



Canadian Environmental Protection Act

Priority Substances List
Assessment Report No. 3

Chlorobenzene



Government
of Canada

Gouvernement
du Canada

Environment
Canada

Environnement
Canada

Health
Canada

Santé
Canada



**PRIORITY SUBSTANCES LIST
ASSESSMENT REPORT NO. 3**

CHLOROBENZENE

Government of Canada
Health and Welfare Canada
Environment Canada

Also available in French
under the title: *Loi canadienne
sur la protection de l'environnement,
Liste des substances d'intérêt prioritaire,
Rapport d'évaluation n° 3:
Chlorobenzène*

©Minister of Supply and Services Canada 1992
Cat. No. EN40-215/3E
ISBN 0-662-19949-9



Printed on
Recycled Paper

TABLE OF CONTENTS

Overview of Findings	v
1.0 Introduction.....	1
2.0 Summary of Critical Supporting Data.....	3
2.1 Identity and Physical/Chemical Properties	3
2.2 Production and Uses.....	3
2.3 Sources and Releases	3
2.4 Environmental Fate and Concentrations	4
2.4.1 Fate.....	4
2.4.2 Concentrations.....	4
2.5 Toxicokinetics and Metabolism.....	6
2.6 Mammalian Toxicology.....	6
2.7 Effects on Humans	8
2.8 Effects on the Environment.....	8
3.0 Assessment of "Toxic" under CEPA.....	10
3.1 Entry.....	10
3.2 Exposure.....	10
3.3 Effects	12
3.3.1 Human Health	12
3.3.2 Environment.....	13
3.4 Conclusions	13
3.4.1 Paragraph 11(a) -- Effects on the Environment	13
3.4.2 Paragraph 11(b) -- Effects on the Environment on which Human Life Depends.....	14
3.4.3 Paragraph 11(c) -- Effects on the Human Life or Health.....	14
3.4.4 General Conclusions	14
4.0 Recommendations for Research	15
5.0 References.....	16

Overview of Findings

Chlorobenzene (also referred to as monochlorobenzene (MCB), the term most often used by the scientific community and throughout this report) is used in Canada in a variety of ways that lead to the direct (via pesticide application) and indirect (via effluents, emissions, and leachate waters) entry of this substance into the Canadian environment. These releases result in measurable or predicted concentrations of monochlorobenzene in the various media to which humans and other organisms may be exposed, albeit at low levels.

Concentrations predicted in surface water are lower by six orders of magnitude than those which induce adverse effects in the most sensitive aquatic species, the largemouth bass, following long-term exposure. The average level found in a study of raw effluent from selected organic chemical manufacturing facilities is six times lower than this effect level while the highest concentrations recorded in these raw effluent samples were 80 times higher than the level found to be acutely lethal to aquatic invertebrates.

Studies on effects of monochlorobenzene following short- or long-term exposure of wildlife were not identified. However, the effect levels reported in inhalation studies conducted in laboratory animals are considered relevant to wild mammals. The highest measured airborne concentration in Canada to which wild mammals may be exposed is more than five orders of magnitude less than the lowest reported effect level in the longest term inhalation study in laboratory animals.

Because of its short persistence in the atmosphere, and relatively low rate of release, monochlorobenzene is not associated with ozone layer depletion. Also, the magnitude of any potential indirect effects of monochlorobenzene on global warming and photochemical smog formation, although difficult to quantify, is not believed to be significant.

Data on concentrations of monochlorobenzene to which humans are exposed in food are limited, though it is likely that intake from this source is negligible compared to that from air. Based on data on concentrations of monochlorobenzene in air and drinking water, total average daily intakes of monochlorobenzene for various age groups in the general population have been estimated. These estimated intakes are much less than (from approximately 60 to 175 times) the intake to which it is believed that a person can be exposed over a lifetime without deleterious effect (i.e., the Tolerable Daily Intake derived on the basis of the most relevant study in laboratory animals exposed by the most appropriate route).

Based on these considerations, the Ministers of Environment Canada and of Health and Welfare Canada have concluded that concentrations of monochlorobenzene present in the environment do not constitute a danger in Canada to the environment or to the environment on which human life depends or to human life or health. Therefore, monochlorobenzene is not considered to be "toxic" as defined under section 11 of the *Canadian Environmental Protection Act (CEPA)*.

1.0 Introduction

CEPA requires the federal Ministers of Environment Canada and of Health and Welfare Canada to prepare and publish a Priority Substances List that identifies substances, including chemicals, groups of chemicals, effluents and wastes which may be harmful to the environment or constitute a danger to human health. The Act also requires both Ministers to assess these substances and determine whether they are "toxic" as interpreted in section 11 of the Act which states:

"[...] a substance is toxic if it is entering or may enter the environment in a quantity or concentration or under conditions

- (a) having or that may have an immediate or long-term harmful effect on the environment;
- (b) constituting or that may constitute a danger to the environment on which human life depends; or
- (c) constituting or that may constitute a danger in Canada to human life or health."

Substances which are assessed to be "toxic" according to this section may be placed on Schedule I of the Act, and considered for possible development of regulations to control any aspect of their life cycle, from the research and development stage through manufacture, use, storage, transport and ultimate disposal.

The assessment of whether monochlorobenzene is "toxic", as interpreted in CEPA, was based on the determination of whether it **enters** or may enter the Canadian environment in a concentration or quantities or conditions that could lead to **exposure** of humans or other biota to the degree where adverse **effects** could result.

The assessment of whether monochlorobenzene is "toxic" to human health under CEPA, is based principally on documentation prepared by staff of Health and Welfare Canada (HWC) for the International Programme on Chemical Safety (IPCS). Between 1984 and 1987, original data relevant to the assessment of risks to health associated with exposure to the chlorinated benzenes (excluding hexachlorobenzene) were reviewed by staff of Health and Welfare Canada in the preparation of a draft IPCS Environmental Health Criteria Document (EHC). The current assessment has been updated and expanded to emphasize data most relevant to the assessment of the risks associated with exposure of Canadians to monochlorobenzene in the general environment.

In preparation of the World Health Organization (WHO)-IPCS document, a wide variety of scientific databases were searched to update information provided in earlier contractors' reports, including an annotated bibliography on the monochlorobenzenes (excluding hexachlorobenzene) by Peter Strahlendorf (1978), and a criteria document on monochlorobenzenes (including hexachlorobenzene) by Michael Holliday and Associates (1984a; 1984b). Additional information was identified during peer review of the draft Environmental Health Criteria Document by IPCS focal points and a task group of experts which met in June 1990. More recently, in February 1991, a search of Enviroline, Chemical Abstracts, Pollution Abstracts, Environmental Bibliography, IRIS, MEDLINE and BIOSIS databases to identify recent data relevant to assessment in particular, of the risks to Canadians, was conducted

Data relevant to assessment of whether MCB is "toxic" to the environment were identified through the evaluation of existing review documents, supplemented with information from published reference texts and literature identified through on-line searches of commercial databases (ASFA, BIOSIS, CAB Abstracts, Chemical Abstracts, CESARS, CIS, Enviroline, Hazardous Substances Database and IRPTC) conducted in November 1990. Although much of the research on monochlorobenzene has been conducted outside of Canada, data on sources, use patterns, fate and effects of monochlorobenzene on the Canadian environment were emphasized, where available.

Data relevant to assessment of whether MCB is "toxic" to human health obtained after completion of these sections of the report (i.e., May 1991) were not considered for inclusion. Similarly, data relevant to assessment of whether MCB is "toxic" to the environment obtained after February 1992 have not been incorporated.

