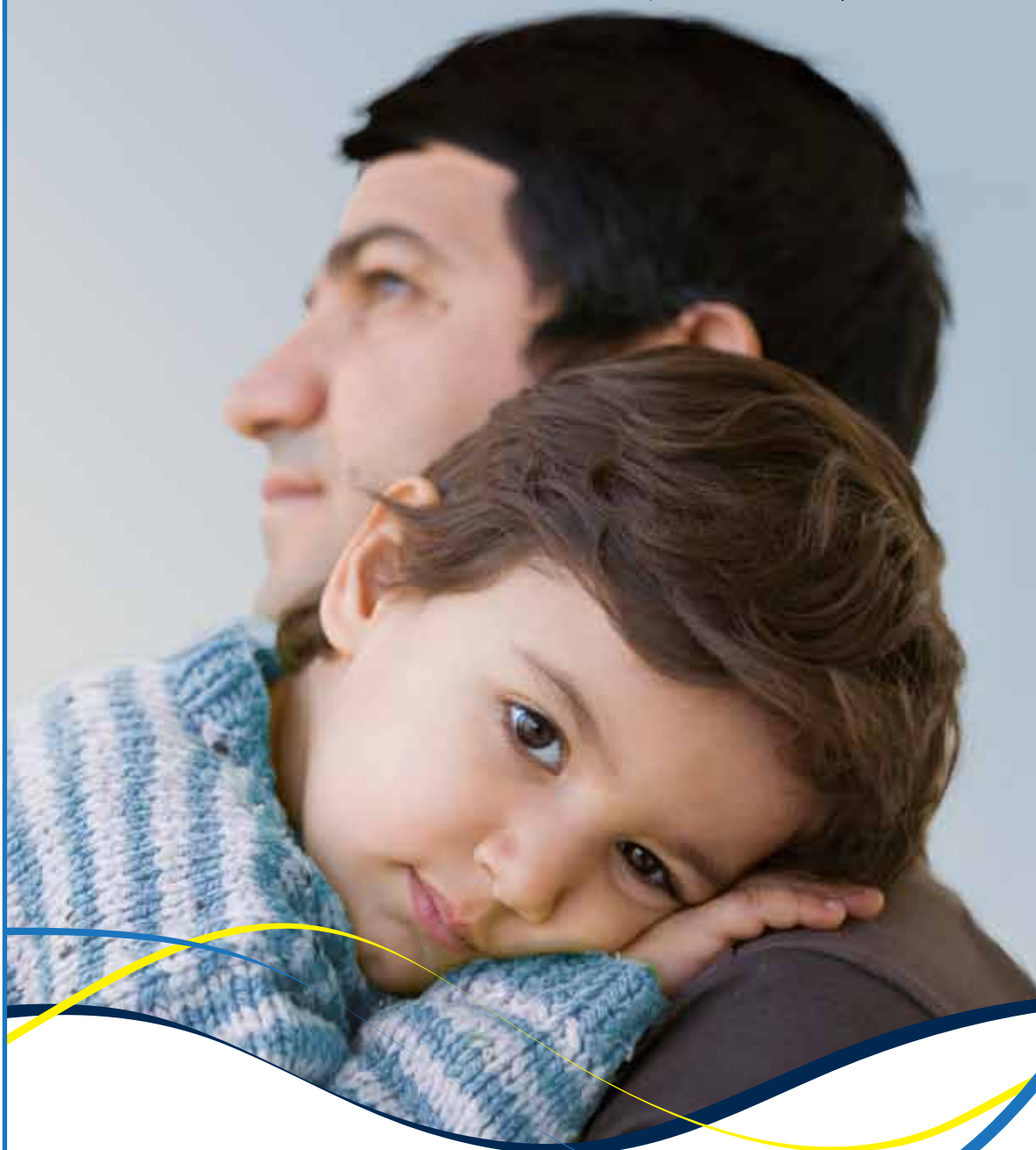


PCBs

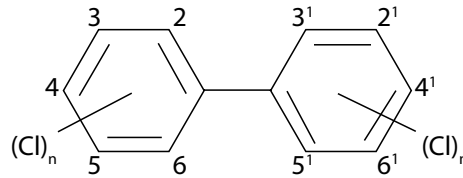
POLYCHLORINATED BIPHENYLS (PCBs) GUIDE



 **Metamatrix**
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PCBs PROFILE INTERPRETIVE GUIDE



For the PCBs measured in the Metamatrix profile, $n=2$ or 3 , so there are 4 or 5 chlorine atoms in various positions of the phenyl rings.

