

Historical Changes in Serum PCB and DDT Levels in an Environmentally-Exposed Cohort

Mary E. Hovinga¹, MaryFran Sowers^{**}, and Harold E. B. Humphrey[†]

^{*}Department of Epidemiology, School of Public Health, University of Alabama at Birmingham, Birmingham, Alabama 35294, USA; ^{**}Department of Epidemiology, School of Public Health, University of Michigan, Ann Arbor, Michigan 48109, USA and [†]Michigan Department of Public Health, Lansing, Michigan 48960, USA

Abstract. A previously characterized cohort of 115 Great Lakes fish eaters and 95 non-fishing controls was re-examined in 1989 to evaluate changes that had occurred in serum PCB and DDT levels since the 1982 study. Substantial and significant decreases in mean serum DDT levels had occurred in both fish eaters (25.8 ppb vs 15.6 ppb) and controls (9.6 ppb vs 6.8 ppb)² over this time period. In contrast, only a slight decrease in serum PCB levels was observed, and in fish eaters only. No association between individual changes in serum PCB or DDT levels and self-reported changes in Great Lakes fish consumption was observed. The findings from this longitudinal examination of serum PCB and DDT levels confirm earlier cross-sectional surveys of western populations, and demonstrate that the prohibition of DDT has been successful in reducing the level of DDT contamination in human populations.

Widespread and increasing use of polychlorinated biphenyls (PCBs) and DDT from the 1940's through the 1960's resulted in extensive environmental contamination. These organochlorine compounds are not only exceptionally stable and persistent in the physical environment, but, because of their lipophilic nature, have the ability to biomagnify in aquatic and terrestrial foodchains (Risebrough *et al.* 1968). Examination of a variety of wildlife has clearly demonstrated that extensive contamination of fish, bird, and mammalian species by PCBs and DDT has occurred (Risebrough *et al.* 1968; Dustman and Stickel 1969; Horn *et al.* 1979; Veith *et al.* 1981; Foley *et al.* 1988).

Contamination has also been demonstrated in humans. Measurable levels of PCBs and DDT have been found in human adipose tissue, blood, and breast milk throughout the world (Quimby *et al.* 1965; Zavon *et al.* 1969; Mes *et al.* 1982; Frank

et al. 1988; Skaare *et al.* 1989; Kutz *et al.* 1991). A primary route of human exposure has been the consumption of contaminated foods. In the Great Lakes region, consumption of contaminated fish has been identified as an important exposure route (Humphrey 1983a; Schwartz *et al.* 1983; Fiore *et al.* 1989). In a 1982 study by the Michigan Department of Public Health (Humphrey 1983b), individuals who consumed 24 or more pounds of sport-caught Great Lakes fish had a mean serum PCB level three times greater than that of the non-fishing controls. The mean serum DDT level for the fish eaters was more than twice that of the control group. Further analysis of this study population demonstrated that fish consumption was a major determinant of the elevated PCB and DDT levels in the fish eater group.

The environmental persistence and biomagnification of PCBs and DDT, coupled with evidence of serious adverse health effects in wildlife populations, prompted a reduction in the use of DDT and PCBs in the United States. The use of DDT was banned in 1972, and PCB production was halted in 1977. The effects of these prohibitions have become apparent in the environment. In the Great Lakes region, sampling data suggest that the degree of DDT and PCB contamination in fish declined sharply in the 1970's, and has plateaued more recently (Schmitt *et al.* 1985; D'Itri 1988; Schmitt *et al.* 1990). Surveys of dairy milk in Canada have suggested a similar pattern (Frank and Braun 1989).

The current study examined whether the observed declines in environmental DDT and PCB levels were also detectable in human populations. Surveys monitoring contaminant levels in human adipose tissue, blood, and breast milk in the United States (Kutz *et al.* 1991), Canada (Frank *et al.* 1988), and Norway (Skaare *et al.* 1989), suggest that DDT levels have been declining since the early 1970s while PCB levels have remained stable. However, these studies have been cross-sectional, and did not allow longitudinal follow-up of individuals, or identification of the relevant sources of exposure. Re-contact of the 1982 Michigan Department of Public Health study population provided an opportunity for the current study to evaluate historical changes in serum DDT and PCB levels within individuals who had a previously identified route of environmental exposure.

