A Reassessment of the Nomenclature of Polychlorinated Biphenyl (PCB) Metabolites

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Polychlorinated biphenyls (PCBs) are a widespread class of persistent organic chemicals that accumulate in the environment and humans and are associated with a broad spectrum of health effects. PCB biotransformation has been shown to lead to two classes of PCB metabolites that are present as contaminant residues in the tissues of selected biota: hydroxylated (HO) and methyl sulfone (MeSO₂) PCBs. Although these two types of metabolites are related structures, different rules for abbreviation of both classes have emerged. It is important that a standardized nomenclature for the notation of PCB metabolites be universally agreed upon. We suggest that the full chemical name of the PCB metabolite and a shorthand notation should be adopted using the International Union of Pure and Applied Chemistry's chemical name/original Ballschmiter and Zell number of the parent congener, followed by the assignment of the phenyl ring position number of the MeSO2- or HO-substituent. This nomenclature provides a clear, unequivocal set of rules in naming and abbreviating the PCB metabolite structure. Furthermore, this unified PCB metabolite nomenclature approach can be extended to the naming and abbreviation of potential metabolites of structurally analogous contaminants such as HO-polybrominated biphenyls and HO-polybrominated diphenyl ethers. Key words: hydroxylated metabolites, methyl sulfone metabolites, nomenclature, polychlorinated biphenyls. Environ Health Perspect 112:291-294 (2004). doi:10.1289/ehp.6409 available via http://dx.doi.org/ [Online 3 December 2003]

Nomenclature of Polychlorinated Biphenyls

Polychlorinated biphenyls (PCBs) are a class of chemical compounds in which 1–10 chlorine atoms are attached to a biphenyl backbone. Theoretically, 209 discrete congeners are possible (Ballschmiter and Zell 1980). However, PCB technical mixtures are composed of a smaller suite of congeners (Frame et al. 1996), and only about 80–100 PCB congeners are of actual environmental relevance (de Voogt et al. 1990).

The full chemical notation for these 209 possible PCB congeners is inconvenient, and therefore various shorthand notations have been developed and adopted (Erickson 1997, 2001). Ballschmiter and Zell (1980) originally introduced a system (BZ) in which congeners were arranged in the ascending numerical order based on the number of chlorine atoms and their substitution pattern on the biphenyl base structure. Minor theoretical discrepancies in the BZ naming system were later corrected (Ballschmiter et al. 1992; Guitart et al. 1993; Schulte and Malisch 1983). The BZ system of PCB shorthand notation was subsequently recognized by the International Union of Pure and Applied Chemistry (IUPAC) (U.S. Environmental Protection Agency 2003) and is the generally accepted notation used by scientists who perform congener-specific PCB research.

Metabolism of PCBs

PCBs that accumulate in biota are subject to elimination processes that are facilitated by

processes including enzyme-mediated degradation. The mechanism and kinetics of PCB biotransformation depend on a number of factors, including the metabolic capacity of the organism and the PCB congener structure. PCB biotransformation has been shown to lead to two classes of PCB metabolites that are present as contaminant residues in the tissues of biota that have been studied: hydroxylated (HO) and methyl sulfone (CH₃SO₂) PCBs (Letcher et al. 2000a). CH₃SO₂-PCBs are generally referred to as MeSO₂-PCBs. The numbers of animals and populations where tissue residues of these PCB metabolites have been characterized remains small, but HO-PCBs and MeSO₂-PCBs are emerging as common contaminant phenomena in wildlife and humans and are of increasing importance in risk assessments of exposure to PCBs (Bennett et al. 2002; Campbell et al. 2003; Chu et al. 2002, 2003; Guvenius et al. 2002; Hoekstra et al. 2003; Hovander et al. 2002; Letcher et al. 2000a, 2000b; Li et al. 2003; Sandala et al., in press; Sandau et al. 2000a, 2000b, 2002; Stapleton et al. 2001). In some species and tissues, HO-PCB and MeSO₂-PCB concentrations may be in a similar or higher range with respect to the concentrations of the parent PCBs. Furthermore, congeners of these two classes of PCB metabolites, which are present as contaminant residues, have demonstrated biologic and toxicologic activity-for example, endocrine-related activity (Brouwer et al. 1998; Letcher et al. 2000a, 2002).

