DATA EVALUATION RECORD - SUPPLEMENT

XDE-570 (FLORASULAM)

Study Type: OPPTS 870.4100b [§83-1b]; Chronic Toxicity in Dogs

Work Assignment No. 4-1-128 G (MRID 46808229)

Prepared for
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Disclaimer

This Data Evaluation Record my have been altered by the Health Effects Division subsequent to signing by Dynamac Corporation personnel

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See TXR # 0054348 for previous DER

This supplement contains:

- New cover page
- New executive summary

STUDY TYPE: Chronic toxicity - dog [feeding]; OPPTS 870.4100b [' 83-1b]; OECD 452.

PC CODE: 129108 **DP BARCODE:** D331116
TXR#: 0054348

TEST MATERIAL (PURITY): XDE-570 (Florasulam; 99.3% a.i.)

SYNONYMS: *N*-(2,6-Difluorophenyl)-8-fluoro-5-methoxy(1,2,4)triazolo(1,5-*c*)pyrimidine-2-sulfonamide; XR-570; XRD-570; DE-570

CITATION: Stebbins, K. E., and K. T. Haut (1997) XDE-570: one year dietary toxicity study in beagle. The Toxicology Research Laboratory, Health and Environmental Sciences, The Dow Chemical Company, Midland, MI. Laboratory Project Study ID: 960018, October 30, 1997. MRID 46808229. Unpublished.

SPONSOR: Dow AgroSciences Canada, Inc., 2100-450 1 St. SW, Calgary, AB, Canada

EXECUTIVE SUMMARY: In a chronic toxicity study (MRID 46808229), XDE-570 (Florasulam; 99.3% a.i.; Lot No. 940714) was administered in the diet to four purebred beagle dogs/sex/dose at dose levels of 0, 0.5, 5, or 100 mg/kg/day for 52 weeks. Severe body weight loss and reduced food consumption were observed in both sexes at 100 mg/kg/day during the first three months of the study; therefore, the high dose was reduced to 50 mg/kg/day in both sexes beginning on Study Day 105 (Week 15).

No adverse treatment-related effects were observed on mortality, clinical signs, food efficiency, ophthalmoscopic examinations, hematology, urinalysis, organ weights, or gross or microscopic pathology.

At 100 mg/kg/day, both sexes exhibited loss of body weight accompanied by reduced food consumption. Following reduction of the high dose to 50 mg/kg/day, the females continued to exhibit both decreased (not significant [NS]) body weights (decr. 17% at Week 52) and food consumption, resulting in decreased (NS) overall (Week 0-52) body weight gains (decr. 68%).

Male body weights and food consumption at Week 52, and overall body weight gains were similar to controls.

Additionally at 100 mg/kg/day, males and females had increased ($p<=0.05$) alkaline phosphatase (incr. 233-783%) and alanine aminotransferase (incr. 268-390%) after 3 months of dosing. Alkaline phosphatase continued to be elevated ($p<=0.05$) in both sexes through 12 months of dosing (incr. 141-354%). No corroborating findings were observed in organ weights, or gross or microscopic pathology; therefore, these findings were considered equivocal.

The LOAEL is 100/50 mg/kg/day, based on decreased body weights, body weight gains, and food consumption in the females. The NOAEL is 5 mg/kg/day.

This study is classified as **acceptable/guideline** and satisfies the guideline requirements (OPPTS 870.4100b, OECD 452) for a chronic toxicity study in the dog.

COMPLIANCE: Signed and dated GLP Compliance, Quality Assurance, and Data Confidentiality statements were provided.