¹ Address correspondence to Dr. Mary E. Hovinga.

² ppb = ng/g

Table 1. Comparison of mean baseline characteristics of Entire Cohort and Southern Region Fish eaters and Controls

	1982 Fish eaters		1982 Controls	
	Entire Cohort	Southern Region	Entire Cohort	Southern Region
N	572	197	419	162
1982 age (yrs)	45.9	45.6	44.1	42.3
% male	65.0	63.5	43.2	49.4
1982 Quetelet Index	2.61	2.56	2.49	2.47
1982 annual fish meals	58.6	55.7	NR	NR
1982 serum PCB (ppb)	21.1	19.9	7.2	6.8
1982 serum total DDT (ppb)	28.8	24.5*	10.6	9.7

*p < 0.05

NR = not recorded in 1982 dataset

Natural log transformation used for mean and signif. testing

Methods

Study Population

The 1982 study by the Michigan Department of Public Health (MDPH) identified and enrolled 572 fish eaters and 419 controls within eleven communities along the Lake Michigan shoreline. Fish eaters were defined as individuals who typically consumed 24 or more pounds of sport-caught Great Lakes fish annually. These individuals were identified and recruited through MDPH staff visits to area marinas, piers, bait shops, sporting goods stores, and fishing clubs, as well as referrals through other participants. Controls were defined as individuals whose self-reported annual consumption of sport-caught Great Lakes fish was less than 6 pounds. Approximately 1,000 households were contacted by random digit dialing to obtain 419 age- and region-matched controls.

Resource limitations restricted the 1989 follow-up study to the Southern Region of the 1982 MDPH study. This region contained 36.2% of the entire 1982 cohort, and included 197 fish eaters (34.4% of all fish eaters) and 162 controls (38.7% of all controls). A comparison of 1982 baseline characteristics is presented in Table 1. The Southern Region eligible population was demographically comparable to the Entire Cohort population. Although mean serum PCB and DDT levels were slightly lower for the Southern Region groups, only the difference in mean serum DDT levels between fish eaters was statistically significant. This difference was not viewed as a biologically important difference.

Letters were sent to the 1982 Southern Region subjects which described the current study and invited continued participation. MDPH staff members conducted personal interviews and collected blood specimens from participating subjects. The questionnaire used in the 1989 personal interview was similar to the 1982 MDPH questionnaire. Information was obtained on the frequency, amount, and changes in sport-caught Great Lakes fish consumption, possible sources of occupational exposures, and lifestyles and demographic characteristics. Study participation was voluntary, without compensation, and was accompanied by the signing of an informed consent statement approved by both Human Subjects Review Committees of The University of Michigan and the Michigan Department of Public Health.

Contact was made with 169 fish eaters (85.8%) and 127 controls (78.4%). Of these eligible participants, 115 fish eaters (68.1%) and 95 controls (74.8%) were full participants—individuals who completed the personal interview and provided a blood specimen. Twenty-six fish eaters (15.4%) and 21 controls (16.5%) declined further participation in the study. The remaining 28 fish eaters and 16 controls agreed to

Table 2. Comparison of mean baseline characteristics of Southern Region and Full Participant Fish eaters and Controls

	Fish eaters		Controls	
	Southern Region	1989 Full Participants	Southern Region	1989 Full Participants
N	197	115	162	95
1982 age (yrs)	45.6	46.0	42.3	44.3
% male	63.5	64.3	49.4	46.3
1982 Quetelet Index	2.56	2.60	2.47	2.48
1982 annual fish meals	55.7	53.5	NR	NR
1982 serum PCB (ppb)	19.9	20.5	6.8	6.6
1982 serum total DDT (ppb)	24.5	25.8	9.7	9.6

NR = not recorded in 1982 dataset

Natural log transformation used for mean and signif. testing

complete the questionnaire but were unable or unwilling to provide a blood specimen. Table 2 presents the 1982 baseline characteristics of the Southern Region cohort and the 1989 Full Participant sample. No significant differences between the regional cohort and the full participant sample were observed, for either fish eaters or controls.

