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OCT 4 2001

DATE OF APPROVAL LETTER: _____

FREEDOM OF INFORMATION SUMMARY

SUPPLEMENTAL ABBREVIATED NEW ANIMAL DRUG APPLICATION

ANADA 200-066

AGRIMYCIN[®] 343 (oxytetracycline HCl)

“revised withdrawal period for turkeys and swine”

Sponsored by Agri Laboratories, Ltd.

FOIS

I. GENERAL INFORMATION:

ANADA: 200-066

Sponsor: Agri Laboratories, Ltd.
P.O. Box 3103
St. Joseph, MO 64503

Generic Name: Oxytetracycline HCl

Trade Name: AGRIMYCIN® 343 SOLUBLE POWDER

Marketing Status: OTC

Effect of the Supplement: Revised withdrawal period for turkeys and swine.

II. INDICATIONS AND DOSAGES FOR USE:

Turkeys: 200 to 400 mg per gallon for control of hexamitiasis caused by *Hexamita meleagridis*, susceptible to oxytetracycline.

400 mg per gallon for control of infectious synovitis caused by *Mycoplasma synoviae*, susceptible to oxytetracycline.

25 mg/lb body weight in GROWING TURKEYS for control of complicating bacterial organisms associated with bluecomb (transmissible enteritis, coronaviral enteritis), susceptible to oxytetracycline.

Swine: 10 mg/lb body weight FOR THE CONTROL AND TREATMENT OF THE FOLLOWING DISEASES IN SWINE – Bacterial enteritis caused by *Escherichia coli* and *Salmonella choleraesuis*, susceptible to oxytetracycline. FOR BREEDING SWINE: Leptospirosis (reducing the incidence of abortions and shedding of leptospira) caused by *Leptospira pomona*, susceptible to oxytetracycline.

III. DRUG EFFECTIVENES AND TARGET ANIMAL SAFETY:

Under the provisions of the Federal Food, Drug, and Cosmetic Act, as amended by the Generic Animal Drug and Patent Term Restoration Act, (53 FR 50460, December 15, 1988, First GADPTRA Policy Letter) an abbreviated new animal drug application (ANADA) may be submitted for a generic version of an approved new animal drug (pioneer product). New target animal safety data, drug effectiveness data, and human food safety data (other than tissue residue data) are not required for approval of an ANADA. An ANADA relies on the target animal safety, drug effectiveness, and human food safety data in the pioneer's new animal drug application. The ANADA sponsor is required to show that the generic product is bioequivalent to the pioneer. ANADA's for drug products for food-producing animals will generally be required to include bioequivalence and tissue residue studies. A tissue residue study will generally be required to accompany clinical end-point and pharmacologic end-point bioequivalence studies, and blood level bioequivalence studies that can not quantify the concentration of the drug in blood throughout the established withdrawal period. For certain dosage forms, the agency will grant a waiver from conducting an *in vivo* bioequivalence study (55 FR 24645, June 18, 1990; Fifth GADPTRA Policy Letter; Bioequivalence Guideline, October 10, 2000).

Based upon the formulation characteristics of the generic product, AGRIMYCIN[®] 343 was granted a waiver from conducting an *in vivo* bioequivalence study and approved on July 15, 1994. The generic and pioneer products contain the same active and inactive ingredients and are soluble powders.

IV. HUMAN FOOD SAFETY:

A. TOLERANCES

Tolerances are established in 21 CFR 556.500 for the sum of residues in tissues of swine and turkeys as follows: 2 ppm in muscle, 6 ppm in liver, 12 ppm in fat and kidney. The acceptable daily intake (ADI) for total oxytetracycline residues is 25 micrograms per kilogram of body weight per day.

B. WITHDRAWAL TIME:

When a waiver from the requirement of an *in vivo* bioequivalence study is granted, the withdrawal times are those previously assigned to the pioneer product. The withdrawal time for pioneer oxytetracycline soluble powders (NADA 130-435, Alpharma, Inc.'s Oxytet Soluble[®]) is established under 21 CFR 520.1660d. Alpharma, Inc. received approval on November 29, 2000, for a zero day withdrawal time for turkeys and swine. This supplemental ANADA requests the same zero-day withdrawal period for turkeys and swine based on the waiver

granted to the original approval of Agri Laboratories, Ltd.'s ANADA. No new data were required for the reduced withdrawal times in swine and turkeys.

