

4-26-83

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

OFFICE OF
PESTICIDES AND TOXIC SUBSTANCES

~~MAY 6 1983~~

April 26, 1983

SUBJECT: PP#2F2707/2H5355: "Free Standing" Summary;
Larvadex (Cyromazine) in Eggs, Meat, Fat and Meat
Byproducts of Poultry.

Caswell #167R

WAB

THRU: William Butler, Section Head #3
Toxicology Branch (TS-769)

William Butler
#-26-83

TO: Timothy A. Gardner
Registration Division (TS-767)

Tolerances Requested: 0.4 ppm in eggs, meat, fat and meat
byproducts of poultry.

Recommendation: Based upon the existing toxicity data base,
the requested permanent tolerances can be supported.

1. Summary of Data Reviewed for the previous temporary
tolerances:

- Oral LD50 rat - 3387 mg/kg
- Oral LD50 mouse - 2029 mg/kg
- Oral LD50 rabbit - 1467 mg/kg
- Acute dermal LD50 rat >3100 mg/kg
- Sensitization, guinea pig - non-sensitizing
- Ames test - negative
- 90 day rat feeding NOEL 30 ppm
- Teratology rat - NOEL teratology >600 mg/kg
- NOEL-fetotoxicity 300 mg/kg; LEL = 600 mg/kg
- 6 month dog - NOEL 30 ppm

2. New Data submitted for the requested permanent tolerances:

- Teratology rabbit - NOEL >75 mg/kg/day
- 2 generation reproduction in rats - reproductive NOEL
1000 ppm
- Mutagenicity - Dominant Lethal - mice negative
- Nucleus anomaly in Chinese Hamster - negative
- 2 year chronic and oncogenicity in rat-
Oncogenicity NOEL - >3000 ppm
- Systemic NOEL - 30 ppm
- Oncogenicity in mice - oncogenicity NOEL >3000 ppm
- Systemic NOEL - 50 ppm

3. No data gaps.
4. Only the following temporary tolerances existed:

Poultry	0.200	ppm
Eggs	0.200	ppm
Cattle	0.100	ppm
Hogs	0.100	ppm
Sheep	0.100	ppm

5. Data to Set Tolerances:

Study No. 382-081, a two year feeding and oncogenicity study in the rat can be used as a basis for the ADI. The NOEL of 30 ppm (1.5 mg/kg) in this study and a safety factor of 100 yields an ADI of 0.015 mg/kg/day and an MPI of 0.90 mg/day (60 kg).

6. A previous TOX approved action, 9G2230, produced a TMRC of 0.0334 mg/day (1.5 kg) which occupied 3.71% of the ADI and contained some of the same RAC's as requested in the current action.

The present action increases the percentage of the ADI occupied from 3.71% to 5.61% and the TMRC from 0.0334 mg/day (1.5 kg) to 0.0505 mg/day (1.5 kg).

7. There are no pending RPAR's against larvadex.
8. There are no other relevant considerations in setting the tolerance.

Stephanie P. April, Ph.D.

4/24/83

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