Laboratory Methods

Serum PCB and DDT analyses were performed by the MDPH Environmental Laboratory using the modifications of the Webb-McCall packed-column gas chromatography methodology developed for the 1982 fish eater study (Webb and McCall 1973; Needham *et al.* 1981; Humphrey 1983b). Peaks were identified by relative retention times and quantified by mean weight percent factors (Sawyer 1978), using Aroclor® 1260 as the reference standard. The total serum DDT value was obtained by multiplying the serum *p,p'*DDE content by 1.114 and adding this value to the serum *p,p'*DDT content. The detection limits for PCB and total DDT analyses were 3 ppb and 1 ppb, respectively. Samples with serum PCB levels below the detection limit ($n = 7$) were set to 2.0 ppb in subsequent analyses. All samples had total serum DDT levels above the detection limit.

Testing of randomly selected duplicates representing 10% of serum samples demonstrated good reliability of laboratory analysis. The technical error (Kahn and Sempos 1989) for PCB measurement was 1.4, which represented 8.1% of the total variation among individuals. The technical error for total DDT measurement was 0.9 and represented 6.3% of the total variation among individuals.

Data Analysis

Paired 1982 and 1989 serum PCB and DDT measurements were available for 111 fish eaters and 90 controls. The difference between the two time point measurements was calculated for each subject. For descriptive purposes, these intra-individual changes over time in serum PCB and DDT levels were treated as categorical variables. Each value was categorized as increased, decreased, or no change. In order to diminish the influence of trivial differences, the "no change" category was broadened to include historical changes which were small relative to the 1982 fish eater or control values.

For analytic purposes, the intra-individual changes over time in serum PCB and DDT levels were treated as continuous variables. The means of these intra-individual differences were calculated separately

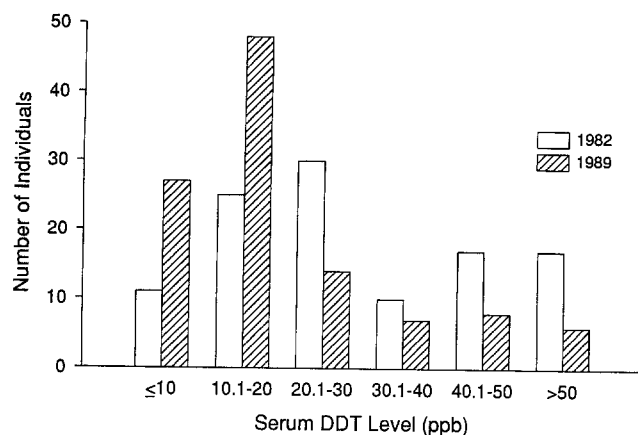


Fig. 1. Distribution of serum DDT levels of fish eaters in 1982 and 1989

for fish eaters and controls, and paired t-tests were performed to determine whether a statistically significant within-group change had occurred between 1982 and 1989. Additionally, the means of the intra-individual changes in serum PCB and DDT levels were compared across the fish eater and control groups using two-sided t-tests. Because the 1982 baseline serum PCB and DDT levels were substantially higher in the fish eater group than in the control group, the intra-individual changes were examined as both the absolute change in serum PCB and DDT levels, as well as the relative change, expressed as percent change from the 1982 level. An alpha of 0.05 was used for all tests of statistical significance.

Changes in fish consumption were determined from questionnaire data. An individual assessment of change in Great Lakes fish consumption since the 1982 study was self-reported in the 1989 questionnaire as increased, decreased, or remained the same. The associations between intra-individual changes in PCB and DDT levels and changes in fish consumption were evaluated using multiple linear regression. Models were constructed which contained the individual difference in serum PCB or DDT level from 1982 to 1989 as the dependent variable. The primary independent variable of interest in these models was the categorical change in Great Lakes fish consumption. Age, sex, and the baseline 1982 serum PCB or DDT level were also included as control variables in the regression models. All statistical analyses were conducted using programs available through SAS (SAS Institute 1985).

Results

The comparisons of the 1982 and 1989 serum DDT distributions are shown in Figures 1 and 2 for the fish eater and control groups, respectively. Since the 1982 survey, there has been an apparent downward shift of the DDT contamination level in both fish eaters and controls. Over the same time period, no changes were seen in the serum PCB distributions in either the fish eater or the control group.