WHY MEASURE PCBs?

Within a span of about 20 years, more than 1.2 billion pounds of polychlorinated biphenyls (PCBs) were produced in the United States. They were used extensively as coolants and lubricants for electrical transformers and in

many other industrial uses. The production of PCBs in the United States ceased in 1977 due to build-up in the environment and association with severe health problems. Much of U.S. soil and groundwater were already heavily polluted. PCBs are very resistant to degradation by heat, chemical, or biological attack. PCBs were attractive for industrial applications

because they outlasted the equipment in which they were used. However, once introducing that property into the environment, PCBs remained extremely persistent, resisting decomposition for hundreds of years. Thus, the levels of these persistent organic pollutants (POPs) built up more and more with each year of their use. Once POPs are introduced into the environment, they undergo biomagnification as illustrated in Figure 1. PCBs have been found to have potential soil-to-human biomagnification factors of over one million.¹

Over 200 different PCBs were made before 1977, and some consumer products may still contain PCBs including old fluorescent lighting fixtures, electrical devices or appliances containing PCB capacitors, electrical transformers, as well as old microscope oil and

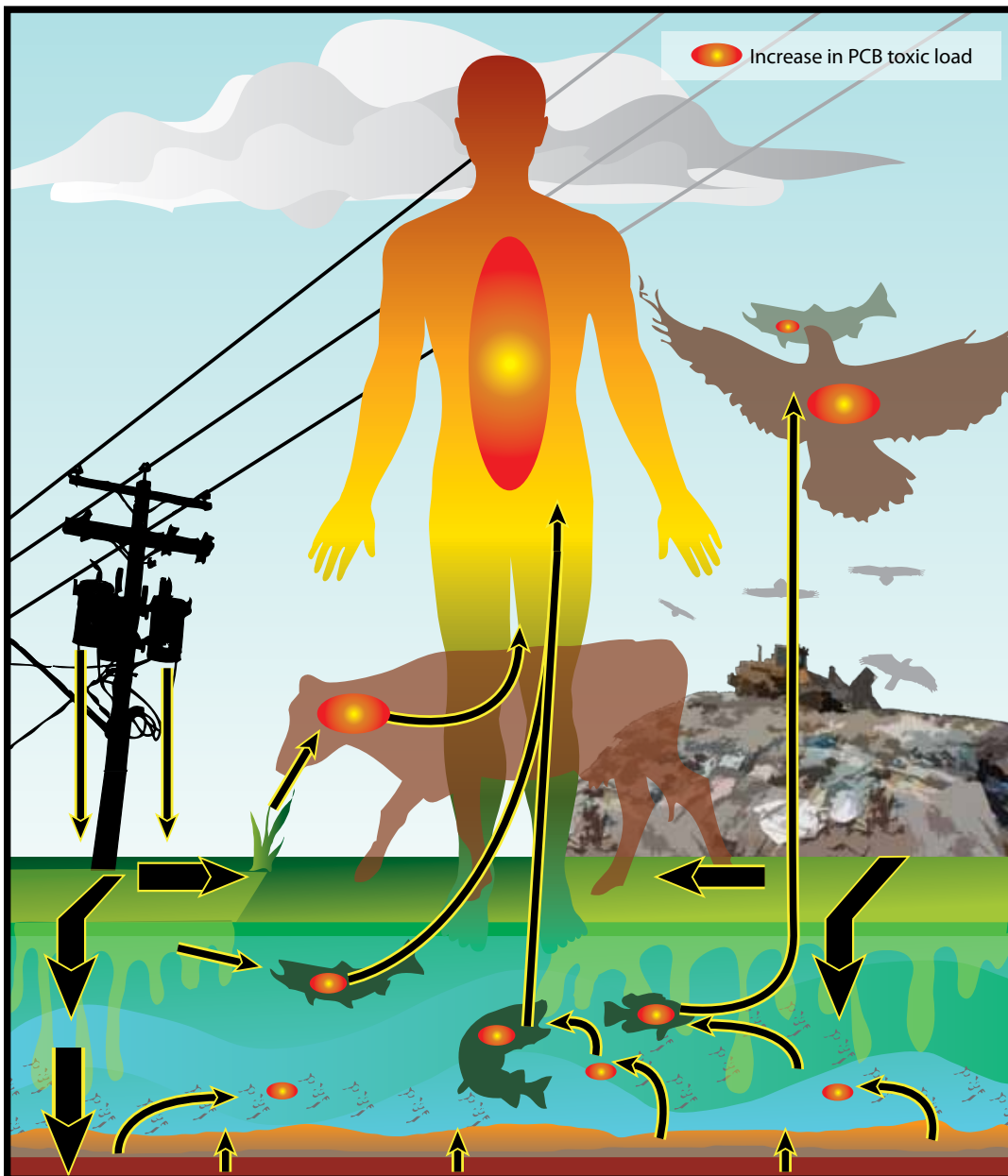


FIGURE 1 . Bioaccumulation and biomagnification of toxins that are slowly eliminated from exposed life forms.

Table 1								
PCB	CDC 50th		CDC 75th		CDC 90th		CDC 95th	
	ppb	ng/g lipid	ppb	ng/g lipid	ppb	ng/g lipid	ppb	ng/g lipid
Dioxin-like PCBs								
# 118	0.032	5.19	0.066	10.4	0.143	21.8	0.216	31.3
# 126	89.8*	14.7*	159*	24.8*	308*	46.7*	475*	68.7*
# 156	0.021	3.29	0.048	7.0	0.075	11.4	0.103	15.3
# 169	<LOD	<LOD	133*	19.5*	203*	31.0*	269*	40.6*
Non-Dioxin-like PCBs								
# 74	0.027	4.36	0.058	8.72	0.104	15.8	0.153	22.3
# 138	0.095	15.1	0.206	30.5	0.359	55.4	0.477	75.3
# 153	0.135	20.8	0.283	43.3	0.477	71.8	0.624	97.1
# 180	0.114	18.0	0.246	37.1	0.409	63.7	0.534	81.5
#194	<LOD	<LOD	.070	11.1	.123	18.1	.162	23.7