C. REGULATORY METHODS

The regulatory method for determination of oxytetracycline in tissues is a microbiological assay procedure using *Bacillus cereus* var. *mycoides* (ATCC 11778) suspension and is found in the FDA publication "Antibiotic Residues in Milk, Dairy Products, and Animal Tissues: Methods, Reports, and Protocols" revised October 1968, reprinted December 1974.

V. AGENCY CONCLUSIONS

This supplemental ANADA satisfies the requirements of section 512 of the Federal Food, Drug and Cosmetic Act and implementing regulations at Part 514 of Title 21, Code of Federal Regulations (21 CFR 514) to demonstrate that AGRIMYCIN[®] 343 (oxytetracycline HCl), is safe and effective for use in turkeys and swine for the approved indications, when administered in water at the approved dose.

The agency has determined under 21 CFR 25.33(a)(1) that this action is of a type that does not individually or cumulatively have a significant impact on human environment. Therefore, neither an environmental assessment nor an environmental impact statement is required.

Under section 512(c)(2)(F)(iii) of the FFDCA, this approval for food-producing animals does not qualify for marketing exclusivity.

VI. APPROVED PRODUCT LABELING (attached)

- A. Pioneer Facsimile label – OXYTET Soluble 3.09 lbs (1400 g)
- B. Generic Facsimile labels – AGRIMYCIN[®] 343 - 50 packets 4.78 oz (135.5g)
AGRIMYCIN[®] 343 - 5 lbs (2.27 kg).

Copies of applicable labels may be obtained by writing to the:

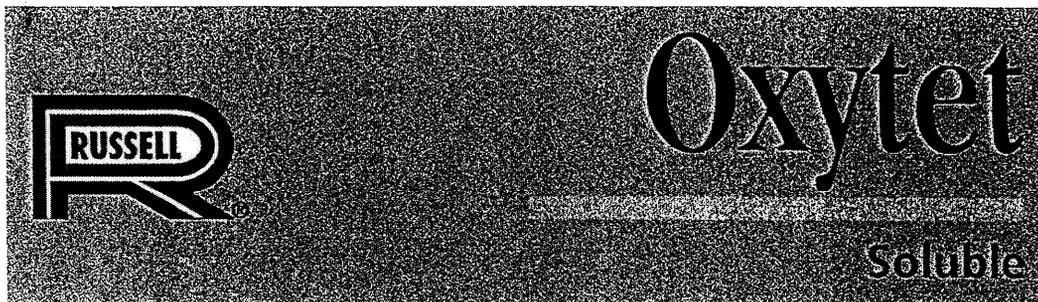
Food and Drug Administration
Freedom of Information Staff (HFI-35)
5600 Fishers Lane
Rockville, MD 20857

Or requests may be sent via fax to: (301) 443-1726. If there are problems sending a fax, call (301) 827-6500.

MIXING DIRECTIONS FOR WATER PROPORTIONERS CHICKENS — TURKEYS — SWINE

Dosage and indications for use in water proportioners - Add 1 pack to the following amount of water to make a stock solution for use in a proportioner set to meter at the rate of one ounce per gallon.

	Disease	Treatment Level	Gallons/Stock Pack
C H I C K E N S T U R K E Y S	Control of infectious synovitis caused by <i>Mycoplasma synoviae</i> , susceptible to oxytetracycline.	200 - 400 mg	20-10
	Control of chronic respiratory disease (CRD) and air sac infections caused by <i>Mycoplasma gallisepticum</i> and <i>Escherichia coli</i> , susceptible to oxytetracycline.	400 - 800 mg	10-5
	Control of fowl cholera caused by <i>Pasteurella multocida</i> , susceptible to oxytetracycline.	400 - 800 mg	10-5
	Control of hexamitiasis caused by <i>Hexamita meleagridis</i> , susceptible to oxytetracycline.	200 - 400 mg	20-10
	Control of infectious synovitis caused by <i>Mycoplasma synoviae</i> , susceptible to oxytetracycline.	400 mg	10
S W I N E	GROWING TURKEYS — Control of complicating bacterial organisms associated with bluecomb (transmissible enteritis, coronaviral enteritis), susceptible to oxytetracycline.	25 mg/lb body weight	Varies with age & water consumption
	FOR THE CONTROL AND TREATMENT OF THE FOLLOWING DISEASES IN SWINE — Bacterial enteritis caused by <i>Escherichia coli</i> and <i>Salmonella choleraesuis</i> , susceptible to oxytetracycline. Bacterial pneumonia caused by <i>Pasteurella multocida</i> , susceptible to oxytetracycline. FOR BREEDING SWINE: Leptospirosis (reducing the incidence of abortions and shedding of leptospira) caused by <i>Leptospira pomona</i> , susceptible to oxytetracycline.	10 mg/lb body weight	Varies with age & water consumption



