The pericardial sac contained blood-tinged fluid. The lungs were engorged with blood and the respiratory mucosa was congested. There was marked congestion of the kidneys and petechiae were found on the surface of the bladder mucosa of both sheep. The liver was friable and swollen and there was congestion of the intestinal mucosa. At necropsy on chickens there were enlarged livers and congestion of the intestinal mucosa and kidneys.

The recommended application rates for linuron range from 1 to 3 pounds actual per acre either as a preemergent or a postemergent spray on crops. At these rates, there would be no hazard to cattle,

sheep, or chickens.

3-Phenyl-1,1-dimethylurea (fenuron)

Cattle and sheep were dosed by drench, chickens by capsule (table 11). One sheep dosed at 100 mg./kg. was the most susceptible of all the test animals and was poisoned after 8 doses. Both cattle and chickens had ill effects only at 500 mg./kg. Cattle were poisoned by 2, whereas chickens showed decreased weight gains after 10.

Signs of poisoning in cattle and sheep were anorexia, depression, and ataxia. In the one yearling poisoned at 500 mg./kg., tympanites occurred. At necropsy on the sheep, there were hemorrhages on the surface of the epicardium, the lungs and liver were congested, and the liver

was enlarged.

Fenuron is recommended for brush control in pastures and rangelands. As a spot treatment only, the application rate is 0.0156 pound (2 tablespoonfuls) of a 25-percent pelletized formula-tion per square foot. This rate could be a hazard to cattle, sheep, and chickens if a large area were to be treated.

3-(3,4-Dichlorophenyl)-1,1-dimethylurea (diuron)

Cattle and sheep were desed by either drench or capsule, chickens by capsule (table 12). One

^{*} Average results of 5 treated chickens.

TABLE 11.—Results of multiple oral dosing of cattle, sheep, and chickens with 8-phenyl-1,1-dimethylurea (fenuron) 1

Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks 3
	Num-		
Cattle:	ber	_	
250	10	Drench	NIE.
500	3	do	Poisoned after 2 and survived, 20-percent weight loss.
Sheep:			
50	10	Capsule	NIE
100	íŏ	Drench	Poisoned after 8, with 9-percent weight loss after 10, died 8 days
250	10	do	after last dose. [‡] Poisoned and survived, 13-percent weight loss.
500	5	do	Poisoned after 3 and sacrificed.
Chickens:			•
250	10	Canaule	47-percent weight gain.
500			38-percent weight gain.
Controls			

Dybar®, 25 percent pellets, E. I. DuPont de Nemours and Co., Wilmington, Del.
 NIE indicates no ili effects apparent.

yearling dosed at 100 mg./kg. by drench was poisoned; another dosed by capsule had no ill effects. One sheep dosed at 100 mg/kg. by drench was poisoned after 2 doses and had a significant weight loss during the next 8 days. Chickons dosed at 10 mg./kg. showed a significant decrease in weight gain. One chicken died at 100 mg./kg. after 10; the 4 surviving chickens showed weight loss

Signs of poisoning in the yearling and the sheep were anorexia, depression, dyspnea, and prostration. In the 2 poisoned sheep, an uncoordinated gait was observed. Both sheep recovered after treatment was discontinued. At

necropsy on chickens, there was congestion of the intestinal mucosa and an enlarged, congested liver.

Application rates for diuron most commonly range from 0.2 to 9.6 pounds actual per acre. Rates in excess of 1 pound per acre would be hazardous for chickens. The 9.6-pound rate would not be hazardous for cattle and sheep. A rate of 80 pounds actual per acre is used in irrigation ditches and would be highly hazardous for all three test species.

TABLE 12.—Results of multiple oral dosing of cattle, sheep, and chickens with 3-(3,4-dichlorophenyl)-1,1-dimethylurea (diuron)

Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks *
	Num-	······································	
Cattle:	ðer		****
50	10	Drench	NIE.
100			Poisoned and survived, 9-percent weight loss.
100	10	Capsule	NIE.
Sheep:			
25	10	Drench	Do.
δ0			
50		Drench	Do.
50		do	
100	10	4~	Poisoned and survived,
10011111	#	40	12-percent weight loss.
250	1	do	Poisoned and survived, 8-percent weight loss.
Chickens:			
5	10	Capsule	58-percent weight gain.
10	10	do	37-percent weight gain.
25	10	do	37-percent weight gain.
50	า๊ก	do	14-percent weight gain.
100	10	do	I poisoned and died after
	••		10, 17-percent weight loss in survivors.
250	G.	do	All died after 8 or 9.
Controle	•		50 normant waight sein
Controls			50-percent weight gain

¹ Karmer®, 80 percent wettable powder, E. I. DuPont de Nemours and Co., Wilmington, Del.

² NIE indicates no ill effects apparent.

Thiocarbamate Compounds

S-2.3-Dichloroallyl diisopropylthiocarbamate (diallate)

Cattle and sheep were dosed by either capsule or drench, chickens by capsule (table 13). The toxic dosage was 25 mg./kg. for cattle and sheep.

One yearling was poisoned after 1 dose by drench; another was poisoned after 5 by capsule. In comparison, 1 sheep dosed at this rate was poisoned after 5 doses by drench; another was poisoned after 1 by capsule. One sheep dosed at 25 mg./kg. had no ill effect from 10 by drench;

Complications of pneumonia.

Average results of 5 treated chickens.

Average results of 5 treated chickens.

whereas the other 2 sheep dosed at this rate, 1 by drench and 1 by capsule, were poisoned. This, presumably, would be due to individual tolerance. Chickens dosed at 125 mg./kg. or higher had aignificant reduction in weight gain.

Signs of poisoning in cattle and sheep were muscular spasms, ataxia, and depression, generally followed by prostration. A delayed alopecic effect (loss of hair or wool—either partial or complete) was seen in the more chronically affected animals. at times 60 days after dosing was discontinued. The most prominent sign of poisoning in chickens was anorexia, resulting in decreasing weight gains, weight loss, or death.

At necropsy on cattle and sheep, gross lesions consisted of a congested and friable liver, congestion of the kidneys with petechine on the surface of the bladder mucosa, and, in one sheep, blood-engorged lungs and hemorrhagic intestinal mucosa. At necropsy on chickens, the kidneys were congested, the liver was swollen, and the

intestinal mucosa was hemorrhagic.

Application rates for diallate range from 1.25 to 2 pounds actual per acre, which would not be hazardous for the three test species. However, only a modest increase to 3 pounds actual per acre would be hazardous for cattle and sheep.

S-2.3.3-Trichloroallyl diisopropylthiocarbamate (triallate)

Cattle and sheep were dosed by either drench or capsule, chickens by capsule (table 14). A yearling was poisoned at 25 mg./kg. after 5 doses by drench but survived 5 additional doses, with significant weight loss. One sheep was poisoned at 50 mg./kg. after 2 by capsule, but survived 8 additional doses, with severe weight loss. In contrast, 2 sheep dosed at 50 mg./kg. by drench had no ill effects after 10 and 60 consecutive doses. Chickens dosed at 175 mg./kg. had a significant reduction in average weight gains.

Signs of poisoning in cattle and sheep were anorexia and subsequent large percentages of weight loss. A long recuperative period usually followed toxicosis. Muscular spasms, diarrhea, and ataxia were seen in a poisoned sheep that recovered. At dosages over 100 mg./kg., chickens had anorexia, resulting in reduced weight gains. All chickens were fatally affected at 250 mg./kg., showing congestion of the liver and kidneys and hemorrhagic intestinal mucosa at necropsy.

Application rates for triallate range from 1 to 1.25 pounds actual per acre. At such rates it would not be a hazard to the three test species.