hydraulic fluids. In an Environmental Working Group study with volunteers that lead healthy lives and do not work with chemicals, the subjects contained an average of 91 compounds – most of which did not exist 75 years ago. Bill Moyers, a well-know journalist and news commentator, participated in the study. Of the 48 different PCBs tested, 31 were present in his blood.² The CDC is also measuring PCB levels in the NHANES trial and to date has been looking for 15 different PCBs.³ Of those fifteen, ten are found fairly often. Of those ten, six have known health effects published in the medical literature. These six PCBs, all frequently found in people and with documented health effects, are the ones measured in the Metamatrix PCBs Profile.

United States population distribution values (50th, 75th, 90th, and 95th percentile cutoffs) for the PCBs that the CDC measured in the 2009 Fourth National Report on Human Exposure to Environmental Chemicals are shown in Table 1. Their levels include both parts per billion (ppb) in the serum and lipid-adjusted values. The values highlighted in yellow were chosen for measurement in this panel. While PCB numbers 74, 170, 187, and 194 are also found with some regularity, Metamatrix does not measure them due to lack of evidence regarding health effects.

TABLE I. Percentile cutoff values found in the Fourth National Report on Human Exposure to Environmental Chemicals conducted by the U.S. CDC reported in 2009. LOD stands for Limit of Detection. *measured in parts per quadrillion (ppq), and picogram per gram (pg/g) lipid.

DIOXIN-LIKE POLYCHLORINATED BIPHENYLS

- PCB 118 (2,3',4,4',5-Pentachlorobiphenyl)
- PCB 126 (3,3',4,4',5-Pentachlorobiphenyl)
- PCB 156 (2,3,3',4,4',5-Hexachlorobiphenyl)
- PCB 169 (3,3',4,4',5,5'-Hexachlorobiphenyl)

NON-DIOXIN-LIKE POLYCHLORINATED BIPHENYLS

- PCB 74 (2,4,4',5-Tetrachlorobiphenyl)
- PCB 138 (2,2',3,4,4',5'-Hexachlorobiphenyl)
- PCB 153 (2,2',4,4',5,5'-Hexachlorobiphenyl)
- PCB 180 (2,2',3,4,4',5,5'-Heptachlorobiphenyl)

THE BENEFIT OF SEEING BOTH “PPB” AND “NG/G LIPID” VALUES

The levels reported in parts per billion (ppb) are reflective of the amount of toxin present in the serum, mostly found in the lipoprotein and albumin fractions.⁴ This level most likely reflects the recirculation of toxins from the adipose tissue due to diurnal lipolysis cycles. Loss of adipose tissue due to stress, rigorous exercise, or weight loss causes increased serum levels from the release of stored PCBs. Current exposures from air and food can also be detected in the serum.