OXYTETRACYCLINE HCl AN ANTIBIOTIC

This packet contains 512 grams of oxytetracycline HCl and will make:

2560 gallons (9690 L) containing 200 mg oxytetracycline HCl per gallon

1280 gallons (4845 L) containing 400 mg oxytetracycline HCl per gallon

640 gallons (2422 L) containing 800 mg oxytetracycline HCl per gallon

This packet will treat 51,200 lbs of swine

RECOMMENDED PACKET STORAGE CONDITIONS:

Store below 77° F (25° C)

See back panel for instructions

NADA #130-435, Approved by FDA

For Use in Drinking Water Only
Not for Use in Liquid Type B Medicated Feeds
Keep Out of Reach of Children

Net Weight
3.09 lbs (1400 g)

Restricted Drug. Use Only as Directed (CA)

Take Time



Observe Label Directions

List #2205



Animal Health Division

Alpharma Inc.
 One Executive Drive, Fort Lee, New Jersey 07024

FDA AHF-039 0007

DIRECTIONS FOR USE: Mix fresh solutions daily. Use as sole source of drinking water. Do not mix this product directly with milk or milk replacers. Administer 1 hour before or 2 hours after feeding milk or milk replacers. Consult a poultry diagnostic laboratory or poultry pathologist for diagnosis and advice on dosage for chickens and turkeys. As a generalization, 200 chickens will drink 1 gallon of water per day for each week of age. Turkeys will consume twice that amount. Administer up to 5 days to swine and 7 to 14 days for chickens and turkeys. **NOTE:** The concentration of drug required in medicated water must be adequate to compensate for variation in the age of the animal, feed consumption rate, and the environmental temperatures and humidity, each of which affects water consumption. **CAUTION:** Use as sole source of oxytetracycline. Not to be used for more than 5 consecutive days in swine or 14 consecutive days in chickens and turkeys.

WARNING - Do not feed to birds producing eggs for human consumption. LIVESTOCK REMEDY. NOT FOR HUMAN USE. HAZARDOUS REMEDY.



MIXING INSTRUCTIONS FOR WATER MEDICATION
CHICKENS — TURKEYS — SWINE
DOSAGE AND INDICATIONS FOR USE IN WATER PROPORTIONERS—Add the following amount to one gallon of stock solution when proportioner is set to meter at the rate of one ounce per gallon.

	DISEASE	TREATMENT LEVEL	PACKS/GAL. STOCK SOL.
C H I C K E N S	Control of infectious synovitis caused by <i>Mycoplasma synoviae</i> , susceptible to oxytetracycline.	200 - 400 mg	1/4-1/2 (34-68 g)
	Control of chronic respiratory disease (CRD) and air sac infections caused by <i>Mycoplasma gallisepticum</i> and <i>Escherichia coli</i> , susceptible to oxytetracycline.	400 - 800 mg	1/2-1 (68-135.5 g)
	Control of low cholera caused by <i>Pasteurella multocida</i> , susceptible to oxytetracycline.	400 - 800 mg	1/2-1 (68-135.5 g)
T U R K E Y S	Control of hexamitiasis caused by <i>Hexamita meleagridis</i> susceptible to oxytetracycline.	200 - 400 mg	1/4-1/2 (34-68 g)
	Control of infectious synovitis caused by <i>Mycoplasma synoviae</i> susceptible to oxytetracycline.	400 mg	1/2 (68 g)
S W I N E	GROWING TURKEYS—Control of complicating bacterial organisms associated with bluecomb (transmissible enteritis, coronaviral enteritis) susceptible to oxytetracycline.	25 mg/lb. body weight	Varies with age & water consumption
	FOR THE CONTROL AND TREATMENT OF THE FOLLOWING DISEASES IN SWINE: Bacterial enteritis caused by <i>Escherichia coli</i> and <i>Salmonella choleraesuis</i> , susceptible to oxytetracycline. Bacterial pneumonia caused by <i>Pasteurella multocida</i> , susceptible to oxytetracycline. FOR BREEDING SWINE: Leptospirosis (reducing the incidence of abortions and shedding of leptospiral) caused by <i>Leptospira pomona</i> , susceptible to oxytetracycline.	10 mg/lb. body weight	Varies with age & water consumption