Qualitatively, these same results were observed when the serum PCB and DDT level changes were treated categorically. The data presented in Table 3 demonstrate that over 60% of both fish eaters and controls experienced a decrease in serum DDT levels from 1982 to 1989, while only 7% (14/201) of the entire group exhibited an increase. In contrast, no apparent trends in serum PCB changes were seen in either fish eaters or controls.

The population mean serum PCB and DDT levels for 1982 and 1989 are presented in Table 4, in conjunction with the

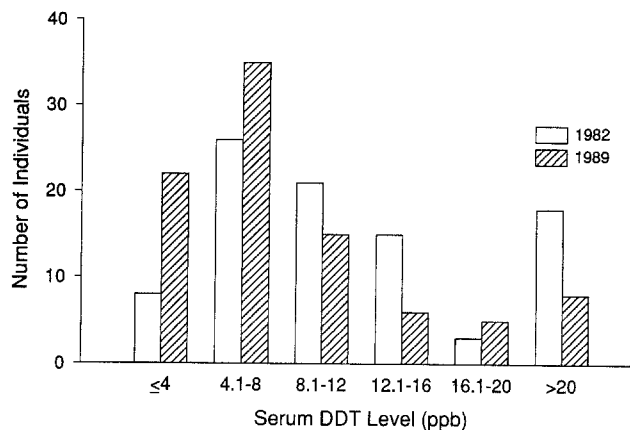


Fig. 2. Distribution of serum DDT levels in non-fishing controls in 1982 and 1989

Table 3. Distribution of serum level change categories for PCB and DDT in fish eaters and controls

	Serum PCB		Serum DDT	
	Fish eaters	Controls	Fish eaters	Controls
N	111	90	111	90
% increased	21.6 (24)	31.1 (28)	6.3 (7)	7.8 (7)
% no change	43.2 (48)	36.7 (33)	28.8 (32)	28.9 (26)
% decreased	35.1 (39)	32.2 (29)	64.9 (72)	63.3 (57)

Number of individuals in parentheses

Table 4. Mean serum PCB and DDT levels in fish eaters and controls, 1982 vs 1989

	Fish eaters			Controls		
	1982	1989	p-Value*	1982	1989	p-Value*
Serum PCB (ppb)	20.5	19.0	0.026	6.6	6.8	0.79
Serum DDT (ppb)	25.8	15.6	0.0001	9.6	6.8	0.0002

Values represent 111 fish eaters and 90 controls

*p-values for paired t-test of mean differences

results from the paired t-test analysis of the mean intra-individual historic changes. A substantial decrease in the overall population mean serum DDT level was apparent in both the fish eater and control groups. The paired t-test analysis revealed that the mean intra-individual decreases in serum DDT levels from 1982 to 1989 were statistically significant for both fish eater and control groups. In contrast, only small intra-individual decreases in serum PCB levels from 1982 to 1989 were observed, and only the mean intra-individual decline seen in the fish eater group was statistically significant.

Further comparisons of the mean intra-individual differences in serum PCB and DDT levels were made across the fish eater and control strata. For serum PCB levels, the mean intra-individual decrease seen in the fish eater group (-2.1 ppb) was not significantly different from that observed in the control group (-0.1 ppb). For serum DDT levels, the mean intra-individual decrease for the fish eater group (-13.8 ppb) was significantly

Table 5. Distribution of self-reported changes in fish consumption for fish eaters and controls

	Fisheaters	Controls
% increased	5.2 (6)	9.5 (9)
% no change	37.3 (43)	66.3 (63)
% decreased	57.4 (66)	24.2 (23)

Number of individuals in parentheses

greater than that observed in the control group (-3.1 ppb). However, when the relative magnitude of the serum DDT change was examined by expressing the intra-individual difference as the percent change from the 1982 level, there was no statistically significant mean difference between fish eaters and controls (-29.7% vs. -26.3% , respectively).

A substantial decrease was observed in the amount of Great Lakes fish consumed by the fish eater group, relative to the 1982 study. Questionnaire data indicated that this decrease resulted from a reduction in the amount of fish caught, and was not due to concern over toxins in the fish. The distributions of the self-reported changes in fish consumption are presented in Table 5. Despite these changes, the mean annual Great Lakes fish consumption of the fish eater group remained significantly higher than that of the control group (21.6 pounds vs. 1.6 pounds).

