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8EHQ-91-1303
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ORIGINAL
8EHQ-0894-1303
August 21, 1994



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Attention: Section 8(e) Coordinator
Office of Pollution Prevention and Toxics
U.S. Environmental Protection Agency
401 M Street, S.W.
Washington, D.C. 20460

Dear Coordinator:

(cymoxanil)

TSCA 8(e): [REDACTED]

This letter informs you of findings in a set of tests done in Japan to meet Japan-specific requirements for the above referenced material. The study was conducted in mice, rats, guinea pigs and rabbits to assess the general pharmacology of the test compound and included doses which were lethal as well as doses without effect. Doses were delivered both by gavage and intravenous administration. Effects which may be reportable were seen in the tests involving mice and rabbits. These are described below.

1. Effects on the central nervous system:

In the general behavior test in mice, orally administered doses of 30 and 100 mg/kg were without effect. At 300 mg/kg a slight, transient, decreased reactivity and spontaneous motor activity (SMA) were apparent at only 1 hr. after dosing. A dose of 1000 mg/kg caused passivity and decreases in alertness, visual placing, grooming, reactivity, SMA, touch response and in body temperature; 2 of 3 mice died within 2 hours of administration, but the surviving mouse had recovered by 4 hours post dosing and showed no toxicologic signs at 24 hours post dosing.

Oral administration of the test material to mice at a dose of 300mg/kg prolonged hexobarbital-induced sleep time and caused a synergistic effect on electrically-induced convulsions.

2. Effects on the respiratory and cardiovascular system:

Intravenously administered test compound given to anesthetized rabbits at a dose of 10 mg/kg caused decreased blood pressure and increased respiratory rate and respiratory flow. Doses of 0.1 and 1.0 mg/kg caused no effects on respiration, blood pressure, heart rate, or ECG.

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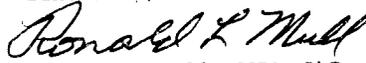
Document Processing Center (TS-790)
August 11, 1994
Page -2-

3. Effects on the gastrointestinal system:

Orally administered doses of 300 mg/kg to mice caused significant decreases in intestinal transport in mice and death of 2 of 8 within one hr. of dosing. Doses of 30 and 100 mg/kg had no effect on intestinal transport.

Under these test conditions, these effects may be reportable based on the EPA guidance regarding TSCA section 8 (e) criteria.

Sincerely,



Ronald L. Mull, DVM. PhD
Regulatory Toxicologist

RLM/ckz