This IRIS Summary has been removed from the IRIS database and is available for historical reference purposes. (July 2016)

Diphenylamine; CASRN 122-39-4

Health assessment information on a chemical substance is included in IRIS Human health assessment information on a chemical substance is included in the IRIS database only after a comprehensive review of toxicity data, as outlined in the IRIS assessment development process. Sections I (Health Hazard Assessments for Noncarcinogenic Effects) and II (Carcinogenicity Assessment for Lifetime Exposure) present the conclusions that were reached during the assessment development process. Supporting information and explanations of the methods used to derive the values given in IRIS are provided in the guidance documents located on the IRIS website.

STATUS OF DATA FOR Diphenylamine

File First On-Line 01/31/1987

<table>
<thead>
<tr>
<th>Category (section)</th>
<th>Assessment Available?</th>
<th>Last Revised</th>
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<tbody>
<tr>
<td>Oral RfD (I.A.)</td>
<td>yes</td>
<td>01/31/1987</td>
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<tr>
<td>Inhalation RfC (I.B.)</td>
<td>not evaluated</td>
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<tr>
<td>Carcinogenicity Assessment (II.)</td>
<td>not evaluated</td>
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I. Chronic Health Hazard Assessments for Noncarcinogenic Effects

I.A. Reference Dose for Chronic Oral Exposure (RfD)

Substance Name — Diphenylamine
CASRN — 122-39-4
Last Revised — 01/31/1987

The oral Reference Dose (RfD) is based on the assumption that thresholds exist for certain toxic effects such as cellular necrosis. It is expressed in units of mg/kg-day. In general, the RfD is an estimate (with uncertainty spanning perhaps an order of magnitude) of a daily exposure to the human population (including sensitive subgroups) that is likely to be without an appreciable risk of deleterious effects during a lifetime. Please refer to the Background Document for an
elaboration of these concepts. RfDs can also be derived for the noncarcinogenic health effects of substances that are also carcinogens. Therefore, it is essential to refer to other sources of information concerning the carcinogenicity of this substance. If the U.S. EPA has evaluated this substance for potential human carcinogenicity, a summary of that evaluation will be contained in Section II of this file.

I.A.1. Oral RfD Summary

<table>
<thead>
<tr>
<th>Critical Effect</th>
<th>Experimental Doses*</th>
<th>UF</th>
<th>MF</th>
<th>RfD</th>
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<tbody>
<tr>
<td>Decreased body weight gain, and increased liver and kidney weights</td>
<td>NOEL: 0.01% of diet (2.5 mg/kg/day)</td>
<td>100</td>
<td>1</td>
<td>2.5E-2 mg/kg/day</td>
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<tr>
<td>2-Year Dog Feeding Study</td>
<td>LEL: 0.1% of diet (25 mg/kg/day)</td>
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<td></td>
<td></td>
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<tr>
<td>Thomas et. al., 1967</td>
<td></td>
<td></td>
<td></td>
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</table>

*Conversion Factors -- 0.01% (1 ppm) = 0.025 mg/kg/day (assumed dog food consumption)

I.A.2. Principal and Supporting Studies (Oral RfD)


Eight male and 8 female purebred beagles were fed 0.01, 0.1, and 1% diphenylamine in the diet for 2 years. Decreased weight gain and anemia were noted at 0.1 and 1%. At 1%, increases in liver and kidney weights were observed.

I.A.3. Uncertainty and Modifying Factors (Oral RfD)

UF — An uncertainty factor of 100 was used to account for inter- and intra- species differences.

MF — None

I.A.4. Additional Studies/Comments (Oral RfD)
Data Considered for Establishing the RfD

1) 2-Year Feeding - Dog: Principal study - see discussion above

2) 2-Year Feeding (oncogenic) - rat: NOEL=3.1 mg/kg/day; LEL=31 mg/kg/day (kidney lesions) (Chemley Products, 1967)

3) 2-Generation Reproduction - rat: Fetotoxic NOEL=125 mg/kg/day; Fetotoxic LEL=250 mg/kg/day (reduced litter size and weight of young) (Food and Agriculture Organization of the United Nations, 1970)

Data Gap(s): Teratology - Rat; Teratology - Rabbit

I.A.5. Confidence in the Oral RfD

Study — Medium
Database — Medium
RfD — Medium

The principal study appears to be of good quality and is given a medium confidence rating. Since the database on chronic toxicity is supportive but incomplete, it is given a medium confidence rating. Medium confidence in the RfD follows.

I.A.6. EPA Documentation and Review of the Oral RfD

Source Document — This assessment is not presented in any existing U.S. EPA document.

Other EPA Documentation — Pesticide Registration Files

Agency Work Group Review — 07/22/1986

Verification Date — 07/22/1986

Screening-Level Literature Review Findings — A screening-level review conducted by an EPA contractor of the more recent toxicology literature pertinent to the RfD for Diphenylamine conducted in August 2003 identified one or more significant new studies. IRIS users may request the references for those studies from the IRIS Hotline at hotline.iris@epa.gov or 202-566-1676.

I.A.7. EPA Contacts (Oral RfD)
Please contact the IRIS Hotline for all questions concerning this assessment or IRIS, in general, at (202)566-1676 (phone), (202)566-1749 (FAX) or hotline.iris@epa.gov (internet address).

I.B. Reference Concentration for Chronic Inhalation Exposure (RfC)

Substance Name — Diphenylamine
CASRN — 122-39-4

Not available at this time.

II. Carcinogenicity Assessment for Lifetime Exposure

Substance Name — Diphenylamine
CASRN — 122-39-4

Not available at this time.

III. [reserved]
IV. [reserved]
V. [reserved]

VI. Bibliography

Substance Name — Diphenylamine
CASRN — 122-39-4

VI.A. Oral RfD References


VI.B. Inhalation RfC References

None

VI.C. Carcinogenicity Assessment References

None

VII. Revision History

Substance Name — Diphenylamine
CASRN — 122-39-4

<table>
<thead>
<tr>
<th>Date</th>
<th>Section</th>
<th>Description</th>
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<td>10/28/2003</td>
<td>I.A.6.</td>
<td>Screening-Level Literature Review Findings message has been added.</td>
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VIII. Synonyms

Substance Name — Diphenylamine
CASRN — 122-39-4
Last Revised — 01/31/1987

- 122-39-4
- Aniline, N-phenyl-
- Anilinobenzene
- Benzenamine, N-phenyl-
- Benzene, anilino-
• Benzene, (phenylamino)-
• Big Dipper
• C.I. 10355
• DFA
• Diphenylamine
• DPA
• N,N-Diphenylamine
• No Scald
• N-Phenylaniline
• N-Phenylbenzenamine
• Scaldip