Although review articles were consulted where considered appropriate, all original studies which form the basis for the determination of "toxic" under CEPA have been critically evaluated by the following staff of the Departments of National Health and Welfare (effects on human health) and of the Environment (effects on the environment):

B. Elliott (Environment Canada)
C. Fortin (Environment Canada)
M. Giddings (Health and Welfare Canada)
K. Lloyd (Environment Canada)
M.E. Meek (Health and Welfare Canada)

In this report, a brief summary of the conclusions which will appear in the *Canada Gazette* is presented. In addition, an extended summary of the technical information which is critical to the assessment, and which is included in greater detail in a Supporting Document, is presented in section 2. The assessment of whether monochlorobenzene is "toxic" under CEPA is presented in section 3.

Sections of the Supporting Document related to the assessment of environmental effects were peer reviewed by a number of experts from Environment Canada and Agriculture Canada and by Dr. Barry Oliver from Zenon Environmental Laboratories (Burnaby, B.C.). The sections related to assessment of effects on human health were approved by the Standards and Guidelines Rulings Committee of the Bureau of Chemical Hazards of Health and Welfare Canada. The Assessment Report was reviewed and approved by Environment Canada/Health and Welfare Canada CEPA Management Committee.

Copies of this Assessment Report and the unpublished Supporting Document are available upon request from:

Environmental Health Centre
Room 104
Health and Welfare Canada
Tunney's Pasture
Ottawa, Ontario, Canada
K1A 0L2

Commercial Chemicals Branch
Environment Canada
14th Floor, Place Vincent Massey
351 Saint-Joseph Boulevard
Hull, Quebec, Canada
K1A 0H3

2.0 Summary of Critical Supporting Data

2.1 Identity and Physical/Chemical Properties

Monochlorobenzene (CAS No. 108-90-7) is a monocyclic aromatic compound with one hydrogen atom on the benzene ring substituted with one chlorine. It is produced by chlorination of benzene in the liquid phase with a catalyst. Monochlorobenzene is a colourless liquid at ambient temperature with a relatively high vapour pressure (1 573.2 Pa), moderate octanol-water partition coefficient (log 2.8) and moderate to low water solubility (497.9 mg/L @ 25°C). Technical grade monochlorobenzene is typically 99% pure with < 0.05% benzene and < 0.1% dichlorobenzenes as contaminants. Analytical methods used to quantify monochlorobenzene in environmental media include gas chromatography/mass spectrometry with flame ionization or electron capture detection.

2.2 Production and Uses

Based on the results of a survey of commercial uses, monochlorobenzene is not produced in Canada (Camford, 1991). Over the last five years, imports of monochlorobenzene into Canada averaged 63 000 kg/year from sources in the United States, West Germany and the United Kingdom. Canadian demand for monochlorobenzene was reported to have remained steady for the last five years, averaging approximately 60 000 kg/year. For 1990, the latest year for which data were available, the demand was 50 000 kg broken down as follows: 29 000 kg as a carrier in pesticides; 20 000 kg in the formulation of rubber polymers; and 1 000 kg as a carrier for textile dyes. Based on the results of the Camford survey, demand over the next five years is expected to be similar. At the time of writing this report, there was only one pesticide in Canada (under temporary registration) in which monochlorobenzene is used as a carrier solvent.

2.3 Sources and Releases

Based on the quantities of monochlorobenzene reported to be used as a carrier solvent for pesticides, and considering that all the monochlorobenzene used for this purpose is released directly onto soil during application, this represents the largest single source of release of monochlorobenzene to the Canadian environment. Quantitative data on other sources of release in Canada have not been identified; such sources are believed to include effluents from certain organic chemical manufacturing facilities, and leachate from disposal in landfill sites (Camford, 1991; OME, 1992; Jackson *et al.*, 1985, 1991; Reinhard *et al.*, 1984).

Another potential source identified, but not confirmed in Canada, is emissions from waste incinerators. The formation and emission of monochlorobenzene as a product of incomplete combustion has been documented in pilot- and full-scale tests with hazardous waste incinerators and co-fired boilers in the United States (U.S. EPA, 1991). Monochlorobenzene was one of the most prevalent products of incomplete combustion emitted in at least 75% of all tests, with a geometric mean emission rate of 494.8 µg/min from hazardous waste incinerators. For full-scale boilers, a geometric mean emission rate of 2 732.8 µg/min was reported. Monochlorobenzene is stable at temperatures up to 700°C as a pure compound, and up to 900°C when in a mixture with other chlorinated compounds (Graham *et al.*, 1986).

Monochlorobenzene is included in the group of substances termed "Volatile Organic Compounds" (VOCs). Currently, there are initiatives by the Federal and Provincial Governments directed at limiting the release of these substances from industrial and other sources as the means of controlling ground-level ozone (CCME, 1990).

2.4 Environmental Fate and Concentrations

2.4.1 Fate

Based on its relatively high vapour pressure and moderate to low water solubility, it is likely that the atmosphere plays an important role in the distribution and ultimate fate of monochlorobenzene (Mackay *et al.*, 1979; Mackay and Shiu, 1990; Garrison and Hill, 1972; Callahan *et al.*, 1979; Mackay and Yeun, 1983; Thomas, 1982). Once released to the atmosphere, either directly or by volatilization from other media, monochlorobenzene is expected to photooxidize relatively quickly in a reaction with hydroxyl radicals producing phenols and their subsequent degradation products (Atkinson *et al.*, 1985). It has been reported that 18.5% of the airborne monochlorobenzene irradiated over a 17-hour period was oxidized by this reaction (Singh *et al.*, 1983; Atkinson *et al.*, 1985). The hydroxyl radical consumed by this reaction may impact negatively on the atmospheric fate of a number of greenhouse gases since it represents an important sink for these substances (IPCC, 1990). Photolysis, another prevalent atmospheric process, is considered an unlikely transformation process because monochlorobenzene does not absorb in the range of wavelengths reaching the surface of the earth (Dulin *et al.*, 1986).

The bulk of the monochlorobenzene released onto soil during, for example, pesticide application, is expected to volatilize, either directly or subsequently, to the atmosphere. The one monochlorobenzene-containing pesticide under temporary registration in Canada is used on dry bulb onions which are typically grown in rich organic soils (muck soils). No studies have been found concerning the fate of monochlorobenzene in these soils. Based on its organic carbon sorption coefficient (K_{oc} of 389) [Schwarzenbach and Westall, 1981], the soil mobility potential of monochlorobenzene is medium (McCall *et al.*, 1981). It was reported that monochlorobenzene volatilizes readily from sandy soil (0.15% organic matter) [Wilson *et al.*, 1981].

On the basis of several studies in which the behaviour of monochlorobenzene in the aquatic environment has been examined, it has been concluded that none of microbiological degradation, hydrolysis or photolysis were significant transformation processes (Morrison and Boyd, 1987; Ellington *et al.*, 1988; Dulin *et al.*, 1986; Lee and Ryan, 1979). Furthermore, monochlorobenzene was not found to bioconcentrate in significant quantities in aquatic biota. Bioconcentration factors of 70 and 50 have been reported for exposure periods of 72 and 24 hours for the golden ide (*Leuciscus idus melanotus*) and the green alga (*Chlorella*) respectively (Freitag *et al.*, 1985).