All fat-soluble toxins, including PCBs, are carried in the lipid fraction of the serum, mostly in low-density lipoprotein particles (LDL). Since levels of PCBs change in direct proportion to blood lipid levels, improper test interpretations can result from examining only the concentrations in blood. Measurement of cholesterol and

triglycerides in the blood serum from the same specimen used to perform the PCB testing allows calculation of total lipid level. The PCB concentrations can then be expressed as nano gram per gram (ng/g) lipid. This method of correction has been shown to generate results that reflect adipose tissue levels of organotoxin compounds.⁵ PCBs stored in adipose tissue are a result of bio-accumulation over the lifespan. When lipolysis occurs (with fasting, exercise, stress, weight loss, and saunas) a portion of these toxins accompany the cholesterol and triglycerides that are released from the body's fat stores. By measuring the amount of lipids present in the blood and adjusting the amount of toxins to that lipid level, one is able to get a very accurate idea of just how much of those toxins are in storage. Effective cleansing or detoxification protocols will eventually produce low levels of PCBs, representing lower overall burden of these persistent and accumulating toxins.

A recent exposure may be detected as high ppb in serum, but low or undetectable ng/g lipid levels. This combination indicates that the PCBs in blood have not had time to redistribute to the adipose tissue, a process that occurs over several weeks.