MFD. FOR: AGRI LABORATORIES, LTD.
 St. Joseph, MO 64503

AGRIMYCIN®-343

Oxytetracycline Hydrochloride Soluble Powder

antibiotic for control of specific diseases in poultry and swine

Each packet contains 102.4 grams of oxytetracycline HCl and will make:
 512 gallons (1936 L) containing 200 mg oxytetracycline HCl per gallon
 256 gallons (969 L) containing 400 mg oxytetracycline HCl per gallon
 128 gallons (484 L) containing 800 mg oxytetracycline HCl per gallon

Each packet will treat 10,240 pounds of swine.

ANADA 200-066, Approved by FDA

NET CONTENTS: 50 PACKETS
 Each containing 4.78 oz (135.5 g)

AgriLabs®

If improvement is not noted within 24 to 48 hours, consult a poultry veterinarian or poultry diagnostic laboratory.

DIRECTIONS FOR USE: Mix fresh solutions daily - Use as sole source of drinking water - Do not mix this product directly with milk or milk replacers. Administer one hour before or two hours after feeding milk or milk replacers.

Consult a poultry diagnostic laboratory or poultry pathologist for diagnosis and advice on dosage for chickens and turkeys. As a generalization, 200 chickens will drink one gallon of water per day for each week of age. Turkeys will consume twice that amount. Administer up to 5 days to swine and 7 to 14 days for chickens and turkeys. Note: The concentration of drug required in medicated water must be adequate to compensate for variation in the age of the animals, feed consumption rate, and the environmental temperature and humidity, each of which affects water consumption.

CAUTION: USE AS SOLE SOURCE OF OXYTETRACYCLINE—NOT TO BE USED FOR MORE THAN 5 CONSECUTIVE DAYS IN SWINE AND 14 CONSECUTIVE DAYS IN CHICKENS AND TURKEYS.

WARNING: DO NOT FEED TO BIRDS PRODUCING EGGS FOR HUMAN CONSUMPTION. DISCONTINUE TREATMENT OF SWINE 10 DAYS AND TURKEYS 5 DAYS PRIOR TO SLAUGHTER.

RECOMMENDED STORAGE: STORE BELOW 77°F (25°C) FOR USE IN DRINKING WATER ONLY—NOT FOR USE IN LIQUID FEED SUPPLEMENTS

**FOR USE IN ANIMALS ONLY
 KEEP OUT OF REACH OF CHILDREN**

Rev. 1098

TAKE TIME  OBSERVE LABEL DIRECTIONS

Lot No.:

Exp. Date



DISCONTINUE TREATMENT OF CHICKENS, TURKEYS AND SWINE 0 DAYS PRIOR TO SLAUGHTER.

DISCONTINUE TREATMENT OF CHICKENS,
TURKEYS AND SWINE 0 DAYS PRIOR TO SLAUGHTER.

DIRECTIONS FOR USE: Mix fresh solutions daily - Use as sole source of drinking water - Do not mix this product directly with milk or milk replacers. Administer one hour before or two hours after feeding milk or milk replacers. Administer to all birds in the flock. As a generalization, 200 chickens will consume one gallon of water per day for each week of age. Turkeys will consume twice that amount. Administer up to 5 days to swine and 7 to 14 days for chickens and turkeys. Note: The concentration of drug required in medicated water must be adequate to compensate for variation in the age of the animal, feed consumption rate, and the environmental temperature and humidity, each of which affects water consumption.