2.4.2 Concentrations

The presence of monochlorobenzene in atmospheric samples from urban, suburban and industrial sites across Canada has been confirmed in a recent ongoing monitoring study (Environment Canada, 1991a, unpublished). Mean concentrations of monochlorobenzene in 608 (24-hour) samples of ambient air from 18 sites in five provinces during the period October

1988 through April 1990 ranged from 0.10 to 0.21 $\mu\text{g}/\text{m}^3$; the overall mean value was 0.15 $\mu\text{g}/\text{m}^3$. For the 11 urban and one rural sites included in this program with data for at least 25 days during 1988-89, median concentrations ranged from 0.05 to 0.13 $\mu\text{g}/\text{m}^3$. Daily maximum concentrations ranged from 0.15 to 1.74 $\mu\text{g}/\text{m}^3$ with the higher values recorded at sites having nearby industrial areas. Although data on concentrations of monochlorobenzene found in indoor air in Canada are not available, limited information from other countries indicates that they are similar to those in ambient air (Lebret, 1985; Pellizzari *et al.*, 1986).

Monochlorobenzene was not detected (detection limit of 0.5 to 1.0 $\mu\text{g}/\text{L}$) in extensive monitoring of surface and raw drinking water supplies across Canada between May 1984 and October 1988 (Environment Canada, 1991b) nor in two samples of river water in Canada (Oliver and Bothen, 1980). There is evidence that monochlorobenzene may be produced during treatment of raw drinking water supplies by reaction of chlorine (or one of its aqueous species) with organic material (both natural and man-made). Although levels were too low to permit quantification (detection limit of 1.0 $\mu\text{g}/\text{L}$), the frequency of detection of monochlorobenzene was less for samples of raw than for treated water at 30 Canadian water treatment plants (5 and 18%, respectively) [Otson *et al.*, 1982a; 1982b]. In treated water samples from 30 water treatment plants across Canada, monochlorobenzene was detected in 16/90 samples; mean concentrations were less than 1 $\mu\text{g}/\text{L}$, and the maximum value recorded was 5 $\mu\text{g}/\text{L}$ (Otson *et al.*, 1982b).

In several studies, measurable concentrations of monochlorobenzene have been reported in industrial effluents, and in leachate and ground water near specific sites that received industrial wastes. Monochlorobenzene was detected in samples from effluents being discharged from four organic chemical manufacturing plants near the St. Clair and St. Lawrence Rivers and Lake Ontario monitored between October 1989 and July 1990 (OME, 1992, unpubl.). Concentrations ranged from 0.2 to 50.0 $\mu\text{g}/\text{L}$ with a weighted average for all seven sites ($N = 68$) of 7.4 $\mu\text{g}/\text{L}$. Reinhard *et al.* (1984) reported concentrations of monochlorobenzene in leachate water samples taken in the vicinity of a landfill site in North Bay, Ontario, ranging from 16-33 $\mu\text{g}/\text{L}$. In a study of ground water under another landfill site in Gloucester, Ontario, levels in 530 samples were reported to range from 0.1 to 315 $\mu\text{g}/\text{L}$ (Jackson *et al.*, 1985; 1991). High concentrations of monochlorobenzene ranging from 280 to 5 310 $\mu\text{g}/\text{L}$ were found in samples from six monitoring wells situated within 12 metres of four former wastewater treatment ponds near Elmira, Ontario (CH2M Hill Engineering Ltd., 1991). Disposal of industrial organic wastes near Ville Mercier, Quebec, has resulted in contamination of the surrounding ground water with monochlorobenzene (Pakdel *et al.*, in press). Concentrations of monochlorobenzene in 16 samples from sand and gravel and bedrock aquifers at this site ranged from 224 to 1 382 $\mu\text{g}/\text{L}$, and 445 to 1 787 $\mu\text{g}/\text{L}$, respectively. Waste disposal practices ceased in 1980 and 1972 at the Gloucester and Ville Mercier sites, respectively. Waste disposal at the tar pits and lagoons located near Elmira was suspended in 1979 and 1986, respectively. The Gloucester, Ville Mercier and Elmira sites are undergoing or are targeted for remediation.

Monochlorobenzene was not detected in nine sediment samples from Lake Ontario (detection limit of 1.5 $\mu\text{g}/\text{g}$) [Oliver and Bothen, 1982]. Concentrations of monochlorobenzene ranging from 3.4 to 138.1 $\mu\text{g}/\text{kg}$ were detected in soil samples collected during sampling for ground water in the wastewater treatment pond area near Elmira, Ontario (CH2M Hill Engineering Ltd., 1991). The highest levels of soil contamination (3.4 to 138.1 $\mu\text{g}/\text{kg}$) of monochlorobenzene occurred approximately five metres from a treatment pond, at depths of 6.1 to 6.7 m at the base of one of the shallow aquifers. Reports of concentrations of monochlorobenzene in biota in Canada were not identified.

In view of the possible relative significance of monochlorobenzene released to the environment resulting from pesticide application to soil, and the lack of field data, concentrations of monochlorobenzene that may occur in the soil, the atmosphere, and other media were predicted based on models developed by Mackay and Shiu (1990). Two scenarios were examined: one in which concentrations of monochlorobenzene in the immediate vicinity of pesticide application (near-field model) were predicted, and the other in which concentrations in the broader region surrounding the treated field (regional model) were estimated. Details including modelling parameters, assumptions and results are presented in the Supporting Document to this report. For the near-field model, the soil concentration following application of the pesticide was predicted to be 5.5 µg/g of soil. Airborne concentrations of monochlorobenzene over the field were predicted to be 1 µg/m³. For the regional model under worst-case conditions, the following concentrations were also predicted: 8.4 x 10⁻⁴ µg/m³ in air, 4.8 x 10⁻⁵ µg/L in surface water, 0.93 µg/g in soil and 3.0 x 10⁻⁷ µg/g in sediment. For the air compartment, where comparison can be made, the predicted concentrations are considerably less than those measured during monitoring programs.

Information on concentrations of monochlorobenzene in food has not been identified. Monochlorobenzene has, however, been detected in five of eight samples of human breast milk in the United States, although levels were not quantified (Pellizzari *et al.*, 1982).

2.5 Toxicokinetics and Metabolism

Monochlorobenzene is readily absorbed through the lungs and gastrointestinal tract. Given its lipophilic nature, it is likely that it is also absorbed well through the skin; however, quantitative data on uptake are not available for any of these routes of exposure. Once absorbed, monochlorobenzene is rapidly distributed to many tissues with concentrations being greatest in adipose tissue. It is primarily metabolized by oxidative reactions involving the mixed-function oxidase mediated enzymes to ortho-, meta- or parachlorophenols, the glutathione, glucuronic acid or sulphate conjugates of which are excreted in the urine (Selander *et al.*, 1975; Smith-Lindsay *et al.*, 1972; Yoshida and Hara, 1985). Reactive intermediates of the metabolism of monochlorobenzene, possibly arene oxides and chlorophenols, can bind to cellular proteins; binding of these metabolites appears to be correlated with necrotic pathological damage in the kidneys and liver of rodents (U.S. EPA, 1985).

2.6 Mammalian Toxicology

Monochlorobenzene is acutely toxic following administration by all routes of exposure examined to date (i.e., dermal, oral, intraperitoneal). Acute exposure to monochlorobenzene by inhalation causes sensory irritation of the respiratory system after several minutes; prolonged exposure (several minutes to several hours) causes narcosis and central nervous system depression which can be lethal. Lethal concentrations which killed 50% of the animals (LC₅₀s) in male rats and female mice have been reported to be 13 490 and 8 581 mg/m³, respectively (Bonnet *et al.*, 1979; 1982). Lethal doses which killed 50% of the animals (LD₅₀s) for ingestion (gavage in corn oil) were approximately 4 000 mg/kg in rats (both sexes); mice were more sensitive, with 100% lethality above 1 000 mg/kg and 2 000 mg/kg for males and females, respectively (NTP, 1983; Kluwe *et al.*, 1985). Systemic effects following acute or short-term exposure include damage to the liver and kidneys, and effects on bile and pancreatic flow.