Table 13.—Results of multiple oral dosing of cattle, sheep, and chickens with S-2,3-dichloroallyl discopropulthiocarbamate (diallate)1

Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks ?	Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks *
Cattle:	Num- ber	· · · · · · · · · · · · · · · · · · ·			Num-		
10		Drench	NITE	Sheep-Con:		Committee	Defeated often 4 and
25	2	do	Poisoned after 1 and survived, 5-percent weight loss.	25	19	Capaule	Poisoned after 1 and survived, 20-percent weight loss, partial alopecia.
25	10	Capsule	Poisoned after 5 and survived, alopecia of	50			Poisoned after 3 and sacrificed.
50	8	do	the tail 60 days after last dose, 17-percent weight loss. Poisoned after 7 and survived, alopeda of	100	3	do	Poisoned and survived, complete alopecia within 21 days after last dose, 14-percent weight loss.
			the tail 60 days after last dose, 11-percent weight loss.	250 Chickens:	2	do	Poisoned after 1 and died.
50	6	Drench	Poisoned after 3 and survived, 20-percent	50 100	10 10	Capsule	64-percent weight gain. 51-percent weight gain.
			weight loss.	125	10	do	36-percent weight gain.
Bheep:	••		*****	150	10	do	22-percent weight gain.
25 25	10	do	NIE. Poisoned after 5 and	175 250	10	do	9-percent weight loss. All died after 7 to 10.
20	10		survived, 7-percent weight loss.				55-percent weight gain.

¹ Avadex®, 45.7 percent emulsifiable concentrate, Monsanto Chemical Co., St. Louis, Mo.

^{*} NIE indicates no apparent ill effects.

Average results of 5 treated chickens.

Table 14.—Results of multiple oral dosing of cattle, sheep, and chickens with S-2,3,3-trichloroallyl diisopropylthiocarbamate (triallate)¹

Animal and dosage received (mg./kg.)	Doccs	Means of dosing	Results and remarks ³	Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks?
Cattle:	Num-			Sheep—Con:	Num- ber		
10 25	ber 10 10		NIE. Poisoned after 5 and survived, 5-percent	100	10	Drench	Poisoned after 1 and survived, 21-percent weight loss.
50	10	Capsule	weight loss, long recuperative period. Poisoned after 4 and	100	4.	do	Poisoned and survived, 20-percent weight loss.
50	10	•	survived, 12-percent weight loss. Poisoned after 3 and	250	2	do	Poisoned and survived, 11-percent weight loss.
		37101104424	survived, 16-percent weight loss.	Chickens:			
Sheep: 25 50 50	10	do do Capsule		50	10 10 10 10	do do	59-percent weight gain. 55-percent weight gain. 10-percent weight gain. 17-percent weight gain. All died after 7 to 10. 41-percent weight gain.

¹ Avadex BW6, 46.3 percent emulsifiable concentrate, Monsanto Chemical Co., St. Louis, Mo.

Triazine Compounds

2-Chloro-4-ethylamino-6-isopropylamino-striazine (atrazine)

Cattle and sheep were dosed by either drench or capsule, chickens by capsule (table 15). The toxic dosage for cattle was 25 mg./kg. after 8 doses by drench and 2 by capsule. The toxic dosage for sheep was 5 mg./kg. No lesser dosage was tried. However, one sheep received 199 consecutive doses at 50 mg./kg. before it was poisoned and died. Chickens given 10 at 50 mg./kg. had a significant reduction in weight gains.

Signs of poisoning in cattle and sheep were muscular spasms of varying intensity in the hind quarters, a stilted gait and stance, and anorexia. In 2 sheep dosed at 100 mg./kg., when dosing was continued after moderate signs of poisoning developed, apparent improvement was observed until unexpected death occurred after 8 and 10 doses.

At necropsy, lesions of the yearling and the sheep varied somewhat. Petechiae on the surface of the epicardium and congestion of the kidneys, liver, and lungs were generally present. On occasion, the liver was light brown and friable. Some, but not all, of the dead animals had enlarged adrenals, as reported previously (10).

Application rates for atrazine range from 0.4 to 6.4 pounds actual per acre. Rates of less than 1 pound would be hazardous for sheep. Rates of 3 pounds actual per acre would be hazardous for cattle. The 6.4-pound rate would be hazardous for chickens.

2-Chloro-4,6-bis(ethylamino)-s-triazine (simazine)

Cattle were dosed by drench, chickens by capsule, and sheep by either drench or capsule (table 16). Two yearlings were poisoned at a dosage of 25 mg./kg. after 3 and 10 doses. One sheep was poisoned at 50 mg./kg. after 17 and died after 31, whereas another was poisoned after 10 and survived with an 18-percent weight loss. Chickens dosed at 50 mg./kg. 10 times showed a reduced weight gain.

Signs of poisoning in cattle and sheep were anerexia, depression, muscular spasms, and dyspnea of increasing intensity. Weakness and an uncoordinated gait were commonly observed in severely poisoned animals.

At necropsy, lesions in cattle and sheep generally were congestion of the lungs and kidneys;

NIE indicates no ill affects apparent.
 Average results of 5 treated chickens.

TABLE 15.—Results of multiple oral dosing of cattle, sheep, and chickens with 2-chloro-4-ethylamino-6-isopropylamino-s-triazine (atrazine)

Doses Means of Results and remarks and dosage dosing received (mg./kg.) Number Cattle: Drench... NIE. 10 10.... ___do____ Poisoned after 8 and 25_____ 10 survived, 13-percent weight loss. 10 Capsule Poisoned after 2 and survived, 13-percent 25..... weight loss. ...do Poisoned after 1 and survived, 19-percent weight loss. Drench... Poisoned and survived.

Drench... Poisoned after 1 and 100_____ 250_____ died. Sheep: 10 ...do Poisoned and survived, 5.... 13-percent weight loss. 10 ...do..... Poisoned after 8 and 10_____ survived, 7-percent weight loss. Capsule .. Poisoned after 9 and 25_____ survived, 5-percent weight loss. 10 Drench... Poisoned after 8 and 25_____ survived, 15-percent weight loss. 199 ...do..... Poisoned after 192 and 50_____ died. 10 ...do Poisoned and sacrificed, 50-----19-percent weight loss. Capsule... Poisoned after 3 and 50_____ survived, 23-percent weight loss. Drench... Poisoned after 3 and 100_____ died. Capsule.. Poisoned after 1 and 100_____ died. _do____ Do. 100_____ Drench ... Do. 250_____ 400_____ 2 ...do Do. Chickens:* Capsule ... 61-percent weight gain. 10 25_____ 10 ...do 24-percent weight gain. 50.... 10 ...do..... 20-percent weight gain, 10 ...do..... 2-percent weight loss, 45-percent weight gain. 100_____ 250_____ Controls....

TABLE 16.—Results of multiple oral dosing of cattle, sheep, and chickens with 2-chloro-4,8-bis (ethylamino) s-triesine (simazine) 1

Animal and dosage received (mg./kg.)	Dooes	Means of doeing	Results and remarks
	Num-		
Oatile:	ber_		
10	10	Drench	NIE.
25			Poisoned after 3 and aurvived, 5-percent weight loss.
25	10	do	Poisoned and survived, 12-percent weight loss.
80	10	do	Poisoned after 3 and aurvived, 21-percent weight loss, long recuperative period.
100	7	do	Poisoned after 3 and sacrificed.
250	3	do	Poisoned after 1 and survived, 11-percent weight loss.
Sheep:			weight tops.
25	10	do	NIE.
50		do	Poisoned after 17, died 14 days later, 13-
60	10	do	percent weight loss. Poisoned and survived, 18-percent weight
100	14	do	loss. Poisoned after 4 and died.
100	10	Capaule	Poisoned, with 13- percent weight loss, died 4 days after last dose.
250	8	Drench	Poisoned after 1 and survived, 9-percent weight loss.
250	10	Capsule	Poisoned after 3 and died.
400	9	Drench	Do.
Chickens:			
25	10	Capsule	52-percent weight gain.
50	10	do	29-percent Weight gain.
100	10	do	20-percent weight gain.
250	10	do	percent weight gain
Controls	*****		in survivors. 43-percent weight gain.