CAUTION: USE AS SOLE SOURCE OF OXYTETRACYCLINE - NOT TO BE USED FOR MORE THAN 5 CONSECUTIVE DAYS IN SWINE AND 14 CONSECUTIVE DAYS IN CHICKENS AND TURKEYS.

WARNING: DO NOT FEED TO BIRDS PRODUCING EGGS FOR HUMAN CONSUMPTION. DISCONTINUE TREATMENT OF SWINE 14 DAYS AND TURKEYS 0 DAYS PRIOR TO SLAUGHTER.

RECOMMENDED STORAGE: STORE BELOW 77°F (25°C)
FOR USE IN DRINKING WATER ONLY - NOT FOR USE IN LIQUID FEED SUPPLEMENTS



AGRIMYCIN® 343

Oxytetracycline Hydrochloride Soluble Powder

antibiotic for control of specific diseases in poultry and swine

This packet contains 102.4 grams of Oxytetracycline HCl and will make:
512 gallons (1938 L) containing 200 mg Oxytetracycline HCl per gallon
256 gallons (969 L) containing 400 mg Oxytetracycline HCl per gallon
128 gallons (484 L) containing 800 mg Oxytetracycline HCl per gallon

This packet will treat 10,240 pounds of swine
FOR USE IN ANIMALS ONLY

KEEP OUT OF REACH OF CHILDREN

ANADA 200-066, Approved by FDA

NET WEIGHT: 4.78 oz (135.5g)

Mfg. for: Agri Laboratories, Ltd.
St. Joseph, Missouri • 64503

If improvement is not noted within 24 to 48 hours,
consult a poultry veterinarian or poultry diagnostic laboratory.

DOSE AND INDICATIONS FOR USE IN WATER PROPORTIONERS - Add the following amount to one gallon of fresh solution when proportioner is set to meter at the rate of one pound per gallon.

DISEASE	TREATMENT LEVEL	PACKAGAL STOCK SOL.
Control of infectious enteritis caused by <i>Mycobacterium avium</i> , susceptible to erythromycin.	200-400 mg	16-17 (24-48 g)
Control of chronic respiratory disease (CRD) and air sac infections caused by <i>Mycoplasma gallisepticum</i> and <i>Escherichia coli</i> , susceptible to oxytetracycline.	400-800 mg	17-1 (66-135.5 g)
Control of foot lesions caused by <i>Pasteurella multocida</i> , susceptible to oxytetracycline.	400-800 mg	17-1 (66-135.5 g)
Control of heresmitiasis caused by <i>Neisseria meningitidis</i> , susceptible to oxytetracycline.	200-400 mg	17-1 (24-48 g)
Control of infectious enteritis caused by <i>Mycobacterium avium</i> susceptible to erythromycin.	400 mg	17 (66g)
GROWING TURKEYS - Control of campylobacteriosis associated with <i>Salmonella</i> (transmissible enteritis, <i>Escherichia coli</i>) susceptible to erythromycin.	25 mg/lb. body weight	Varies with age & water consumption
FOR THE CONTROL AND TREATMENT OF THE FOLLOWING DISEASES IN SWINE: Bacterial enteritis caused by <i>Escherichia coli</i> and <i>Salmonella choleraesuis</i> , susceptible to oxytetracycline. Bacterial pneumonia caused by <i>Streptococcus pneumoniae</i> , susceptible to erythromycin. Bacterial meningitis caused by <i>Streptococcus pneumoniae</i> , susceptible to erythromycin. Bacterial meningitis caused by <i>Escherichia coli</i> and <i>Streptococcus pneumoniae</i> , susceptible to erythromycin. Bacterial meningitis caused by <i>Streptococcus pneumoniae</i> , susceptible to erythromycin.	10 mg/lb. body weight	Varies with age & water consumption



If improvement is not noted within 24 to 48 hours, consult a poultry veterinarian or poultry diagnostic laboratory.

MIXING INSTRUCTIONS FOR WATER MEDICATION
CHICKENS—TURKEYS—SWINE
DOSAGE AND INDICATIONS FOR USE IN WATER
PROPORTIONERS—Use the enclosed scoop to add the following amount to one gallon of stock solution when proportioner is set to meter at the rate of one ounce per gallon. One level scoop is equal to approximately 34 grams.