In subchronic studies, administration of MCB by inhalation or ingestion to rats, mice, rabbits and dogs has caused reductions in both body weight gain and survival at high doses, and hepatic and renal toxicity, as indicated by increases in serum enzymes, liver and kidney weights, histopathological changes and necrosis (Dilley, 1977; Irish, 1963; NTP, 1983; Knapp *et al.*, 1971). At high doses, depression of bone marrow activity in mice (Zub, 1978) and myeloid depletion of the thymus, spleen or bone marrow in rats and mice (NTP, 1983) have also been observed. In subchronic inhalation studies in which analyses of at least body weight gain, survival, clinical signs of toxicity, clinical chemistry, haematology and histopathology of major organs and tissues have been conducted, no-observed-effect-levels (NOELs) were approximately 1 000 to 2 000 mg/m³ in rats (Irish, 1963). For ingestion, no-observed-effect-levels were 50 to 125 mg/kg in rats, and 125 mg/kg in mice (NTP, 1983; KIuwe *et al.*, 1985).

The lowest reported effect level in inhalation studies conducted to date was 341 mg/m³ (a "marginal toxic concentration") which resulted in increased kidney weight and tubular and interstitial lesions in the kidney, lesions in the adrenal cortex, and small changes in red blood cell parameters in male rats exposed for 24 weeks (Dilley *et al.*, 1977).

The carcinogenicity of monochlorobenzene has been investigated in only one study, which was conducted by the National Toxicology Program in which groups of male and female rats and female mice (50/group) were administered doses of 0, 60, or 120 mg/kg bw daily by gavage in corn oil, five days per week for 103 weeks. Male mice were administered 0, 30, or 60 mg/kg bw on the same schedule (NTP, 1983; KIuwe *et al.*, 1985). There was no convincing evidence in this study of compound-related toxicity in either rats or mice. There was a significant increase in hepatic neoplastic nodules noted in the high-dose group of male rats (120 mg/kg bw). The increase was significant in comparison with both concurrent vehicle and pooled controls, and there was a marginally significant dose-response trend. However, there were no hepatocellular carcinomas in exposed male rats, and analysis of combined data on neoplastic nodules and hepatocellular carcinomas reduced the significance of the observed increase in tumour incidence. No other significant increases in tumour incidence were observed in either rats or mice. It was concluded that the study provided some evidence of carcinogenicity in male F344/N rats, but no evidence of carcinogenicity in either female F344/N rats or B6C3F₁ mice of either sex. The no-observed-effect-levels in this study were 120 mg/kg/day for female rats and mice and 60 mg/kg/day for male rats and mice. The doses administered in this bioassay were not significantly less than those at which toxic effects were observed in the subchronic studies by the same authors (lowest-observed-adverse-effect-level (LOAEL) = 250 mg/kg/day for 13 weeks), indicating little potential for progressive toxicity with continued monochlorobenzene administration beyond 13 weeks.

Monochlorobenzene has not been teratogenic in rats or rabbits, although slight delays in foetal skeletal development (ossification) have been observed in foetuses of pregnant rats exposed by inhalation to 2 864 mg/m³, a concentration which was also toxic to the mothers (John *et al.*, 1984). In the only reproductive study identified, hepatocellular hypertrophy and renal changes were reported in F₀ and F₁ male rats exposed to 150 ppm (683 mg/m³). At 450 ppm (2 048 mg/m³), there was an increase in the incidence of bilateral degeneration of the testicular germinal epithelium of the F₀ adults which was not, however, observed in the F₁ group; its relationship to monochlorobenzene administration is unclear (Nair *et al.*, 1987).

Monochlorobenzene has induced chromosomal aberrations in plants and bacteria but not in mammalian systems. Indeed, limited available data indicate that monochlorobenzene has little genotoxic potential.

2.7 Effects on Humans

Available data on the effects of exposure to monochlorobenzene in humans are restricted to case reports, three limited epidemiological studies of occupationally-exposed populations, and a limited clinical investigation of the threshold for effects on the electrical activity in the brain of a very small number of subjects. Although effects on the nervous system, neonatal development and skin have been reported in occupationally-exposed populations, available studies are inadequate for assessing potential risks associated with exposure to monochlorobenzene due to methodological weaknesses such as a lack of documentation of the nature or magnitude of exposure, concomitant exposure to other compounds and, in some cases, lack of control groups.

2.8 Effects on the Environment

The information available on acute and chronic toxicity of monochlorobenzene includes data for a number of trophic levels from bacteria through to fish in the aquatic environment. Information on toxicity to terrestrial species is very limited. Although no data were found on wild mammals, the toxicity of monochlorobenzene to these organisms can be assessed by extrapolation from the results of toxicity studies conducted with laboratory mammals (reported in section 2.6). No data were available on effects on birds or terrestrial plants.

In the following two paragraphs, the results of acute toxicity studies considered representative of data available for different trophic levels are summarized. For the bacteria, *Pseudomonas putida*, a 16-h toxicity threshold for inhibition of cell multiplication was reported by Bringmann and Kuhn (1980) to be 17 mg/L. Calamari *et al.* (1983) reported a 96-h concentration which was effective in inhibiting growth by 50% (EC₅₀) of 12.5 mg/L for growth inhibition in the algae, *Selenastrum capricornutum*. The latter authors also reported a 24-h LC₅₀ of 4.3 mg/L for *Daphnia magna*. Other authors reported, also for *Daphnia magna*, 48-h LC₅₀s ranging between 5.8 and 25.8 mg/L (Bobra *et al.*, 1985; Hermens *et al.*, 1984; Abernathy *et al.*, 1986; Gersich *et al.*, 1986; Cowgill *et al.*, 1985). For the rainbow trout (*Oncorhynchus mykiss*), 96-h LC₅₀s for tests conducted in flow-through systems and with measured concentrations were reported by Dalich *et al.* (1982) and Hodson *et al.* (1984) to be 4.7 and 7.46 mg/L, respectively.

Only one acute toxicity study was found concerning terrestrial organisms. Neuhauser *et al.* (1986) exposed earthworms (*Eisenia fetida*) by contact with filter papers soaked in graduated concentrations of monochlorobenzene and reported, under these conditions, a 24-h LC₅₀ of 29 µg/cm². However, the protocol followed for this study does not allow for extrapolation of these results to field conditions.

With respect to longer term studies, two reports were found concerning the toxicity of monochlorobenzene on various species of fish exposed continually, in closed, flow-through systems, from shortly after fertilization of the egg to four days after hatching. The lowest reported LC₅₀ for aquatic species was 0.05 mg/L for the largemouth bass (*Micropterus salmoides*). The total exposure period averaged seven days (Birge *et al.*, 1979). Black *et al.* (1982) reported an LC₅₀ of 0.11 mg/L for the same early life stages of rainbow trout (*Oncorhynchus mykiss*), where the total exposure time was approximately 27 days.

In this paragraph, the results of chronic studies conducted with monochlorobenzene on aquatic organisms are summarized. Hermens *et al.* (1984; 1985) reported a 16-day EC₅₀ of

1.1 mg/L for sub-lethal reproductive effects for *Daphnia magna*. They also reported for growth reduction in the same species, a 16-day EC₅₀ of 3.3 mg/L and a 16-day LC₅₀ of 3.9 mg/L. De Wolf *et al.* (1988) reported no-observed-effect-levels for growth and reproductive effects of 0.32 mg/L and 1.0 mg/L, respectively, for *Daphnia magna* that were exposed until the control daphnids had produced four broods. van Leeuwen *et al.* (1990) studied the effects of monochlorobenzene on the growth of the zebra fish (*Brachydanio rerio*). They reported a 28-day no-observed-effect-concentration (NOEC) of 4.8 mg/L.