¹ Simazine, 80 percent wettable powder, Geigy Agricultural Chemicals, Ardsley, N.Y.

Atrasine, 80 percent wettable powder, Geigy Agricultural Chemicals, Ardeley, N.Y.

NIE indicates no ill effects apparent.

Average results of 5 treated chickens.

NIE indicates no ill effects apparent. Average results of 5 treated chickens.

swollen, friable, often light-brown liver; and petechine on the surface of the epicardium. At necropsy, one chicken had an enlarged, congested liver and congestion of the intestinal mucosa.

Application rates for simazine range from 1 to 9.6 pounds actual per acre. Rates in excess of 3 pounds per acre would be hazardous for cattle and in excess of 5 pounds for sheep. The 9.6-pound rate would be hazardous for chickens.

2,4-Bis(isopropylamino)-6-methoxy-s-triazine (prometone)

Both cattle and sheep were more susceptible to doses by drench than by capsule (table 17). Cattle were poisoned at 10 mg./kg., sheep at 25 mg./kg. Chickens dosed at 25 mg./kg. had a

significant reduced weight gain.

Signs of poisoning in cattle and sheep were anorexia and diarrhea, with increased salivation. At necropsy the sheep dosed at 100 mg./kg. had petechiae on the surface of the abomasal mucosa and areas of hemorrhage in the small intestine. The liver was swollen and friable and the kidneys were congested.

Application rates for prometone for use on noncrop areas vary from 9.5 to 57 pounds actual per acre. Rates in excess of 1 pound are hazardous for cattle; rates of 3 pounds or greater are hazard-

ous for sheep and chickens.

2-Chloro-4,6-bis(isopropylamino)-8triazine (propazine)

Cattle dosed by drench at 50 mg./kg. showed evidence of poisoning after 2 doses, but survived 10 with significant weight loss (table 18). A yearling dosed by drench at 25 mg./kg. was poisoned after 3, whereas another dosed by capsule had no ill effects after 10. Three of the 5 sheep dosed at 25 mg./kg. by either capsule or drench were poisoned after 5 to 16. The 4th sheep tolerated 59 by capsule with no ill effect, whereas the 5th sheep lost weight initially but regained it. Chickens were poisoned at 100 mg./kg., with significant reduced weight gains.

Signs of poisoning in cattle and sheep were anorexia and increasing depression and weakness. On occasion, a stilted gait was observed. The sheep dosed at 500 mg./kg. had lost 16 percent of its weight by the time it had received 10 doses, and anorexia continued for 14 days after the 10th dose. This sheep lost a total of 28 percent of its

pre-exposure weight before it died.

At necropsy on sheep, there was congestion of

the kidneys, adrenals, and liver. The mucosa of the abomasum and intestines was reddened. In sheep that died after long periods of toxicosis. there were often blood-engorged lungs, with froth in the respiratory tract. There were excessive amounts of red-tinged fluid in the thoracic and abdominal cavities. The one chicken fatality had congestion of the kidneys and the intestinal mucosa.

Application rates for propazine range from 2 to 4 pounds actual per acre. The maximum rate would be a hazard to cattle and sheep but not to chickens.

TABLE 17.—Results of multiple oral dosing of cattle, sheep, and chickens with 2,4-bis(isopropylamino)-6-methoxy-s-triazine (prometone)

Animal and desage received (mg./kg.)	Doses	Means of dosing	Results and remarks ²
	Num-	· · · · · · · · · · · · · · · · · · ·	
Cattle:	ber		
10	10	Capsule	NIE.
10	10	Drench	Poisoned after 8 and survived, 8-percent weight loss.
25	10	do	Poisoned after 1 and survived, 31-percent weight loss.
50	5	do	Poisoned after 2 and survived, 6-percent weight loss.
Sheep:		•	
10		do	
25	10	do	Poisoned after 4 and survived, 9-percent weight loss.
25	10	Capsule	NIE
50	îŏ		
50	10	Capsule	Poisoned after 1 and survived, 35-percent weight loss.
100	4	Drench	Poisoned after 1 and died.
Chickens: 4			
10			48-percent weight gain.
25	10	go	29-percent weight gain.
50			28-percent weight gain.
100	10	do	22-percent weight gain.
250	10	do	13-percent weight gain.
Controls			43-percent weight gain.

¹ Prometone, 25 percent emulsifiable concentrate, Geigy Agricultural Chemicals, Ardsley, N.Y.

NIE indicates no ill effects apparent.

Preumonia developed 1 day after last doso.

Average results of 5 treated chickens.

TABLE 18 .- Results of multiple oral dosing of cattle, sheep, and chickens with 2-chloro-4,6-bis (isopropylamino)-s-triazine (propazine)1

Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks *	Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks 1
	Num-				Num-		
Cattle:	ber		MID	Sheep-Con:	ðег		
10	10	Drench	NIE.	50	EQ.	do	Poisoned after 8, 11-
25			Poisoned after 3 and a survived.	00	, 43	++= 4 0+=++	percent weight loss, died 12 days after
25		Capsule	Poisoned after 2 and	ŀ			last doso.
50	10		survived, 16-percent weight loss.	60	. 59	do	Poleoned and survived, 22-percent weight loss.
100			Poisoned after 1 and survived, 12-percent weight loss.	50	. 10	Drench	 Poisoned and survived, 13-percent weight
250	7	do	Poisoned after 2 and survived, 21-percent weight loss.	100	. 10	do	loss. Poisoned and survived, 20-percent weight
Obase.				260	. 10	do	Poisoned after 3 and
Sheep: 10	10	Drench	NIE.				survived, 21-percent
25		do	Poisoned and survived,				weight loss.
		_	5-percent weight loss.	500	. 10	do	Poisoned after 1, 16-
25	5	Capsule	Poisoned and died.	1			percent weight lose,
25	21	do	Poisoned after 15 and survived, 19-percent				died 14 days after last dose.
			weight loss during	Chickens:			
	*^	do	trial period.	50	10	Capsule	49-percent weight gain
25 25	₽9 FO	uo	7-percent weight loss	100	10	do	. 28-percent weight gain
<i>x</i> 3			after 8, otherwise NIE.	250			. 1 died after 6, 18- percent weight gain
50			Poisoned after 5 and died.	Controls			in survivors. 33-percent weight gain
50	32	do	Poisoned after 17 and survived, 32-percent weight loss during trial period.				

² Propazine, 80 percent wettable powder, Geigy Agri-cultural Chemicals, Ardsley, N.Y. ³ NIE indicates no ill effects apparent.

2,4-Bis[(3-methoxypropyl)amino]-6-(methylthio)-s-triazine

Cattle were dosed by either drench or capsule, sheep by drench, chickens by capsule (table 19). No signs of poisoning were observed in cattle. Signs of poisoning in sheep dosed at 100 mg./kg. were observed after 1 and 8 doses. Chickens dosed at 250 mg./kg. had reduced weight gain, and 3 dosed at 500 mg./kg. died.