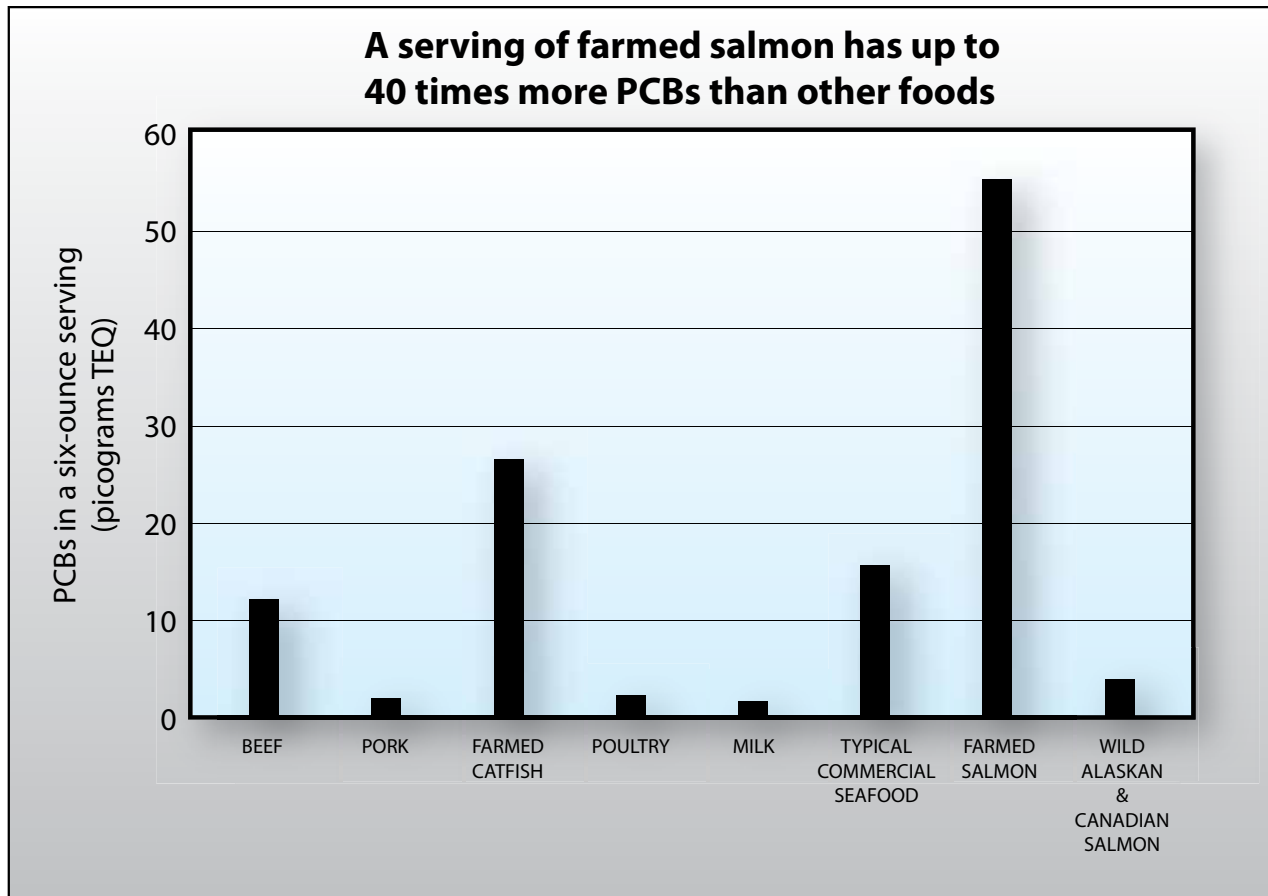
WHAT TO EXPECT

PCBs are a class of chemicals that have been found in virtually everyone tested across the planet. The question is not if one has PCBs, but how many are present and are the PCBs adversely affecting that person's health and vitality. Metamatrix is using the CDC's values found in the Fourth National Report on Human Exposure to Environmental Chemicals to give the clinician a frame of reference when deciding if a patient has higher levels than what is found in the general population.

WHAT ARE THE ADVERSE EFFECTS OF POLYCHLORINATED BIPHENYLS IN HUMANS?

IMMUNE SYSTEM EFFECTS

- PCBs (both dioxin-like and non-dioxin-like) induce apoptosis of monocytes⁶ and thymocytes.⁷
- PCBs cause white cells to have diminished mitogen response, decreased phagocytosis and diminished numbers of CD8+ cells.⁸
- Dioxin-like PCBs can cause thymic atrophy and immunosuppression.⁹
- Dietary PCB exposure can lead to increased rates of certain cancers (especially liver and lung).¹⁰



NEUROLOGIC EFFECTS

- Neonatal exposure of mice to PCBs results in persistent aberrations in spontaneous behavior that worsens as they age.¹¹ PCB exposure also adversely affected learning and memory function when they became adult mice.
- In utero exposure to PCBs affected intellectual functioning in children¹², such as increased cognitive defects, poorer gross motor function, and decreased visual recognition memory.¹³ They also have lower IQ levels and have increased rates of hyperactivity, both of these problems persist after birth.¹⁴⁻¹⁶
- Adults consuming PCB-contaminated fish had increased problems with memory and learning.¹⁷
- Persons exposed to PCB gas had chronic neurological problems including slower reaction time (for both simple and choice reactions), faster sway speeds, diminished color discrimination and visual performances, and constricted visual fields. They also had diminished scores on digit symbols, vocabulary, verbal recall, and embedded memory.¹⁸

ENDOCRINE EFFECTS

- PCBs adversely affect thyroid levels, including elevated anti-thyroid antibodies.¹⁹⁻²¹
- As PCB serum levels increase, the thyroid hormones triiodothyronine (T₃) and thyroxine (T₄) decrease.²²
- Women exposed to PCBs report higher incidence of stillbirth, miscarriage, more abnormal menstrual bleeding, and greater incidence of endometriosis than non-exposed women.^{23,24}
- PCBs and other chlorinated compounds lead to increased risk for type 2 diabetes.²⁵

ADVERSE HEALTH EFFECTS OF SPECIFIC PCBs

- The sum of PCBs 118, 138, 153, and 180 was used to demonstrate negative immune effects in children.²⁶
- PCB 153 was significantly associated with diabetes risk.^{27, 28}
- The sum of PCBs 28, 52, 101, 118, 138, 153, and 180 was associated with low T₄ and elevated Gamma Glutamyl Transpeptidase (GGT).²⁹
- PCB 118 mimics T₃ action.³⁰
- PCBs 153 and 126 can negatively affect the nervous system and thyroid gland.³¹