3.0 Assessment of "Toxic" under CEPA

As described in the Introduction of this report, the following assessment is organized according to the sources of monochlorobenzene, the exposure of humans and other biota, and potential resulting harmful effects.

3.1 Entry

Monochlorobenzene is released directly onto soil during pesticide application, to the general environment via emissions from industrial manufacturing and processing, and in leachate from selected landfill sites. There is also reason to believe that monochlorobenzene is formed and released during incineration of hazardous wastes. Monochlorobenzene has been measured in the atmosphere, in selected industrial effluents, and in ground water and leachate near specific waste disposal sites in Canada. It was also detected in treated drinking water.

3.2 Exposure

Monochlorobenzene was not found in measurable concentrations in Canadian surface waters nor in sediments from the Great Lakes as would be expected due to its physical and chemical properties. Monochlorobenzene was measured in raw effluent from four organic chemical manufacturing plants in Ontario. Concentrations averaged 7.4 µg/L with a reported maximum of 50 µg/L. Monochlorobenzene is not believed to bioaccumulate to any significant extent in biota.

The concentrations of monochlorobenzene in atmospheric samples from across Canada averaged 0.15 µg/m³ (range of mean values, 0.10 to 0.21 µg/m³) with the highest measured concentration being 1.74 µg/m³.

The use of monochlorobenzene as a carrier solvent for pesticides may result in temporarily high concentrations in the top layers of soil as well as in the atmosphere above the treated field. Concentrations following pesticide application were predicted by modelling techniques to be 5.5 µg/g in soil and 1 µg/m³ in air above the field. On a regional scale, concentrations were predicted to be under 1 µg/g and 1 x 10⁻³ µg/m³ in soil and air, respectively.

Since humans are exposed to monochlorobenzene in all media, total intake has been assessed on a multimedia basis. Although available data on concentrations of monochlorobenzene in environmental media to which the general public is exposed are limited, it is possible to estimate the intake of monochlorobenzene from various sources, with the exception of food (Table 1). However, on the basis of available data on concentrations of the other monochlorobenzenes in food, physical/chemical properties, and limited information on levels of monochlorobenzene in air and drinking water, it is likely that the intake of monochlorobenzene from air is greater than that from food or drinking water. For suckling infants, mothers' milk may also be an important source of exposure, though quantitative data are not available to serve as a basis for estimation of intake via this route. Based on available data, and as indicated in Table 1, it has been estimated that the total daily intake of monochlorobenzene (in air and drinking water) ranges from 0.047 to 0.087 µg/kg bw, 0.102 to 0.142 µg/kg bw, 0.081 to 0.131 µg/kg bw, 0.06 to 0.1 µg/kg bw, and 0.051 to 0.081 µg/kg bw for Canadians aged less than 6 months, 6 months to 4 years, 5 to 11 years, 12 to 19 years, and 20 to 70 years, respec-

tively. These estimated intakes which are expected to be typical for the majority of the general population are based on mean values measured in the general environment. Elevated levels present, for example, in ground water as a result of poor waste disposal practices in isolated cases, were not considered relevant to estimation of exposure for the general population.

Table 1 - Estimated Daily Intake ($\mu\text{g}/\text{kg}$) of MCB by Canadians from Various Sources

Medium	Estimated Intake ($\mu\text{g}/\text{kg}\text{-bw}/\text{day}$)				
	0-0.5 yr ^a	0.5-4 yr ^b	5-11 yr ^c	12-19 yr ^d	20-70 yr ^e
Ambient Air ^f	0.03-0.07	0.04-0.08	0.04-0.09	0.04-0.08	0.03-0.06
Drinking Water ^g	<0.017	<0.062	<0.041	<0.020	<0.021
Food	N/A	N/A	N/A	N/A	N/A
Total Intake*	0.047-0.087	0.102-0.142	0.081-0.131	0.06-0.1	0.051-0.081

^a Assumed to weigh 6 kg, breathe 2 m³ of air per day and drink 0.1 L of water per day (Environmental Health Directorate, 1988)

^b Assumed to weigh 13 kg, breathe 5 m³ of air per day and drink 0.8 L of water per day (Environmental Health Directorate, 1988)

^c Assumed to weigh 27 kg, breathe 12 m³ of air per day and drink 1.1 L of water per day (Environmental Health Directorate, 1988)

^d Assumed to weigh 55 kg, breathe 21 m³ of air per day and drink 1.1 L of water per day (Environmental Health Directorate, 1988)

^e Assumed to weigh 70 kg, breathe 20 m³ of air per day and drink 1.5 L of water per day (Environmental Health Directorate, 1988)

^f Based on range of mean concentrations reported in a survey of concentrations in ambient air from 18 Canadian sites in five provinces (0.10-0.21 $\mu\text{g}/\text{m}^3$) [Environment Canada, 1991a, unpublished]

^g Based on a mean concentration of MCB in drinking water of <1.0 $\mu\text{g}/\text{L}$ (Otson *et al.*, 1982a; 1982b)

N/A No data available

* Data on concentrations of MCB in indoor air in Canada were not identified; based on information from other countries, concentrations in indoor air appear to be similar to those in ambient air (Lebret, 1985; Pellizzari *et al.*, 1986).

3.3 Effects

3.3.1 Human Health

Based on the increased incidence of hepatic neoplastic nodules in F344/N male rats observed in the NTP carcinogenesis bioassay (NTP, 1983; Kluwe *et al.*, 1985), monochlorobenzene has been classified in Group IIIB - (possibly carcinogenic to man) of the classification scheme developed for use in the derivation of the "Guidelines for Canadian Drinking Water Quality" (Environmental Health Directorate, 1989).

For compounds classified in Group IIIB, a Tolerable Daily Intake (TDI) is derived on the basis of division of the no-or lowest-observed-(adverse)-effect-level (NOAEL) or LO(A)EL observed in an animal species conducted by the most appropriate route of administration by an uncertainty factor that takes into account, where appropriate, the limited evidence of carcinogenicity. Data on concentrations of monochlorobenzene in food and breast milk are not available, and information on levels in drinking water in Canada are sparse. However, on the basis of available data on concentrations of the other monochlorobenzenes in food, physical/chemical properties, and limited information on levels of monochlorobenzene in air and drinking water, it is likely that for most of the population, the intake of monochlorobenzene from air is greater than that from food or drinking water. Therefore, a TDI based on the results of inhalation studies was derived as follows:

$$\begin{aligned} \text{TDI} &= \frac{341 \text{ mg/m}^3 \times (7/24) \times (5/7) \times 0.144 \text{ m}^3/\text{d}}{5\,000 \times 0.25 \text{ kg}} \\ &= 0.0081 \text{ mg/kg/d (8.1 } \mu\text{g/kg/d)} \end{aligned}$$

where:

- 341 mg/m³ is the lowest reported effect level ("marginal toxic concentration") based on increased kidney weight and tubular and interstitial lesions in the kidney, lesions in the adrenal cortex and small changes in red cell parameters in male rats in the limited available studies (Dilley, 1977);
- 7/24 and 5/7 is the conversion of 7 hours/day, 5 days per week dosing to continuous exposure;
- 0.144 m³/d is the assumed inhaled air volume of rats (NIOSH, 1985);
- 0.25 kg is the assumed body weight of adult rats (NIOSH, 1985);
- 5 000 is the uncertainty factor (x 10 for interspecies variation; x 10 for intraspecies variation; x 10 for less-than-chronic and limited study; x 5 for use of a LOAEL rather than a NOAEL, though effects at the LOAEL were considered to be only marginally adverse).