Anorexia was the most prominent sign of poisoning, but diarrhea and depression were noted

* Complicated by pneumonia, which was successfully treated.

Average results of 5 treated chickens.

at times. A swelling in the mandibular area developed suddenly in a yearling dosed at 50 mg./kg. after the 8th and final dose. This swelling was considered to be caused by the chemical irritation on the pharyngeal mucosa and resulted in dyspnea and partial anorexia. (See p. 5.) A sheep was severely poisoned at 250 mg./kg. after one dose and showed lameness and an uncoordinated gait within one-half hour after it was dosed. It became prostrate, extremely depressed, and moribund during the next 7 hours. Even though we expected the sheep to die, after a

TABLE 19.—Results of multiple oral dosing of cattle, sheep, and chickens with 2,4-bis [(8methoxypropyl) amino]-6-(methylthio)-s-triazine 1

Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks
	Num- ber		
Cattle:			
25		Drench	
50	8	do	27-percent weight loss due to irritation effect. ¹
100	10	Capsule	
Sheep:			
25	10	Drench	Do.
50		do	Do.
100	ĩň	do	Poisoned after 1 and
***************************************	20	22.40.31	survived, 12-percent weight loss.
100	9	do	Poisoned after 8 and survived, 10-percent weight loss.
250	1	do	Poisoned and survived, 10-percent weight loss.
Chickens: 4			
100	10	Capsule	40-percent weight gain.
250	10	do	21-percent weight gain.
500	10	do	3 died after 4 to 7, no weight change in survivors.
Controls		******	43-percent weight gain.

Lambast®, 25.5 percent emulsifiable concentrate, Monsanto Chemical Co., St. Louis, Mo.

NIE indicates no ill effects apparent.

Affected by chemical reaction of the formulation on

period of several days of partial anorexia, it made a slow but uneventful recovery. At necropsy on three chickens dosed at 500 mg./kg., congestion of the intestinal mucosa and the kidneys was seen.

Registration of this herbicide was cancelled in 1968; however, 10 pounds actual per acre would be hazardous for sheep but not for cattle or

2.4-Bis(isopropylamino)-6-(methylthio)-striazine

Cattle and sheep were dosed by drench, with poisoning at 50 mg./kg. for one yearling after 2 doses and at 100 mg./kg. for 2 sheep after 6 and 8 (table 20). A significant reduction in weight gain occurred at 50 mg./kg. for chickens dosed by capsule.

Signs of poisoning in cattle and sheep were anorexia and depression. The severely poisoned yearling that died after 4 doses at 100 mg./kg. had an uncoordinated gait before prostration and death. The sheep given 9 doses at 100 mg./kg. had a

similar uncoordinated gait. At necropsy on one sheep, congestion of the respiratory tract mucosa and blood-engorged lungs were seen, with the subcutaneous tissue surrounding the larynx and upper traches filled with reddish, edematous infiltrate. The liver was friable and light brown, the kidneys were congested, and the adrenals were enlarged. Lesions in the yearling were similar to those in the sheep.

Application rates for this herbicide range from 0.64 to 6 pounds actual per acre. The maximum rate would be hazardous for cattle and chickens but not for sheep.

TABLE 20.—Results of multiple oral dosing of cattle, sheep, and chickens with 2,4-bis(isopropylamino)-6-(methylthio)-s-triazine 1

Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks a
---	-------	-----------------	-----------------------

eri val.	Num- ber		
Cattle:	10	Drench	NIE.
50			Poisoned after 2 and survived, 6-percent weight loss.
100	4	do	Poisoned after 1 and died.
Sheep:			
25	10	do	NIE.
50	10	do	Do.
100	9	do	Poisoned after 8 and survived, 8-percent weight loss.
100	10	do	Poisoned after 6 and died.
Chickens:			
25	10	Capsule	48-percent weight gain.
50	10	do	44-percent weight gain.
100	10	do	34-percent weight gain.
250			Poisoned after 2, 4- percent weight loss.
Controls			49-percent weight gain.

Caparol® or Prometryne®, 80 percent wettable powder, Geigy Agricultural Chemicals, Ardsley, N.Y.

NIE indicates no ill effects apparent.

the pharyngeal mucosa, resulting in mandibular area enlargement accompanied by partial anorexia and dyspnea. Average results of 5 treated chickens.

Average results of 5 treated chickens.

Benzoic Acid Compounds

Animal

2-Methoxy-3,6-dichlorobenzoic acid (dicamba), dimethylamine salts

Cattle were poisoned at 250 mg./kg. after 2 doses by drench, whereas sheep were poisoned at 500 mg./kg. after 1 (table 21). Chickens dosed by capsule at 50 mg./kg. had significant average decreased weight gain after 10; however, those dosed at 100 mg./kg. had an average increased

weight gain.

Signs of poisoning in cattle and sheep were convulsions, tympanites, and prostration. At necropay on sheep, lymph nodes throughout the body were enlarged and hemorrhagic and there were areas of hemorrhage on the surface of the epicardium. The liver and kidneys were congested, the spleen was enlarged, and there was redness of the intestinal mucosa.

Application rates for dicamba range from 0.125 to 1 pound actual acid equivalent per acre. These rates would not be a hazard to the three test species.

TABLE 21.—Results of multiple oral dosing of cattle, sheep, and chickens with 2-methoxy-3,6dichlorobenzoic acid (dicamba), dimethylamine salts 1

Animal and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks *
	Num	•	
Cattle:	ber		
100		Dronoh	NIE.
250	2	do	Poisoned and survived.
250	10	do	NIE.
Sheep:	_		
100	`10	do	Do.
200	7	do	Do.
250	10	do	Do.
500			Poisoned and died,
500		do	
1,000	1	do	
Chickens:			
50	10	Capsule	43-percent weight gain.
100			55-percent weight gain.
250			45-percent weight gain.
500			49-percent weight gain.
Controls			53-percent weight gain.

Banvol D®, 4 lb./gal. emulsifiable concentrate, Velsical Chemical Corp., Chicago, Ill.
NIE indicates no ill effects apparent.

2,3,6-Trichlorobenzoic acid (2,3,6-TBA) and related polychlorobenzoic acids, dimethylamine salts

Cattle dosed by drench at 500 mg./kg. were poisoned by 1 and 2 doses, with one yearling surviving 8 (table 22). Reactions of sheep drenched at different dosage levels varied, although a significant weight loss occurred at 25 mg./kg. after 10. Chickens dosed at 250 mg./kg. by pipette had a significant reduction in weight gain; in contrast, chickens dosed at 500 mg./kg. had an insignificant reduction.

Signs of poisoning in cattle and sheep usually

TABLE 22.—Results of multiple oral dosing of oattle, sheep, and chickens with 2,3,6-trichlorobenzoic goid (2,3,6-TBA) and related polychlorobenzoic acide, dimethylamine salts 1

and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks *
	Num	-	
Cattle:	061		
250	10	Drench	NIE.
500			Poisoned after 2 and survived.
\$00	1	go	Poisoned and died 6% months after dose.
Sheep:			
10	10	do	NIE.
25	10	do	5-percent weight loss.
60	10	do	Poisoned, 15-percent weight loss, sacrificed 60 days after last dose.
80	10	do	
100		do	
100		do	
250			Poisoned after 3, 9-percent weight loss, died 2 days after last dose.
250	10	do	Poisoned and died 2 days after last dose, 6-percent weight loss.
Chickens:			·· - -
100	10	Pipette	43-percent weight gain.
250		do	30-percent weight gain.
500		do	45-percent weight gain.
Controls			
* · · · · · · · · · · · · · · · ·			-

¹ Trysben®, 26.1 percent emulsifiable concentrate, E. I. DuPont de Nemours and Co., Wilmington, Del.

