

Supplement to HED Document No. 000065 & 007865 - DER for Accession No. 097233 (amendment MRID No. 41026604): 3-Generation Reproduction Study in Rats. This supplement provides an Executive Summary to upgrade the original DER.

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Reregistration Branch 4 (7509C)
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DATA EVALUATION RECORD

STUDY TYPE: Reproduction Study - Rat; OPPTS 870.3800 [§83-4]

DP BARCODE: D262725
P.C. CODE: 111901

SUBMISSION CODE: S548748
TOX. CHEM. NO.: 497AB

TEST MATERIAL (PURITY): Imazalil technical (98.8%)

SYNONYMS: R23979; Fungaflor Technical; 1-[2-(2,4-dichlorophenyl)-2-(2-propenyloxy)ethyl]-1H-imidazole

CITATION: Anonymous, 1978. Oral Three Generation Study in Wistar Rats. Study number 7336. Janssen Pharmaceutica Research Laboratories, 2340 Beerse, Belgium. March 3, 1978. Accession No. 097233 (Amendment MRID No. 41026604). Unpublished.

SPONSOR: Janssen Pharmaceutica N.V., 2340 Beerse, Belgium

EXECUTIVE SUMMARY: In a 3-generation reproduction study (Accession No. 097233 and amendment MRID 41026604) imazalil (98.8%) was administered in the diet to 10 male and 20 female Wistar rats per dose at approximately 0, 50, 200 or 800 ppm at the time of mating and continuously thereafter. The mating ratio was 1:2 for the F0 and 1:3 for the subsequent generations. Two litters per generation were produced. During the first mating, only 40% of the high dose females delivered viable litters. However, this finding was considered incidental since 100% of the dams of this dose group produced a normal number of litters from the second mating.

No treatment related effects were noted in the parental as well as developmental parameters measured. These included body weight of dams, food consumption, fertility, number of litters, litter size, pup viability, pup body weight or malformations, with the exception of the results of the first mating in the first generation at the high dose.

LOAEL for parental (systemic and reproductive) as well as developmental toxicity is > 800 ppm based on lack of effect in parameters measured. The NOAEL is ≥800 ppm.

The original review of the study (HED # 000065) classified this study as core minimum. Reevaluation of the study (HED # 007865) determined the study contained many deficiencies and was downgraded to **supplementary**. The study was not conducted under GLP. No test chemical stability data was reported. No analyses were reported of the test material in the diets administered. Weekly body weights of males and females were reported only for the third week of gestation for the three generations. Food consumption was reported only for the same periods and for lactation. No necropsy data were reported for the parents of any generation. Pup necropsies were conducted on the second litters of the third generation only. Dosing did not commence prior to mating. Therefore, the study is **unacceptable** for OPPTS 870.3800 (83-4) guideline on reproduction and fertility.

COMPLIANCE: GLP, Quality Assurance statements were not provided. This study predates GLP requirements.

IMAZALIL

Reproduction Study OPPTS 870.3800 (S83-4)

SignOff Date:	2/8/00
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