DISEASE	TREATMENT LEVEL	SCOOPS/GAL. STOCK SOL.
C Control of infectious synovitis caused by <i>Mycoplasma synoviae</i> susceptible to oxytetracycline	200 - 400 mg	1-2 (34-68 g)
H Control of chronic respiratory disease (CRD) and air sac infections caused by <i>Mycoplasma paratuberculosis</i> and <i>Escherichia coli</i> susceptible to oxytetracycline	400 - 800 mg	2-4 (68-135.5 g)
I Control of low cholera caused by <i>Pasteurella multocida</i> susceptible to oxytetracycline	400 - 800 mg	2-4 (68-135.5 g)
K Control of hexamitiasis caused by <i>Hexamita meleagridis</i> susceptible to oxytetracycline	200 - 400 mg	1-2 (34-68 g)
E Control of infectious synovitis caused by <i>Mycoplasma synoviae</i> susceptible to oxytetracycline	400 mg	2 (68 g)
T GROWING TURKEYS —Control of complicating bacterial organisms associated with bluecomb (transmissible enteritis, coronavirus enteritis) susceptible to oxytetracycline.	25 mg/lb. body weight	Varies with age & water consumption
U FOR THE CONTROL AND TREATMENT OF THE FOLLOWING DISEASES IN SWINE: Bacterial enteritis caused by <i>Escherichia coli</i> and <i>Salmonella choleraesuis</i> , susceptible to oxytetracycline. Bacterial pneumonia caused by <i>Pasteurella multocida</i> susceptible to oxytetracycline. FOR BREEDING SWINE: Leptospirosis (reducing the incidence of abortions and shedding of leptospira) caused by <i>Leptospira pomona</i> , susceptible to oxytetracycline.	10 mg/lb. body weight	Varies with age & water consumption

MFD. FOR AGRI LABORATORIES, LTD.
 St. Joseph, MO 64503

AGRIMYCIN®-343

Oxytetracycline Hydrochloride Soluble Powder

antibiotic for control of specific diseases in poultry and swine

Each pail contains 1.715 kilograms of oxytetracycline HCl and will make:
 8.575 gallons (32.460 L) containing 200 mg oxytetracycline HCl per gallon
 4.288 gallons (16.230 L) containing 400 mg oxytetracycline HCl per gallon
 2.144 gallons (8.115 L) containing 800 mg oxytetracycline HCl per gallon

Each pail will treat 171,500 pounds of swine.

ANADA 200-066, Approved by FDA

NET WEIGHT:
5 lb (2.27 kg)

AgriLabs®

DIRECTIONS FOR USE: Mix fresh solutions daily - Use as sole source of drinking water - Do not mix this product directly with milk or milk replacers. Administer one hour before or two hours after feeding.

Consult a poultry diagnostic laboratory or poultry pathologist for diagnosis and advice on dosage for chickens and turkeys. As a generalization, 200 chickens will drink one gallon of water per day for each week of age. Turkeys will consume twice that amount. Administer up to 5 days to swine and 7 to 14 days for chickens and turkeys.

Note: The concentration of drug required in medicated water must be adequate to compensate for variation in the age of the animals, feed consumption rate, and the environmental temperature and humidity, each of which affects water consumption.

CAUTION: USE AS SOLE SOURCE OF OXYTETRACYCLINE—NOT TO BE USED FOR MORE THAN 5 CONSECUTIVE DAYS IN SWINE AND 14 CONSECUTIVE DAYS IN CHICKENS AND TURKEYS.

WARNING: DO NOT FEED TO BIRDS PRODUCING EGGS FOR HUMAN CONSUMPTION. DISCONTINUE TREATMENT OF SWINE 10 DAYS AND TURKEYS 6 DAYS PRIOR TO SLAUGHTER.

RECOMMENDED STORAGE: STORE BELOW 77°F (25°C) FOR USE IN DRINKING WATER ONLY. NOT FOR USE IN LIQUID FEED SUPPLEMENTS

FOR USE IN ANIMALS ONLY
KEEP OUT OF REACH OF CHILDREN

1098

TAKE TIME TO OBSERVE LABEL DIRECTIONS



Lot No.

Exp. Date:

DISCONTINUE TREATMENT OF CHICKENS, TURKEYS AND SWINE 0 DAYS PRIOR TO SLAUGHTER.