NIE indicates no ill effects apparent.

^{*}Average results of 5 treated chickens.

Average results of 5 treated chickens.

were anorexia, depression, diarrhea, and lameness. The yearling acutely poisoned after a single dose at 500 mg./kg. became chronically affected, with lameness persisting for 6% months before death. Partial anorexia continued for 6 weeks in 1 sheep with appearances of abdominal pain while cating. The animal became weaker until prostration and a moribund condition developed. Then, 60 days after the last dose, the sheep was sacrificed.

At necropsy, the chronically affected yearling had edematous infiltration of the brisket, thoracic, and inner thigh muscles. At necropsy on sheep,

edema was found in the tissues surrounding the upper respiratory tract, in the pericardial sac, and in the peritoneal cavity. The liver was friable and enlarged and the kidneys appeared congested, with petechiae on the surface of the cortex. There was congestion of the intestinal and respiratory tract mucosa. The lungs were engorged with blood.

Registration of 2,3,6-TBA for use on croplands and rangelands was cancelled in 1968; however, rates in excess of 3 pounds actual per acre would be hazardous for sheep but would be no hazard

to cattle and chickens.

Animal

Miscellaneous Compounds

5-Bromo-3-sec-butyl-6-methyluracil (bromacil)

Cattle and sheep were dosed by either drench or capsule, chickens by capsule (table 23). A yearling dosed at 250 mg./kg. by capsule was poisoned after one dose. One sheep dosed at 50 mg./kg. by drench was not poisoned after 10; another dosed at 50 mg./kg. by capsule was poisoned, along with weight loss. Chickens dosed at 250 mg./kg. had a significant reduction in average weight gain after 10.

Signs of poisoning in cattle and sheep were anorexia and depression, although tympanites and an uncoordinated gait were occasionally observed in some.

Application rates for bromacil range from 1.6 to 20 pounds actual per acre. Rates in excess of 5 pounds would be hazardous for sheep but a rate of 20 pounds would not be hazardous for cattle and chickens.

5-Bromo-3-isopropyl-6-methyluracil (isocil)

Cattle were dosed by drench, sheep by either drench or capsule, chickens by capsule (table 24). Cattle dosed at 50 mg./kg. were poisoned after 7 doses. Dosing was stopped after 9 when complications of pneumonia developed. A sheep dosed at 100 mg./kg. by drench was poisoned after 1 but survived 10, with significant weight loss; but a sheep dosed at 100 mg./kg. by capsule had no ill effects after 10. Chickens dosed at 250 mg./kg. had a reduced weight gain after 10.

Signs of poisoning in cattle and sheep were anorexia and depression, which were often accompanied by tympanites, an uncoordinated gait, and prostration. Convulsions were observed before death.

At necropsy enlarged, hemorrhagic lymph nodes were found in cattle and sheep. There were hemorrhages on the surface of the epicardium and lungs were congested. The upper respiratory tract mucosa was reddened. In one animal the

intestinal mucosa was congested; in another, petechiae on the surfaces of the liver and kidneys were present. At necropsy, an enlarged and congested liver, capsular hemorrhages in the spleen, and swollen, congested kidneys were seen in the chicken.

Table 23.—Results of multiple oral dosing of cattle, sheep, and chickens with 5-bromo-3-sec-butyl-6methyluracil (bromacil) 1

and dosage Doses Means of Results and remarks 1

received (mg./kg.)	Doses	dosing	1000min and 10mm as
	Num-		
Cattle:	ber		
100	10	Drench	NIE.
100	10	Capsule	Do.
250	10	do	Poisoned after 1 and survived, 14-percent weight loss.
Sheep:			
25	10	Drench	NIE.
50	10		Poisoned and survived, 8-percent weight loss.
50	10	Drench	NIE.
100	10	Capsule	
100	10		Poisoned and survived, 9-percent weight loss.
250	5	do	Poisoned after 3 and survived, 9-percent weight loss.
250	10	Capsule	Poisoned after 8 and survived, 8-percent weight loss.
Chickens:			
100	10	do	49-percent weight gain.
250		do	
500			24-percent weight gain.
Controls			48-percent weight gain.

Hyvar X®, 80 percent wettable powder, E. I. DuPont de Nemours and Co., Wilmington, Del.
 NIE indicates no ill effects apparent.

Average results of 5 treated chickens.

and dosage

Animal

Table 24.—Results of multiple oral dosing of cattle, sheep, and chickens with 5-bromo-3-isopropyl-6methyluracil (isocil)1

and dosage received (mg./kg.)	Doses	Means of desing	Results and remarks
	Num-		
Cattle:	ber		
25		Drench	
50			Poisoned after 7, 11- percent weight loss.
100	10	do	6-percent weight loss.
250	2	do	Poisoned and died.
Sheep:			
50		do	
100	10	do	Poisoned after 1 and survived, 15-percent weight loss.
100	10	Capaule	NIE.
100	10	Drench	Poisoned and survived, 15-percent weight loss.
250	4	do	Poisoned after 1 and died.
250	3	Capsule	Poisoned and died.
Chickens: 4			
100	10	do	49-percent weight gain.
250	10	do	38-percent weight gain.
500			1 died after 7, 12- percent weight gain in survivors.
Controls			47-percent weight gain.

Hyvar Isocii[®], 80 percent wettable powder, E. I. DuPont de Nemours and Co., Wilmington, Del.

NIE indicates no ill effects apparent.

Application rates for isocil to noncrop areas range from 1.6 to 26 pounds actual per acre. Rates in excess of 5 pounds actual per acre would be hazardous for cattle and 10 pounds for sheep; however, even the maximum rate (26 pounds) would not be hazardous for chickens.

Polychlorobicyclopentadiene isomers

Cattle and sheep were dosed by capsule or drench, chickens by capsule (table 25). A yearling was poisoned at 100 mg./kg. after 3 doses but survived; a sheep was poisoned at the same rate after 7 but eventually died. Chickens dosed at

TABLE 25.—Results of multiple oral dosing of cattle, sheep, and chickens with polychlorobicyclopentadiene isomers 1

Results and remarks 2

Doses Means of

(mg./kg.)		dosing	
	Num-	<u> </u>	
Cattle:	ber		
25 *	10	Capsule	NIE.
50 3		do	
100	10	do	Poisoned after 3 and survived, 6-percent weight loss.
250	3	Drench	Poisoned after 2 and died 8 days after last dose.
Sheep:			
25 3	10	Capsule	NIE.
80 8	10	do	Do.
100	10	do	Poisoned after 7, with 13-percent weight loss; died 13 days after last dose, with 34-percent weight loss.
200	1	Drench	
250			Poisoned after 1 and survived.
1,000	1	do	Poisoned and died.
Chickens: 4			
25 4	10	Capsule	40-percent weight gain.
80	10	do	27-percent weight gain.
100	10	do	1 died after 10,
	-•		11-percent weight loss in survivors.
250	9	do	Affected after 3 and
	. •		died after 7 to 9.

Bandane®, 4 lb./gal. emulsifiable concentrate, Velsicol Chemical Corp., Chicago, Ill.
NIE indicates no ill effects apparent.

Controls..... 40-percent weight gain.

50 mg./kg. showed significant weight gain reduction.

Signs of poisoning were anorexia, depression, muscular tremors, and hyperexcitability. In the sheep chronically poisoned at 100 mg./kg., there was an uncoordinated gait, followed by prostration, convulsions, and death.

At necropsy on cattle and sheep, hemorrhages were found along the brain stem and in the anterior

^{*} Toxicity complicated by pneumonia, which developed during treatment.