To determine whether the intra-individual historic changes in serum PCB and DDT levels were associated with changes in fish consumption, multiple linear regression analyses were performed. After controlling for the secondary independent variables of age, sex, and the baseline 1982 contaminant level, no statistically significant associations between either increased or decreased fish consumption and the observed changes in serum PCB and DDT levels were found. This lack of association persisted whether the outcome variable was expressed as either the absolute difference between the 1982 and 1989 contaminant levels, or as the percent change from the 1982 baseline level. The findings of no association between changes in fish consumption and changes in serum PCB or DDT levels were consistent in both fish eater and control groups.

Discussion

The 1989 re-examination of the 1982 MDPH fish eater study population provided a unique opportunity to evaluate historical changes in serum PCB and DDT levels within individuals. Previous studies have generally consisted of cross-sectional surveys examining biological specimens obtained from either autopsy or self-referred populations. This study was able to repeat serum PCB and DDT measurements, using identical analytic methods, in individuals with previously demonstrated contamination from a continuing low-level environmental exposure.

Earlier studies have suggested that DDT levels in western populations have been declining since the 1970's. The results from this study confirm those findings, and support the interpretation that the decline reflects the overall reduction in environmental DDT contamination (Skaare *et al.* 1989; Kutz *et al.* 1991). Since the banning of DDT use in the United States in 1972, a documented decrease in environmental and wildlife

DDT contamination has occurred. In this study, the intra-individual reductions in serum DDT levels were not associated with concurrent decreases in the frequency of the exposure—the consumption of sport-caught Great Lakes fish. This suggests that the observed lower levels may have been due to a decreased intensity of exposure, *i.e.*, lower levels of DDT in both the fish and the environment in general. This interpretation is supported by the observation that when the historic changes were expressed in terms of relative magnitude (percent change from 1982 levels), the decrease in DDT contamination was virtually the same for both the exposed fish eater group and the control group. The greater absolute loss in serum DDT experienced by the fish eaters was reflective of their higher 1982 baseline levels. These findings suggest that the gradual environmental improvement resulting from the banning of DDT has extended to human populations as well.

In contrast, although environmental and wildlife PCB levels have declined in the Great Lakes region, no consistent, comparable decline was evident in either the fish eaters or the controls. This finding agrees with earlier cross-sectional surveys which have observed stable PCB levels over time periods during which DDT levels declined dramatically.

Although the novel use of PCBs was restricted in the United States in 1974, and PCB production was halted altogether in 1977, very large quantities remain in industrial use. Environmental contamination continues to occur, as do human exposures, from a variety of sources. The more recent and less complete restrictions on PCBs, in conjunction with the long half-lives typical of most PCB mixtures, may explain the findings of this and other studies, that the restrictions on PCBs have not as yet been as effective as the restrictions on DDT in reducing the levels of contamination in human populations.

Acknowledgments. This publication is a result of work sponsored by the Michigan Sea Grant College Program, project number R/ES-4, under grant number NA89AA-D-SG083 from the office of Sea Grant, National Oceanic and Atmospheric Administration (NOAA), U.S. Department of Commerce, and funds from the State of Michigan. The U.S. Government is authorized to produce and distribute reprints for governmental purposes notwithstanding any copyright notation appearing hereon.

The authors want to thank the Michigan Department of Public Health field staff of Dave Dietel, Lana Ashley, and Linda Crispino who contacted the cohort and conducted the field clinics under the direction of Marvin Budd. We also wish to thank Robert Welch of the MDPH for the PCB and DDT analyses.