Owing to limitations of the critical study on which this TDI is based, a TDI was also derived based on the more extensive long-term NTP study conducted by the oral route, as follows:

$$\text{TDI} = \frac{60 \text{ mg/kg bw/d} \times 5}{500 \times 7} \sim 0.086 \text{ mg/kg bw/d (86 } \mu\text{g/kg bw/d)}$$

where:

- 60 mg/kg bw/d is the lowest NOEL or NOAEL (male rats and mice) in the only chronic and/or carcinogenesis bioassay (NTP, 1983; Kluwe *et al.*, 1985);

- 5/7 is the conversion of 5 days per week of dosing to 7 days per week;
- 500 is the uncertainty factor (x 10 for interspecies variation; x 10 for intraspecies variation; x 5 for limited evidence of carcinogenicity - i.e., the increase in hepatic neoplastic nodules in male rats in the NTP carcinogenesis bioassay).

This value is less conservative than that derived above on the basis of studies conducted by the most relevant route of exposure (i.e., inhalation).

3.3.2 Environment

For aquatic biota, the most sensitive organism identified was the early stages of the largemouth bass (*Micropterus salmoides*). The reported LC₅₀ was 0.05 mg/L for this species following an exposure period of approximately 7 days. The lowest LC₅₀ reported for acute toxicity of monochlorobenzene was 4.1 mg/L for *Daphnia magna*.

Neither acute nor chronic studies were found for wildlife. However, the effect levels reported in inhalation studies conducted in laboratory animals are considered relevant to wild mammals. The lowest reported effect level in the longest term inhalation study conducted to date was 341 mg/m³ which resulted in increased kidney weight and tubular and interstitial lesions in the kidney, lesions in the adrenal cortex, and small changes in red blood cell parameters in male rats exposed for 24 weeks (Dilley *et al.*, 1977).

3.4 Conclusions

Monochlorobenzene is used in Canada in a variety of applications that lead to the direct (pesticides) and indirect (via effluents, emissions, and leachate waters) entry of this substance into the Canadian environment. These releases result in measurable or predictable concentrations of monochlorobenzene in the various media to which humans and other organisms may be exposed, albeit at low levels.

3.4.1 Paragraph 11(a) - Effects on the Environment

The LC₅₀ (0.05 mg/L) for the most sensitive aquatic species, the largemouth bass (*Micropterus salmoides*), under chronic exposure conditions, is higher than concentrations predicted by computer modelling in surface water by six orders of magnitude. It is also six times higher than the average concentration found in raw effluent from organic chemical manufacturing facilities. The lowest acute IC₅₀ (4.3 mg/L for *Daphnia magna*) is 80 times higher than the highest reported concentration in the same raw effluent.

The lowest reported effect level in the longest term inhalation study conducted to date in laboratory animals was 341 mg/m³. This value is more than five orders of magnitude higher than the highest measured concentration (1.74 µg/m³) and more than six orders of magnitude greater than mean concentrations (0.15 µg/m³) reported in the atmosphere in Canada to which wild mammals may be exposed.

Therefore, on the basis of available data, monochlorobenzene is not considered to be "toxic" as interpreted under paragraph 11(a) of the *Canadian Environmental Protection Act*.

3.4.2 Paragraph 11(b) -- Effects on the Environment on which Human Life Depends

Because of its short persistence in the atmosphere, and relatively low levels of release, monochlorobenzene cannot be associated with ozone layer depletion. Similarly, the magnitude of potential effects on global warming and photochemical smog formation, although difficult to quantify, is not believed to be significant.

Therefore, on the basis of available data, monochlorobenzene is not considered to be "toxic" as interpreted under paragraph 11(b) of the *Canadian Environmental Protection Act*.

3.4.3 Paragraph 11(c) -- Effects on the Human Life or Health

Based on the limited available data, total estimated average daily intakes of monochlorobenzene for various age groups in the Canadian population range from 0.047 to 0.142 $\mu\text{g}/\text{kg}$ bw (Table 1). These estimated average daily intakes are considerably less (from about 60 to 170 times) than the most conservative TDI derived above on the basis of the results of studies by the most appropriate route of exposure (inhalation, 8.1 $\mu\text{g}/\text{kg}$ bw).

Therefore, on the basis of available data, monochlorobenzene is not considered to be "toxic" as interpreted under paragraph 11(c) of the *Canadian Environmental Protection Act*.

3.4.4 General Conclusions

Therefore, on the basis of available data, monochlorobenzene is not considered to be "toxic" as interpreted under paragraphs 11(a), (b) and (c) of the *Canadian Environmental Protection Act*.

4.0 Recommendations for Research

1. To permit a more complete assessment of exposure of the Canadian population to monochlorobenzene, additional monitoring data are desirable, particularly for food and breast milk, though the priority for this research is considered to be low.
2. In one study, monochlorobenzene was toxic to earthworms, though these results could not be extrapolated to predicted field conditions for MCB used in pesticides. Data on the fate and toxicity of MCB in soil as a result of this use are, therefore, desirable; however, the priority for this research is considered to be low.

5.0 References

- Abernethy, S.G., A.M. Bobra, W.Y. Shiu, P.G. Wells, and D. Mackay. 1986. Acute lethal toxicity of hydrocarbons and chlorinated hydrocarbons to two planktonic crustaceans: the key role of organism-water partitioning. *Aquat. Toxicol.* 8(3): 163-174.
- Atkinson, R., S.M. Aschmann, A.M. Winer, and J.N. Pitts. 1985. Atmospheric gas phase loss processes for chlorobenzenes, benzotrifluoride, and 4-chlorobenzotrifluoride and generalization of predictive techniques for atmospheric lifetimes of aromatic compounds. *Arch. Environ. Contam. Toxicol.* 14: 417-425.
- Birge, W.J., J.A. Black, J.E. Hudson, and D.M. Bruser. 1979. Embryo-larval toxicity tests with organic compounds. In: Marking, L.L., and R.A. Kimerle, eds., *Aquatic Toxicology*, ASTM STP 667:131-147.
- Black, J.A., and W.J. Birge. 1982. The aquatic toxicity of organic compounds to embryo-larval stages of fish and amphibians. Research Report No. 133, Water Resources Research Institute, University of Kentucky, Lexington, Kentucky, p. 61.
- Bobra, A., W.Y. Shin, and D. Mackay. 1985. Quantitative structure-activity relationships for the acute toxicity of chlorobenzenes to *Daphnia magna*. *Environ. Toxicol. Chem.* 4: 297-305.
- Bonnet, P., G. Raoult, and D. Gradiski. 1979. Lethal concentration 50 of main aromatic hydrocarbons. *Arch. Mal. Prof.* 40(8-9): 805-810. (in French).
- Bonnet, P., Y. Morele, G. Raoult, D. Zissu, and D. Gradiski. 1982. Determination of the median lethal concentration of the main aromatic hydrocarbons in the rat. *Arch. Mal. Prof.* 43(4): 461-465. (in French).
- Bringmann, G. and R. Kuhn. 1980. Comparison of the toxicity thresholds of water pollutants to bacteria, algae, and protozoa in the cell multiplication inhibition test. *Water Res.* 14: 231-241.
- Calamari, D., S. Galassi, F. Setti, and M. Vighi. 1983. Toxicity of selected chlorobenzenes to aquatic organisms. *Chemosphere* 12(2): 253-262.
- Callahan, M., M. Slimak, N. Gabel, I. May, C. Fowler, R. Freed, P. Jennings, R. Durfee, F. Whitmore, B. Maestri, W. Mabey, B. Holt, and C. Gould. 1979. Water-related environmental fate of 129 priority pollutants, Volume I. Monitoring and Data Support Division, U.S. Environmental Protection Agency (EPA 440/4-79-029a).
- Camford Information Services Inc. 1991. Chlorobenzene CPI Product Profile. Don Mills, Ontario, 4 p.
- CCME. 1990. Canadian Council of Ministers of the Environment. CCME Management Plan for Nitrogen Oxides (NO_x) and Volatile Organic Compounds (VOCs). Canadian Council of Ministers of the Environment, p.176.
- CH2M Hill Engineering Ltd. 1991. Research and development of permanent on-site solutions for contamination of ground water at waste disposal and industrial sites in Canada. Final Report. Waterloo, Ontario.