Average results of 5 treated chickens.

^{*} Technical material was used in more recent trials. Average results of 5 treated chickens.

region of the cerebrum. Meningeal vessels were congested. The liver was friable and congested. Petechiae were seen on the surface of the epicardium. At necropsy on the chronically poisoned sheep, the lungs and the respiratory tract mucosa were congested. The liver was enlarged and friable. the kidneys were reduced in size, and the pancreas was congested. Chickens poisoned had swollen. congested livers and kidneys, and the intestinal mucosa was hemorrhagic.

Application rates for polychlorobicyclopentadiene isomers to noncrop areas, such as lawns and turf, range from 30 to 40 pounds actual per acre. These rates would be highly hazardous for all three

test species.

Animal

anneath has

TABLE 26.—Results of multiple oral dosing of cattle, sheep, and chickens with 3',4'-dichloropropionanilide (propanil) 1

Donules and sameshall

Three Moons of

received (mg./kg.)	Doses	Means of dosing	results and remarks .
	Num-		
Cattle:	ber		
50	10	Drench	NIE.
100	10	do	Do.
250	1	do	Poisoned and survived.
Sheep:			
25	10	do	NIE.
25	10	Capsule	Do.
50	10	Drench	Do.
100	10	do	Poisoned and survived,
			11-percent weight
100	10	Capsulo	
250	ī	Drench	Poisoned and survived,
	_	,	14-percent weight loss.
Chickens:			
10	10	Capsule	48-percent weight gain.
25	10	do	10-percent weight gain.
50	10	do	20-percent weight gain.
100	10	do	3 died after 8 or 9, 13-
			percent weight gain in survivors.
250			All poisoned and died after 2 to 6.
Controls			48-percent weight gain.

¹ Rogue®, 46 percent emulsifiable concentrate, Mon-santo Chemical Co., St. Louis, Mo. 1 NIE indicates no ill effects apparent.

3'.4'-Dichloropropionanilide (propanil)

A yearling was poisoned at 250 mg/kg. after I dose by drench (table 26). One sheep was poisoned at 100 mg./kg. after 10 by drench, but another had no ill effects after 10 by capsule. Chickens dosed at 25 mg./kg. 10 times had a significant reduction in weight gain.

In cattle and sheep, the signs of poisoning at 250 mg./kg. were anorexia, depression, an uncoordinated gait, and prostration; all were observed over a period of a few hours in the 2 animals receiving 1 dose each. Anorexia was the only sign in the sheep less acutely poisoned at

100 mg./kg.

At necropsy, chickens had congested intestinal mucosa, swollen kidneys, and a distended gall

bladder.

The application rate for propanil to rice crops is 6 pounds actual per acre but not to exceed 8 pounds per season. Such rates would not be a hazard to cattle and sheep, but would be to chickens.

2,6-Dichlorobenzonitrile (dichlobenil)

A yearling dosed at 50 mg./kg. by capsule was poisoned after 1 dose (table 27). A sheep dosed at 10 mg./kg. by capsule had no ill effects after 60. A sheep dosed at 25 mg./kg. by capsule was poisoned after 4; a sheep dosed by drench was poisoned after 10. Chickens dosed at 25 mg./kg. by capsule had significant reduced weight gains.

Signs of toxicosis in cattle and sheep were increased salivation, anorexia, and depression. Convulsions were observed before death in some

instances.

At necropsy lesions in cattle and sheep were extensive as in generalized toxemia. There were swollen, hemorrhagic lymph nodes throughout the body and hemorrhages in the muscles, chiefly in the thoracic and cervical areas. Petechiae were on the surface of the epicardium. Congestion of the lungs and upper respiratory tract mucosa were commonly seen. The liver was swollen and congested in cases of acute poisoning, but often light brown and friable in prolonged or chronic poisoning. The kidneys were congested and swollen; however, in the sheep that died 30 days after the last dose, they were firm and reduced in size. The mucosa of the abomasum and of the intestinal tract was congested, whereas the mucosas of the more acutely poisoned shoop were markedly inflamed. At necropsy, chickens were notably emaciated, and

Average results of 5 treated chickens.

and dosage

Animal

TABLE 27.—Results of multiple oral dosing of cattle, sheep, and chickens with 2,6-dichlorobenzonitrile (dichlobenil)

and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks*
	Num-		,
Cattle:	ber	a	27787
10	ĬÕ	Capsule	MIR.
25	ΤŇ	do	Do.
50	Z	do	Poisoned after 1 and died.
Sheep:			•
5	60	do	NIE.
10		do	Do.
10	10	Drench	Do.
25	10	Capsule	Poisoned and survived, 18-percent weight loss.
25	13	do	Poisoned after 4 and died, 29-percent weight loss.
25	10	Drench	Poisoned after 10, 15-percent weight loss, died 20 days after last dose.
50	10	Capsule	Poisoned after 10, 26-percent weight loss, died 2 days after last dose.
50	10	Drench	Poisoned after 6 and died, 14-percent weight loss.
100	1	do	Poisoned and died.
Chickens:			
10	10	Capeule	62-percent weight gain.
25	10	do	11-percent weight gain.
50	10	do	9-percent weight gain.
100	10	do	2 died, 23-percent weight loss in survivors.
Controls			37-percent weight gain.

² Casoron[®], 50 percent wettable powder, Thompson-Hayward Chemical Co., Kansas City, Kans. ³ NIE indicates no ill effects apparent.

the liver, kidneys, and intestinal mucosa were congested.

Application rates for dichlobenil range from 2 to 6 pounds actual per cropland acre and 15 pounds actual per water acre. A 3-pound rate per acre would be hazardous for sheep and chickens and a 6-pound rate for cattle.

4-Amino-3.5.6-trichloropicolinic acid (picloram), potassium salt

Cattle dosed at 500 mg./kg. by drench were poisoned after 8 doses; sheep dosed at 250 mg./kg. by either drench or capsule were poisoned after 9 or 10 (table 28). Chickens dosed at 500 mg./kg. had a significant reduction in average weight gain.

In cattle and sheep, weakness, depression, and anorexia were the most prominent signs of poisoning. A delayed toxicity led to the death of a yearling and a sheep 6 days after the last dose.

At necropsy on the yearling and the sheep, lymph nodes of the head, neck, and thoracic cavity were swollen and hemorrhagic. The lungs and the respiratory tract mucosa were congested.

Table 28.—Results of multiple oral dosing of cattle, sheep, and chickens with 4-amino-3,5,6trichloropicolinic acid (picloram), potassium salt 1

Doses Means of Results and remarks 1

received (mg./kg.)		dosing	
	Num-		· · · · · · · · · · · · · · · · · · ·
Cattle:	ber		
100	10	Drench	NIE.
250	10	do	Do.
590	8	do	Poisoned after 8 and died 6 days after
			last dose, 24-percent weight loss.
Sheep:			
100	10	Capsule	NIE.
100	10	Drench	Do.
250	10		Poisoned after 9 and
2002001070		<u>-</u> -	died 6 days after last dose.
250	16	Drengh	Poisoned and survived,
200111111	••		6-percent weight loss.
250	10	do	
500		do	
•			survived, 17-percent weight loss.
Chickens: 4			
100	10	Pipette	34-percent weight gain.
250		do	
500		do	
Controls			AA

¹ Tordon[®], 24.9 percent emulsifiable concentrate Dow Chemical Co., Midland, Mich. ² NIE indicates no ill effects apparent.

Average results of 5 treated chickens.

[•] Average results of 5 treated chickens.

lamiuA.