- PCB 126 is associated with estrogenic³² and adrenal effects³³ and is immunotoxic.³⁴

EXPOSURE SOURCES

The greatest sources of PCBs exposure come from eating contaminated food.³⁵ The estimated dietary intake of PCBs for an average adult was 0.027 ug/kg/day in 1978 and had declined to <0.001 ug/kg/day by 1991.³⁶ The highest content of PCBs in one dietary study was found in dairy products (especially butter), meat, and fish. Atlantic farmed salmon is the greatest exposure source among fish, as well as fish caught and consumed from the Great Lakes.³⁷

Inhalation of indoor air in buildings with old electrical fixtures is another route of PCB exposure.

Numerous studies from around the globe have consistently documented PCBs in breast milk samples, exposing infants who are breast fed daily.

ACTION STEPS

1. Identify exposure sources and remove them. DO NOT eat Atlantic (farmed) salmon; substitute with wild caught Alaskan salmon. Substitute commercial varieties of butter and margarine with organic butter.
2. Enhance the clearance of persistent toxins from the body with cleansing protocols. Sauna therapy and colonic irrigations have been used to reduce the presence of PCBs and chlorinated pesticides.³⁸
3. Increase the normal bowel excretion of the fat-soluble toxins. Daily use of rice bran fiber (RBF) has been documented in several studies in Japan to increase the clearance of PCBs.³⁹⁻⁴¹ Chlorophyll and all chlorophyll containing foods are also tremendous at increasing the excretion of these fat-soluble persistent toxins in the feces.⁴²⁻⁴⁴ Increasing these foods in the diet, or with supplementation on a daily basis, will slowly increase the excretion of these compounds from the body. In addition to the chlorophyll containing agents, polyphenols found highest in white and green teas have been shown to increase the excretion of fat-soluble toxins.⁴⁵
4. Supplement with high amounts of a combination nutrient and botanical antioxidant to protect the tissues and cells that are under assault from the toxic compounds.

5. Assess detoxification ability with organic acid analysis, amino acid analysis, and genetic predisposition testing. These tests can aid in designing appropriate detoxification protocols for patients.