- Cowgill, U.M., I.T. Takahashi, and S.L. Applegath. 1985. A comparison of the effect of four benchmark chemicals on *Daphnia magna* and *Ceriodaphnia dubia-affinis* tested at two different temperatures. *Environ. Toxicol. Chem.* 4: 415-422.
- Dalich, G.M., R.E. Larson, and W.H. Gingerich. 1982. Acute and chronic toxicity studies with monochlorobenzene in rainbow trout. *Aquat. Toxicol.* 2: 127-142.
- Dc Wolf, W., J.H. Canton, J.W. Deneer, R.C.C. Wegman, and J.L.M. Hermens. 1988. Quantitative structure-activity relationships and mixture-toxicity studies of alcohols and chlorohydrocarbons: reproducibility of effects on growth and reproduction of *Daphnia magna*. *Aquat. Toxicol.* 12: 39-49.
- Dilley, J.V. 1977. Toxic evaluation of inhaled chlorobenzene (monochlorobenzene). National Technical Information Service, U.S. Department of Commerce (PB-276 623).
- Dulin, D., H. Drossman, and T. Mill. 1986. Products and quantum yields for photolysis of chloroaromatics in water. *Environ. Sci. Technol.* 20: 72-77.
- Ellington, J.J., F.E. Stancil, W.D. Payne, and C.D. Trusty. 1988. Measurement of hydrolysis rate constants for evaluation of hazardous waste land disposal, Volume III. Data on 70 chemicals. Office of Research and Development, U.S. Environmental Protection Agency (EPA/600/3-88/028), 29 p.
- Environment Canada. 1991a, unpublished. Measurement program for toxic contaminants in Canadian urban air. River Road Environmental Technology Centre. PMD 91-2: 17.
- Environment Canada. 1991b. Naquadat chlorobenzene data. Water Quality Branch, Environment Canada, p.15.
- Environmental Health Directorate. 1988. Reference Values for Canadian Populations. Draft Report prepared by the Environmental Health Directorate Working Group on Reference Values. Health and Welfare Canada, Ottawa.
- Environmental Health Directorate. 1989. Derivation of maximum acceptable concentrations and aesthetic objectives for chemicals in drinking water. In: Guidelines for Canadian Drinking Water Quality - Supporting Documentation. Health and Welfare Canada, Bureau of Chemical Hazards.
- Freitag, D., L. Ballhorn, H. Geyer, and F. Korte. 1985. Environmental hazard profile of organic chemicals. *Chemosphere* 4(10): 1589-1616.
- Garrison, A.W., and D.W. Hill. 1972. Organic pollutants from mill persistent in downstream waters. *American Dyestuff Report*. (February): 23-25.
- Gersich, F.M., P.A. Blanchard, S.L. Applegath, and C.N. Park. 1986. The precision of daphnid (*Daphnia magna* Straus, 1820) static acute toxicity tests. *Arch. Environ. Contam. Toxicol.* 15: 741-749.
- Graham, J.L., D.L. Hall, and B. Dellinger. 1986. Laboratory investigation of thermal degradation of a mixture of hazardous organic compounds. *Environ. Sci. Technol.* 20(7): 703-710.

- Hermens, J., H. Canton, P. Janssen, and R. de Jong. 1984. Quantitative structure-activity relationships and toxicity studies of mixtures of chemicals with an anaesthetic potency: acute lethal and sublethal toxicity to *Daphnia magna*. *Aquat. Toxicol.* 5: 143-154.
- Hermens, J., E. Broekhuizen, H. Canton, and R. Wegman. 1985. Quantitative structure activity relationships and mixture toxicity studies of alcohols and chlorohydrocarbons: effects on growth of *Daphnia magna*. *Aquat. Toxicol.* 6(3): 209-217.
- Hodson, P.V., D.G. Dixon, and K.L.E. Kaiser. 1984. Measurement of median lethal dose as a rapid indication of contaminant toxicity to fish. *Environ. Toxicol. Chem.* 3(2): 243-254.
- Holliday, M.G., and F.R. Engelhardt. 1984a. Chlorinated benzenes. A criteria review. Prepared for Monitoring and Criteria Division, Bureau of Chemical Hazards, Health and Welfare Canada, Ottawa.
- Holliday, M.G., F.R., Engelhardt, and I. MaClachian. 1984b. Chlorobenzenes: an environmental health perspective. Prepared for Health and Welfare Canada, Ottawa.
- IPCC (Intergovernmental Panel on Climate Change). 1990. Climate change: the IPCC scientific assessment. Houghton, J.T., G. Jenkins, H.H. Ephraums, editors. Cambridge University Press, New York, p.51.
- Irish, D.D. 1963. Halogenated hydrocarbons: II. Cyclic. In: Patty, F.A., ed. *Industrial Hygiene and Toxicology*, 2nd edition. Inter-Science Publishers, New York: 1333-1340.
- Jackson, R.E., S. Lesage, M.W. Priddle, A.S. Crowe, and S. Shikaze. 1991. Contaminant hydrogeology of toxic organic chemicals at a disposal site, Gloucester, Ontario. 2. Remedial investigation. Scientific Series No. 181. National Water Research Institute, Inland Waters Directorate, Environment Canada, p. 68.
- Jackson, R.D., R.J. Patterson, B.W. Graham, J. Bahr, D. Belanger, J. Lockwood, and M. Priddle. 1985. Contaminant hydrogeology of toxic organic chemicals at a disposal site, Gloucester, Ontario. 1. Chemical concepts and site assessment. National Hydrogeology Research Institute Paper No. 23, Inland Waters Directorate, Environment Canada, p. 114.
- John, J.A., W.C. Hayes, T.R. Hanley Jr., K.A., Johnson, T.S. Gushow, and K.S. Rao. 1984. Inhalation teratology study on monochlorobenzene in rats and rabbits. *Toxicol. Appl. Pharmacol.* 76: 365-373.
- Kiuwe, W.M., G. Dill, A. Persing, and A. Peters. 1985. Toxic response to acute, subchronic, and chronic oral administrations of monochlorobenzene to rodents. *J. Toxicol. Environ. Health* 15(6): 745-767.
- Knapp, W.K.J.R., W.M. Busey, and W. Kundzins. 1971. Subacute oral toxicity of monochlorobenzene in dogs and rats. *Toxicol. Appl. Pharmacol.* 19(2): 393. (Abstract)
- Lebret, E. 1985. Air pollution in Dutch homes: an exploratory study in environmental epidemiology. Department of Air Pollution, Department of Environmental and Tropical Health, Wageningen Agricultural University, The Netherlands. Report R- 138, Report 1985-221.