The rumens were filled with undigested feed and livers were engorged with blood and swollen. The adrenals were enlarged and congested. The yearling had pleural adhesions and a congested pancreas. The sheep had hemorrhages throughout the heart and blood-tinged fluid in the pericardial sac. The kidneys and the intestinal mucosa of the sheep were congested.

Registration of picloram for use on cropland was cancelled in 1968; however, a rate in excess of 25 pounds actual per acre would present a hazard

to sheep but not to cattle or chickens.

2,2-Dichloropropionic acid (dalapon), sodium salt

Cattle and sheep were dosed by either drench or capsule, chickens by capsule (table 29). A yearling dosed at 500 mg./kg. by capsule was not poisoned after 10 doses; however, a yearling dosed at 500 mg./kg. by drench, although not poisoned, developed a swelling in the parotid area after 9 days. This swelling was considered to be caused by chemical irritation of the pharyngeal mucose. (See p. 5.) Sheep dosed at 500 mg./kg. by either drench or capsule were poisoned. One sheep dosed at 100 mg./kg. by capsule was poisoned and had a 6-percent weight loss after 10. Another sheep dosed at 100 mg./kg. by capsule 481 times had a 10-percent weight loss after 86 but suffered no ill effects. A sheep dosed at 100 mg./kg. by drench had no ill effects after 10. Chickens dosed at 250 mg./kg. had a significant average reduced weight gain after 10; whereas chickens dosed at 500 mg./kg. had slightly less reduced weight gain than the chickens dosed at

Anorexia and depression were the signs of poisoning in the sheep. Recovery was uneventful

after the dosing regimen was completed.

Application rates for dalapon range from 1.1 to 14.8 pounds actual per acre. These rates would be no hazard to the cattle and chickens, but a rate in excess of 10 pounds would be a hazard to sheep.

TABLE 29.—Results of multiple oral dosing of cattle-sheep, and chickens with \$,2-dichloropropionic acid (dalapon), sodium salt 1

and dosage received (mg./kg.)	Doses	Means of dosing	Results and remarks 2
	Nun- ber		
Cattle:	• • •		
250		Drench	
500	9	do	Irritation effect after
50 0	10	Capsule	
Sheep:			
50	10	Drench	NIE.
100	10	Capsule	Poisoned and survived, 6-percent weight loss.
100	481	do	10-percent weight loss after 86, otherwise NIE.
100	10	Drench	NIE.
250	10	do	Do.
.500			Poisoned and survived, 17-percent weight loss.
500	10	Capsule	Poisoned after 7 and survived, 6-percent weight loss.
Chickens: 4			
100	10	do	43-percent weight gain.
250	10	do	27-percent weight gain.
500	10	do	30-percent weight
Controls		*******	gain. 42-percent weight gain.

Dowpon Grass Killer®, 85 percent wettable powder, Dow Chemical Co., Midland, Mich.
NIE indicates no ill effects apparent.
Affected by chemical reaction of the formulation on the pharyngeal mucosa, resulting in parotid area enlargement accompanied by partial anorexia and dyspnea.

Average results of 5 treated chickens.

COMMENT

These data have been abstracted in table 30, "Summary of dosages of various organic herbicides that cause significant weight loss, reduced weight gain, or poisoning in cattle, sheep, and chickens."

Table 30.—Summary of dosages of various organic herbicides that cause significant weight loss, reduced weight gain, or poisoning in cattle, sheep, and chickens 1

Herbicide	Dosage	Least number of desages for—		osages for—	D -aktatda	Dosage	Least nu	mber of d	osages for-
Derbicide Face	rate	Cattle	Sheep	Chickens	Herbicide	rate	Cattle	Sheep	Chicken
	Mg./kg.					Mg./kg.			
hlorophenoxy compounds:					Phenyl urea compounds:				
2,4-Dichlorophenoxyacetic	f 500		7	10		/ 500	1		
acid (2,4-D), alkanolamine	250	1	10	10	2 /n ('himmhmall 1 1	250		. 2	6
salts (of the ethanoi and	3 200	25 86		*******	3-(p-Chiorophenyi)-1,1- dimethylures (monuron).	{ 100		. 4	8
isopropanol series).	į 100	86	8		mmeentraces (montrout.	50		****	. 10
2,4-Dichlorophenoxyacetic	250	2	2	4	[(25			10
acid (2,4-D), propylene	1 100	•	-	10	}	250		. 8	
glycol butyl ether ester.	(100	******		10	3-(3,4-Dicblorophenyl)-1-	100	8	4	. 8
2,4,5-Trichlorophenoxy-	500			. 5	methory-I-methylures	₹ 50	10	1	10
acetic acid (2,4,5-T),	250	4	-	10	(linuron).	25			10
propylene glycol butyl	100	-	3			10			10
ether ester.	(-00		•	*********	3-Phenyl-1.1-dimethylurea	500	2	3	10
2-(2,4,5-Trichlorophenoxy)-			_		(fenuron).	{ 250		10	
propionic acid (silvex),	250	*****	Ď	10	()	100		8	
propylene glycol butyl	100	19	9	10		250	:	1	.8
ether ester.	ţ				3-(3,4-Dichlorophenyl)-1,1-	100	10	2	10 10
2-Methyl-4-chlorophenoxy-	500	8			dimethylurea (diuron).	50			10 10
acetic acid (MCPA),	250	8	· · · · - Ā · · · ·	10	, , , ,	25		-	10
alkanolamine salts (of the	100	•	•	70	dittanala mate an amadas	(10			10
ethanol and isopropanol	1 100		2		Thiocarbamate compounds:	/ 250			7
scries).						175			10
mide compounds: N.N-Dimethyl-2,2-diphenyl-	f 500			. 10		150			10
acetamide (diphenamid).	250	3		10	S-2,3-Dichloroallyl diisopropyl-	125			10
acecamiae (diphenamia).	1 250	٥	- ;	10	thiocarbamate (diallate).	100			10
2-Chloro-N, N-diallyl-	100		1	1	tuiqear Damiace (diamace).	50	3	ž	
acetamide (CDAA).	1 50	1	;	ģ	1	25	ĭ	ĭ	
tectablide (CDAA).	25	*	1	10		250		ź	-
2-Chloro-N.N-dialiyl-	1 250	•	7	10		200			10
acetamide (CDAA) and	100		î	10	S-2,3,3-Trichloroallyl	175			îŏ
trichlorobenzyl chloride	{ 150	2	2	10	diisopropylthiocarbamate	106		<u>-</u>	
(TCBC).	25	•	ĩ.	*******	(triallate).	50	3	-	
(1000).	, 20		U		1	25	ž	-	

Triasine compounds:			Benzoic acid compounds:				
2-Chloro-1-ethylamino- 6-isopropylamino-s- triszine (atrazine).	400	10 10 10 3 10	2-Methoxy-3,6-dichlorobenzoic acid (dicamba), dimethyl- amine salts. 2,3,6-Trichlorobenzoic acid (2,3,6-TBA) and related polychlorobenzoic acids,	\$500 250 50 500 250 100 50	2 1	3 10 10	10 10 10
2-Chloro-4,6-bis(ethylamino)-s- triazine (simazine).	400	8 1 10 10	dimethylamine salts. Miscellaneous compounds: 5-Bromo-3-sec-butyl-6-methyl- uracil (bromacil).	<pre>500 250 100</pre>	1	10 	. 10 10
2,4-Bis(isopropylamino)-6- methoxy-s-triazine (prome- tone).	250	10 10 10 10 10	5-Bromo-3-isopropyl-6-methyl- uracil (isocil).	500 500 250 100 50	2 16 7	10 1 1	7 10
2-Chloro-4,6-bis (isopropylamino)- s-triazinė (propazinė).	500 1 250 2 3 100 1 10 50 2 5 25 3 5	10	Polychlorobicyclopentadiene isomers.	1,000 250 100 50 250	3	1 7	3 10 . 10
2,4-Bis[(3-methoxypropy!)- amino]-6-(methylthio)-s- triazine.*	500	10	3',4'-Dichloropropionanilide (propanil).	100 50 25	*******	10	. 10 . 10
2,4-Bis(isopropylamino)-6- (methylthio)-s-triazine.	250 100 1 6 50 2	10 10	2,6-Dichlorobenzonitrile (dichlobenil). 4-Amino-3,5,6-trichloropicolinic	100 50 25	1	1 6 4	10 10 10
			acid (picloram), potassium salt.	500 250 500	8	2 9 7	10
			2,2-Dichloropropionic acid (dalapon), sodium salt.*	250 100		10	. 10

¹ Broken lines indicate that either the herbicide was not tested at that rate or the herbicide was not toxic to the animal at that rate.