REFERENCES

1. Jensen AA, Jorgensen KF. Polychlorinated terphenyls (PCTs) use, levels and biological effects. *Sci Total Environ*. Apr 1983;27(2-3):231-250.
2. <http://archive.ewg.org/reports/bodyburden1/>.
3. Exposure Report. www.cdc.gov/exposurereport/pdf/FourthReport.pdf.
4. Noren K, Weistrand C, Karpe F. Distribution of PCB congeners, DDE, hexachlorobenzene, and methylsulfonyl metabolites of PCB and DDE among various fractions of human blood plasma. *Arch Environ Contam Toxicol*. Oct 1999;37(3):408-414.
5. Patterson DG, Jr., Needham LL, Pirkle JL, et al. Correlation between serum and adipose tissue levels of 2,3,7,8-tetrachlorodibenzo-p-dioxin in 50 persons from Missouri. *Arch Environ Contam Toxicol*. Mar 1988;17(2):139-143.
6. Shin KJ, Bae SS, Hwang YA, Seo JK, Ryu SH, Suh PG. 2,2',4,6,6'-pentachlorobiphenyl induces apoptosis in human monocytic cells. *Toxicol Appl Pharmacol*. Nov 15 2000;169(1):1-7.
7. Tan Y, Li D, Song R, Lawrence D, Carpenter DO. Ortho-substituted PCBs kill thymocytes. *Toxicol Sci*. Dec 2003;76(2):328-337.
8. Fournier M, Degas V, Colborn T, et al. Immunosuppression in mice fed on diets containing beluga whale blubber from the St Lawrence estuary and the Arctic populations. *Toxicol Lett*. Mar 15 2000;112-113:311-317.
9. Davis D, Safe S. Immunosuppressive activities of polychlorinated biphenyls in C57BL/6N mice: structure-activity relationships as Ah receptor agonists and partial antagonists. *Toxicology*. Jul 1990;63(1):97-111.
10. Onozuka D, Yoshimura T, Kaneko S, Furue M. Mortality after exposure to polychlorinated biphenyls and polychlorinated dibenzofurans: a 40-year follow-up study of Yusho patients. *Am J Epidemiol*. Jan 1 2009;169(1):86-95.
11. Eriksson P, Fredriksson A. Neurotoxic effects in adult mice neonatally exposed to 3,3',4,4',5-pentachlorobiphenyl or 2,3,3',4,4'-pentachlorobiphenyl. Changes in brain nicotinic receptors and behaviour. *Environ Toxicol Pharmacol*. 1998;5:17-27.
12. Jacobson SW, Fein GG, Jacobson JL, Schwartz PM, Dowler JK. The effect of intrauterine PCB exposure on visual recognition memory. *Child Dev*. Aug 1985;56(4):853-860.
13. Jacobson JL, Jacobson SW. Evidence for PCBs as neurodevelopmental toxicants in humans. *Neurotoxicology*. 1997;18(2):415-424.
14. Chen YC, Guo YL, Hsu CC, Rogan WJ. Cognitive development of Yu-Cheng ("oil disease") children prenatally exposed to heat-degraded PCBs. *Jama*. Dec 9 1992;268(22):3213-3218.
15. Chen YC, Yu ML, Rogan WJ, Gladen BC, Hsu CC. A 6-year follow-up of behavior and activity disorders in the Taiwan Yu-cheng children. *Am J Public Health*. Mar 1994;84(3):415-421.
16. Lai TJ, Liu X, Guo YL, et al. A cohort study of behavioral problems and intelligence in children with high prenatal polychlorinated biphenyl exposure. *Arch Gen Psychiatry*. Nov 2002;59(11):1061-1066.
17. Schantz SL, Gasior DM, Polverejan E, et al. Impairments of memory and learning in older adults exposed to polychlorinated biphenyls via consumption of Great Lakes fish. *Environ Health Perspect*. Jun 2001;109(6):605-611.
18. Kilburn KH. Visual and neurobehavioral impairment associated with polychlorinated biphenyls. *Neurotoxicology*. Aug 2000;21(4):489-499.
19. Gerhard I, Monga B, Krahe J, Runnebaum B. Chlorinated hydrocarbons in infertile women. *Environ Res*. May 1999;80(4):299-310.
20. Langer P, Tajtakova M, Fodor G, et al. Increased thyroid volume and prevalence of thyroid disorders in an area heavily polluted by polychlorinated biphenyls. *Eur J Endocrinol*. Oct 1998;139(4):402-409.
21. Li MH, Hansen LG. Enzyme induction and acute endocrine effects in prepubertal female rats receiving environmental PCB/PCDF/PCDD mixtures. *Environ Health Perspect*. Jul 1996;104(7):712-722.
22. Hagmar L, Rylander L, Dyremark E, Klasson-Wehler E, Erfurth EM. Plasma concentrations of persistent organochlorines in relation to thyrotropin and thyroid hormone levels in women. *Int Arch Occup Environ Health*. Apr 2001;74(3):184-188.
23. Yu ML, Guo YL, Hsu CC, Rogan WJ. Menstruation and reproduction in women with polychlorinated biphenyl (PCB) poisoning: long-term follow-up interviews of the women from the Taiwan Yucheng cohort. *Int J Epidemiol*. Aug 2000;29(4):672-677.
24. Leoni V, Fabiani L, Marinelli G, et al. PCB and other organochlorine compounds in blood of women with or without miscarriage: a hypothesis of correlation. *Ecotoxicol Environ Saf*. Feb 1989;17(1):1-11.
25. Lee DH, Lee IK, Jin SH, Steffes M, Jacobs DR, Jr. Association between serum concentrations of persistent organic pollutants and insulin resistance among nondiabetic adults: results from the National Health and Nutrition Examination Survey 1999-2002. *Diabetes Care*. Mar 2007;30(3):622-628.
26. Weisglas-Kuperus N, Patandin S, Berbers GA, et al. Immunologic effects of background exposure to polychlorinated biphenyls and dioxins in Dutch preschool children. *Environ Health Perspect*. Dec 2000;108(12):1203-1207.
27. Lee DH, Lee IK, Song K, et al. A strong dose-response relation between serum concentrations of persistent organic pollutants and diabetes: results from the National Health and Examination Survey 1999-2002. *Diabetes Care*. Jul 2006;29(7):1638-1644.
28. Rylander L, Rignell-Hydbom A, Hagmar L. A cross-sectional study of the association between persistent organochlorine pollutants and diabetes. *Environ Health*. 2005;4:28.
29. Sala M, Sunyer J, Herrero C, To-Figueras J, Grimalt J. Association between serum concentrations of hexachlorobenzene and polychlorobiphenyls with thyroid hormone and liver enzymes in a sample of the general population. *Occup Environ Med*. Mar 2001;58(3):172-177.