References

- D'Itri FM (1988) Contaminants in selected fishes from the upper Great Lakes. In: Schmidtke NW (ed) Toxic contamination in large lakes. Vol II. Lewis Publishers, Chelsea, MI
- Dustman EH, Stickel LF (1969) The occurrence and significance of pesticide residues in wild animals. *Ann NY Acad Sci* 160:162–172
- Fiore B, Anderson HA, Hanrahan LP, Olson LJ, Sonzogni WC (1989) Sport fish consumption and body burden levels of chlorinated hydrocarbons: A study of Wisconsin anglers. *Arch Environ Health* 44:18–24
- Foley RE, Jackling SJ, Sloan RJ, Brown MK (1988) Organochlorine and mercury residues in wild mink and otter: comparison with fish. *Environ Toxicol Chem* 7:363–374

- Frank R, Rasper J, Smout MS, Braun HE (1988) Organochlorine residues in adipose tissues, blood and milk from Ontario residents, 1976-1985. *Can J Public Health* 79:150-158
- Frank R, Braun HE (1989) PCB and DDE residues in milk supplies of Ontario, Canada 1985-1986. *Bull Environ Contam Toxicol* 42:666-669
- Horn EG, Hetling LJ, Tofflemire TJ (1979) The problem of PCBs in the Hudson River system. *Ann NY Acad Sci* 320:591-609
- Humphrey HEB (1983a) Population studies of PCBs in Michigan residents. In: D'Itri FM, Kamrin MA (eds) *PCBs: Human and environmental hazards*. Butterworth, Boston
- (1983b) Evaluation of humans exposed to water-borne chemicals in the Great Lakes. U.S. Environmental Protection Agency (Report #CR-807192), Washington, DC
- Kahn HA, Sempos CT (1989) *Statistical methods in epidemiology*. Oxford University Press, NY, p 239
- Kutz FW, Wood PH, Bottimore DP (1991) Organochlorine pesticides and polychlorinated biphenyls in human adipose tissue. *Reviews Environ Contam Toxicol* 120:1-85
- Mes J, Davies DJ, Turton D (1982) Polychlorinated biphenyl and other chlorinated hydrocarbon residues in adipose tissue of Canadians. *Bull Environ Contam Toxicol* 28:97-104
- Needham LL, Burse VW, Price HA (1981) Temperature-programmed gas chromatographic determination of PCB and PBB in serum. *J Assoc Offic Anal Chem* 64:1131-1137
- Quimby GE, Armstrong JF, Durham WF (1965) DDT in human milk. *Nature* 207:726-728
- Risebrough RW, Rieche P, Herman SG, Peakall DB, Kirven MN (1968) Polychlorinated biphenyls in the global ecosystem. *Nature* 220:1098-1102
- SAS Institute Inc* (1985) *SAS user's guide: Statistics version, 5th ed.* SAS Institute Inc, Cary, NC
- Sawyer LD (1978) Quantitation of polychlorinated biphenyl residues by electron capture gas-liquid chromatography. *J Assoc Offic Anal Chem* 61:272-281
- Schmitt CJ, Zajicek JL, Ribick MA (1985) National Pesticide Monitoring Program: Residues of organochlorine chemicals in US freshwater fish 1980-81. *Arch Environ Contam Toxicol* 14:225-260
- Schmitt CJ, Zajicek JL, Peterman PH (1990) National Contaminant Biomonitoring Program: Residues of organochlorine chemicals in US freshwater fish, 1976-1984. *Arch Environ Contam Toxicol* 91:748-781
- Schwartz PM, Jacobson SW, Fein G, Jacobson JL, Price HA (1983) Lake Michigan fish consumption as a source of polychlorinated biphenyls in human cord serum, maternal serum, and milk. *Am J Public Health* 73:293-296
- Skaare JU, Tuveng JM, Sande HA (1989) Organochlorine pesticides and polychlorinated biphenyls in maternal adipose tissue, blood, milk, and cord blood from mothers and their infants living in Norway. *Arch Environ Contam Toxicol* 17:55-63
- Veith GC, Kuehl DW, Leonard EN, Welch K, Pratt G (1981) Polychlorinated biphenyls and other organic chemical residues in fish from major watersheds near the Great Lakes. *Pestic Monit J* 5:1-8
- Webb RG, McCall AC (1973) Quantitative PCB standards for electron capture gas chromatography. *J Chromatog Sci* 11:366-373
- Zavon MR, Tye R, Latorre L (1969) Chlorinated hydrocarbon insecticide content of the neonate. *Ann NY Acad Sci* 160:196-200

Manuscript received November 18, 1991 and in revised form January 3, 1992.