¹ No toxic desages found for cattle.

SUMMARY AND CONCLUSIONS

Results of studies of the toxicity of 29 organic herbicides to cattle, sheep, and chickens have been presented. A total of 126 yearling cattle, 190 oneto two-year-old sheep, and 700 six-week-old chickens were studied.

Repeated doses of the herbicides were administered in gelatin capsules or as water-diluted solutions by drench or in pipettes. The usual period of study was 10 days, or until toxicologic effects appeared. However, in a number of instances, longer and shorter term studies were made.

The signs of poisoning by most of the herbicides included anorexia and reduced weight gains. In many instances digestion in the rumen seemed to have been impaired. Macroscopic lesions were somewhat variable and nonspecific. The liver and the kidneys were most frequently involved.

An arbitrary yield of forage and rate of con-sumption were selected to evaluate the hazard for cattle, sheep, and chickens likely to exist under

most severe conditions of use.

The most common rates of application of many of these herbicide formulations are not a hazard to cattle, sheep, or chickens. The maximum rates of application of some approach or surpass a hazardous level in one or more of the test species.

LITERATURE CITED

(1) BUCE, W. B., BINNS, W., JAMES, L., and WILLIAMS, M. G. TRESULTS OF FEEDING OF HESSICIDE-TREATED PLANTS TO CALVES AND SHEEP. Amer. Vet. Med. Assoc. Jour. 138: 320-1961. 323.

(2) DALGAARD-MIKKELSON, S. V., AND POULSON, E. 1962. THE TOXICOLOGY OF HERBICIDES. Pharm. Rev. 14: 225-250.

GRIGSBY, B. H., and FARWELL, E. D.

1950. SOME EFFECTS OF HERBICIDES ON PASTURE AND ON GRAZING LIVESTOCK. Mich. Expt. Sts. Quart. Bul. 32: 378-385. Mich. Agr.

(4) JACKSON, J. B.
1966. TOXICOLOGIC STUDIES ON A NEW HERBICIDE IN SHEEP AND CATTLE. Amer. Jour. Vot. Res. 27: 821-824.

MENEIE, C. M. METABOLISM OF PESTICIDES. U.S. Dept. Int., Fish and Wildlife Serv., Spec. Sci. Rpt. 96, 274 pages. 1966.

PALMER, J. S. CHRONIC TOXICITY OF 2,4-D ALKANOLAMINE BALT TO CATTLE. Amer. Vet. Med. Assoc. 1963. Jour. 143: 398-399.

1964. TOXICITY OF CARRAMATE, TRIABINE, DI-CHLOROPROPIONANILIDE, AND DIALLYLACET-AMIDE COMPOUNDS TO SHEEP. Amer. Vet. Med. Assoc. Jour. 145: 917-920.

TOXICITY OF METHYLURACIL AND BUBSTI-1964. TUTED UREA AND PHENOL COMPOUNDS TO Amer. Vet. Med. Assoc. Jour. BREKP. 145: 787-789.

(9) TOXICOLOGIC EFFECTS OF SILVEX IN TEAR-1964. LING CATTLE. Amer. Vet. Med. Assoc. Jour. 144: 750-755.

and RADELEFF, R. D. (10)(10)

1964. THE TOXICOLOGIC EFFECTS OF CERTAIN FUNGICIDES AND HERBICIDES ON SHEEF AND CATTLE. N.Y. Acad. Sci. Ann. HI: 729-736.

(11) PAYNTER, O. E., TUSING, T. W., McCollister, D. D., and Rowe, V. K.

1960. TOXICOLOGY OF DALAFON SODIUM (2,2-2007) OPPOSITION OF ACAD. SODIUM (2,2-2007) OPPOSITION OF ACAD.

DICHLOROPROPIONIC ACID, BODIUM BALT).
Jour, Agr. and Food Chem. 8: 47-51.

(12) RADELEST, R. D. 1958. ADVANCES IN VETERINARY SCIENCE. THE TOXICITY OF INSECTICIDES AND HERBICIDES TO LIVESTOCK. IV. pp. 265-276. Academic Press, Inc., N.Y.

(13) RERDICIDES AND PLANT GROWTH REGULA-tors. In Veterinary Toxicology, pp. 241-257. Lea and Febiger, Philadelphia, Pa. 1964.

and Bushland, R. C.

1960. THE TOXICITY OF PESTICIDES FOR LIVESTOCK. In The Nature and Fate of
Chemicals Applied to Soils, Plants, and
Asimals. U.S. Agr. Res. Serv. ARS 20-9,
pp. 134-160.

(15) ROWE, V. K., and HYMAS, T. A.

1954. SUMMARY OF TOXICOLOGICAL INFORMATION.

1954. BUHMARY OF TOXICOLOGICAL INFORMATION 1M 2,4-D AND 2,4,6-T TYPE HERBICIDES AND AN EVALUATION OF THE HAZARDS TO LIVESTOCK ASSOCIATED WITH THEIR USE. Amer. Jour. Vet. Res. 15: 622-629.

St. John, L. E., Jr., Wagner, D. C., and Lise, D. J. 1964. The pate of atrabine, kuron, silvex and (16)24.5-T IN THE DAIRY COW. Jour. Dairy Sci. 47: 1267-1270.

U.S. DEPARTMENT OF AGRICULTURE. (17) 1966. U.S.D.A. SUMMARY OF REGISTERED AGRIcultural preticide chemical uses. Ed. 2, Sup. III, 836 pp. [See also subsequent preliminary notices of U.S.D.A. pesticide

U.S. DEPARTMENT OF AGRICULTURE. (18)1987. ACCEPTABLE COMMON NAMES AND CHEMICAL MAMES FOR THE INGREDIENT STATEMENT ON BECONOMIC POISON (PESTICIDE AND PLANT GROWTH REGULATOR) LABELS. [Prepared by R. L. Caswell, Pesticides Regulation Division, U.S. Agr. Res. Serv.) 154 pages.

(19)U.S. DEPARTMENT OF AGRICULTURE. 1967. BUGGESTED GUIDE FOR WEED CONTROL 1967. U.S. Dept. Agr., Agr. Handb. 332, 64 pages.

Willard, C. J. THE STATUS OF HERBICIDE POISONING. North Central Weed Control Coul., 8th Meeting, 1951. Proc., pp. 86-89.
WRIGHT, F. C., PALMER, J. S., and HUNT, L. M.

(21)PRELIMINARY STUDY OF TOXICOLOGIC AND ejochemical effects of bilvex on sheep. Amer. Jour. Vet. Res. 27: 172-176.