30. Fritsche E, Cline JE, Nguyen NH, Scanlan TS, Abel J. Polychlorinated biphenyls disturb differentiation of normal human neural progenitor cells: clue for involvement of thyroid hormone receptors. *Environ Health Perspect.* Jul 2005;113(7):871-876.
31. Costa LG, Fattori V, Giordano G, Vitalone A. An in vitro approach to assess the toxicity of certain food contaminants: methylmercury and polychlorinated biphenyls. *Toxicology.* Jul 31 2007;237(1-3):65-76.
32. Matthews J, Wihlen B, Heldring N, et al. Co-planar 3,3',4,4',5-pentachlorinated biphenyl and non-co-planar 2,2',4,6,6'-pentachlorinated biphenyl differentially induce recruitment of oestrogen receptor alpha to aryl hydrocarbon receptor target genes. *Biochem J.* Sep 1 2007;406(2):343-353.
33. Li LA, Wang PW. PCB126 induces differential changes in androgen, cortisol, and aldosterone biosynthesis in human adrenocortical H295R cells. *Toxicol Sci.* May 2005;85(1):530-540.
34. Riecke K, Schmidt A, Stahlmann R. Effects of 2,3,7,8-TCDD and PCB 126 on human thymic epithelial cells in vitro. *Arch Toxicol.* Jun 2003;77(6):358-364.
35. Zuccato E, Calvarese S, Mariani G, et al. Level, sources and toxicity of polychlorinated biphenyls in the Italian diet. *Chemosphere.* May 1999;38(12):2753-2765.
36. Gunderson EL. Dietary intakes of pesticides, selected elements, and other chemicals: FDA Total Diet Study, June 1984-April 1986. *J AOAC Int.* Jul-Aug 1995;78(4):910-921.
37. Anderson HA, Falk C, Hanrahan L, et al. Profiles of Great Lakes critical pollutants: a sentinel analysis of human blood and urine. The Great Lakes Consortium. *Environ Health Perspect.* May 1998;106(5):279-289.
38. Crinnion WJ. Unpublished research. Southwest College of Naturopathic Medicine.
39. Morita K, Hamamura K, Iida T. [Binding of PCB by several types of dietary fiber in vivo and in vitro]. *Fukuoka Igaku Zasshi.* May 1995;86(5):212-217.
40. Morita K, Hirakawa H, Matsueda T, Iida T, Tokiwa H. [Stimulating effect of dietary fiber on fecal excretion of polychlorinated dibenzofurans (PCDF) and polychlorinated dibenzo-p-dioxins (PCDD) in rats]. *Fukuoka Igaku Zasshi.* May 1993;84(5):273-281.
41. Nagayama J, Takasuga T, Tsuji H, Umehara M, Sada T, Iwasaki T. Active elimination of causative PCDFs/DDs congeners of Yusho by one year intake of FBRA in Japanese people. *Fukuoka Igaku Zasshi.* May 2003;94(5):118-125.
42. Morita K, Matsueda T, Iida T. [Effect of green vegetable on digestive tract absorption of polychlorinated dibenzo-p-dioxins and polychlorinated dibenzofurans in rats]. *Fukuoka Igaku Zasshi.* May 1999;90(5):171-183.
43. Morita K, Matsueda T, Iida T, Hasegawa T. Chlorella accelerates dioxin excretion in rats. *J Nutr.* Sep 1999;129(9):1731-1736.
44. Morita K, Ogata M, Hasegawa T. Chlorophyll derived from Chlorella inhibits dioxin absorption from the gastrointestinal tract and accelerates dioxin excretion in rats. *Environ Health Perspect.* Mar 2001;109(3):289-294.
45. Hsu TF, Kusumoto A, Abe K, et al. Polyphenol-enriched oolong tea increases fecal lipid excretion. *Eur J Clin Nutr.* Nov 2006;60(11):1330-1336.



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