- Lee, R.P., and C. Ryan. 1979. Proceedings of the workshop: microbial degradation of organochlorine compounds in estuarine waters and sediments. Office of Research and Development, U.S. Environmental Protection Agency (EPA-600/9-79-012), 7 p.
- Mackay, D., and W.Y. Shiu. 1990. Physical-chemical properties and fate of volatile organic compounds: an application of the fugacity approach. In: Ram, N.M., R.F. Christman, and K.P. Cantor, eds. Significance and Treatment of Volatile Organic Compounds in Water Supplies. Lewis Publishers, Michigan: 183-201.
- Mackay, D., W.Y. Shiu, and R.P. Sutherland. 1979. Determination of air-water Henry's law constants for hydrophobic pollutants. *Environ. Sci. Technol.* 13(3): 333-337.
- Mackay, D., and A.T.K. Yuen. 1983. Mass transfer coefficients for volatilization of organic solutes from water. *Environ. Sci. Technol.* 17: 211-217.
- McCall, J.P., D.A. Laskowski, R.L. Swann, and H.J. Dishburger. 1981. Measurement of sorption coefficients of organic chemicals and their use in environmental fate analysis. In: Test Protocols for Environmental Fate and Movement of Toxicants, p. 89-109. Proceedings of a Symposium. Association of Official Analytical Chemists. 94th Annual Meeting, October 21-22, 1980. Washington, D.C.
- Morrison, R.T., and R.N. Boyd. 1987. Organic Chemistry, 3rd edition. Allyn and Bacon, Inc., Boston.
- Nair, R.S., J.A. Barter, R.E. Schroeder, A. Knezevich, and C.R. Stack. 1987. A two-generation reproduction study with monochlorobenzene vapor in rats. *Fundamental and Applied Toxicology* 9: 678-686.
- Neuhauser, E.F., R.C. Loehr, and M.R. Malecki. 1986. Contact and artificial Soil tests using earthworms to evaluate the impact of wastes in soil. In: Hazardous and Industrial Solid Waste Testing: Fourth Symposium, ASTM STP 886: 192-203.
- NIOSH. 1985. Registry of Toxic Effects of Chemicals Substances (1983-84). Cumulative supplement to the 1981-82 edition. U.S. Department of Health and Human Services.
- NTP (National Toxicology Program). 1983. NTP Technical Report on the carcinogenesis studies of chlorobenzene (CAS No. 108-90-7) in F344/N rats and B6C3F₁ mice (gavage studies). NTP TR 261, U.S. Department of Health and Human Services, Research Triangle Park, North Carolina, p.228.
- Oliver, B.G., and K.D. Bothen. 1980. Determination of chlorobenzenes in water by capillary gas chromatography. *Anal. Chem.* 52: 2066.
- Oliver, B.G., and K.D. Bothen. 1982. Extraction and clean-up procedures for measuring chlorobenzenes in sediments and fish by capillary gas chromatography. *Internat. J. Environ. Anal. Chem.* 12: 131-139.
- OME (Ontario Ministry of the Environment). 1992, unpublished. Twelve month monitoring data report. Water Resources Branch, Municipal Strategy for Abatement (MISA).
- Otson, R., D.T. Williams, and D.C. Biggs. 1982a. Relationships between raw water quality, treatment and occurrence of organics in Canadian potable water. *Bull. Environ. Contam. Toxicol.* 28(4): 396-403.

- Otson, R., D.T. Williams, and P.D. Bothwell. 1982b. Volatile organic compounds in water at thirty Canadian potable water treatment facilities. *J. Assoc. Off. Anal. Chem.* 65(6): 1370-1374.
- Pakdel, H., G. Couture, C. Roy, A. Masson, J. Locat, P. Gelinas, and S. Lesage. 1991. (in press) Method development for the analysis of toxic chemicals in soil and ground water. The Case of Ville Mercier, P.Q. In: S. Lesage and R.E. Jackson, eds. *Groundwater Quality and Analysis at Hazardous Waste Sites*. Marcel Dekker Inc.
- Pellizzari, E.D., T.D. Hartwell, B.S.H. Harris, R.D. Waddell, D.A. Whitaker, and M.D. Erikson. 1982. Purgeable organic compounds in mother's milk. *Bull. Environ. Contam. Toxicol.* 28: 322-328.
- Pellizzari, E.D., T.D. Hartwell, R.L. Perritt, C.M. Sparacino, L.S. Sheldon, H.S. Zelon, R.W. Whitmore, J.J. Breen, and L. Wallace. 1986. Comparison of indoor and outdoor residential levels of volatile organic chemicals in five U.S. geographical areas. *Environ. Int.* 12(6): 619-623.
- Reinhard, M., N.L. Goodman, and J.F. Barker. 1984. Occurrence and distribution of organic chemicals in two landfill leachate plumes. *Environ. Sci. Technol.* 18(12): 953-961.
- Schwarzenbach, R.P., and J. Westall. 1981. Transport of nonpolar organic compounds from surface water to ground water. Laboratory sorption studies. *Environ. Sci. Technol.* 15(11): 1360-1367.
- Selander, H.G., D.M. Jerina, and J.W. Daly. 1975. Metabolism of chlorobenzene with hepatic microsomes and solubilized cytochrome P-450 systems. *Arch. Biochem. Biophys.* 168: 309-321.
- Singh, H.B., L.J. Salas, R. Stiles, and H. Shigeishi. 1983. Measurements of hazardous organic chemicals in the ambient atmosphere. Office of Research and Development, U.S. Environmental Protection Agency (EPA-600/3-83-001), 82 p.
- Smith-Lindsay, J.R., B.A. Shaw, and D.M. Foulkes. 1972. Mechanisms of mammalian hydroxylation: some novel metabolites of chlorobenzene. *Xenobiotica* 2(3): 215-226.
- Strahlendorf, P.W. 1978. Chlorinated benzenes as potential environmental health hazards: a review. Prepared for Monitoring and Criteria Section, Health Protection Branch, Health and Welfare Canada, Ottawa.
- Thomas, R.G. 1982. Volatilization from soil. In: Lyman, W.J., W.F. Reehl, and D.H. Rosenblatt, eds. *Handbook of Chemical Property Estimation Methods. Environmental Behavior of Organic Compounds*. McGraw-Hill Book Company, Toronto, p. 16-1-16-50.
- U.S. EPA. 1985. Health assessment document for chlorinated benzenes. Final Report. Office of Health and Environment Assessment, Washington, D.C., U.S. Environmental Protection Agency (EPA/600/8-84/015F).
- U.S. EPA. 1991. Minimization and control of hazardous combustion by-products. Risk Reduction Engineering Laboratory, Cincinnati, OH. U.S. Environmental Protection Agency (EPA/600/52-90/039), 399 p.

van Leeuwen, C.J., D.M.M. Adema, and J. Hermens. 1990. Quantitative structure-activity relationships for fish early life stage toxicity. *Aquat. Toxicol.* 16: 321-334.

Wilson, J.T., C.G. Enfield, W.J. Dunlap, R.L. Cosby, D.A. Foster, and L.B. Baskin. 1981. Transport and fate of selected organic pollutants in a sandy soil. *J. Environ. Qual.* 10(4): 501-506.

Yoshida, M., and I. Hara. 1985. Composition of urinary metabolites and variation of urinary taurine levels in rats injected with chlorobenzene. *Industrial Health* 23(3): 239-243.

Zub, M. 1978. Reactivity of the white blood cell system to toxic action benzene and its derivatives. *Acta Biologica Cracoviensia* 21: 163-174.