Nomenclature of HO-PCBs

The published reports on HO-PCBs and MeSO₂-PCBs have generally used IUPAC guidelines to describe the full chemical name of these metabolites. However, the presently used abbreviations for HO-PCB congeners deviate from the general IUPAC naming rules. The HO-functional group is not given numbering priority on the biphenyl backbone; rather, the chlorine pattern on the biphenyl ring determines the congener number according to the BZ or IUPAC PCB numbering rules (Ballschmiter and Zell 1980; Ballschmiter et al. 1992; Guitart et al. 1993; Schulte and Malisch 1983), and the HO-group(s) are numbered thereafter. As a result, an HO-functionality in the meta-position relative to the central carbon-carbon bond of the biphenyl attachment is in either position 3 or position 5. When the HO-substituent is located on the ring with the lowest chlorine numbering priority, its number is primed in the same manner as is done for the chlorine atoms on the same phenyl ring. The HO-metabolite 5'-HO-2,3',4,4'-tetrachlorobiphenyl (three unsubstituted meta-positions in the 3, 5, and 5' positions of the corresponding PCB congener), for example, is therefore uniquely abbreviated to 5'-HO-CB66 using this notation approach (Table 1).

Nomenclature of MeSO₂-PCBs

Similarly, for MeSO₂-PCB congeners, the PCB number is first determined according to the chlorine substitution of the biphenyl by omitting the MeSO₂-functional group. The initial shorthand notation used for MeSO2-PCBs assigned the methyl sulfonyl group based on a higher numbering priority on the biphenvl system than for chlorine atoms, and thus the position of methyl sulfonyl substitution was not primed (Letcher et al. 1995; Weistrand and Norén 1997). For example, using this initial numbering approach, a methyl sulfonyl group in a meta-position would always be assigned to position 3. This nomenclature approach can be problematical, as illustrated by the example of 3-MeSO₂-2,2',4',5-tetrachlorobiphenyl (Table 1),

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which is intended when the short notation 3-MeSO₂-CB49 is used. Although this appears to be the only environmentally relevant possibility, two other metabolites would have exactly the same 3-MeSO₂-CB49 abbreviation if the methyl sulfonyl group were positioned on either of the two free *meta*-positions of the other, 2,4-chloro–substituted, phenyl ring.

Different authors have acknowledged the inconsistencies in the naming of MeSO₂-PCBs and have suggested alternate approaches (Letcher et al. 2000a and references therein). In several recent reports, the number of the MeSO₂-group has been primed or unprimed depending on which phenyl ring this substituent was positioned (Guvenius et al. 2002; Hoekstra et al. 2003; Letcher et al. 2000a, 2000b). Following this revised nomenclature system, 3-MeSÕ₂-2,2′,4′,5-tetrachlorobiphenyl is abbreviated to 3'-MeSO₂-CB49. Because the number of the methyl sulfonyl group is primed, and the other meta-carbon position on the primed phenyl ring is substituted with a chlorine atom, the abbreviation indicates only one structural possibility. Because methyl sulfonyl groups on the most commonly encountered congener residues in biota usually occur on a 2,5- or 2,3,6- chlorine-substituted phenyl ring, one meta-position will normally be occupied by a chlorine atom on the MeSO2substituted phenyl ring. It is clear that this might not be the case for some other, at least theoretical, metabolites.

Recently, Larsson et al. (2002) applied the same nomenclature rules for abbreviation of MeSO₂-metabolites as is normally done for HO-PCBs. That is, the methyl sulfonyl groups are numbered according to their substitution position, after the positions of the chlorine atoms are taken into account based on the revised BZ naming system. For example, Larsson et al. (2002) abbreviated 3-MeSO₂-2,2',3,4',5',6-hexachlorobiphenyl to 5-MeSO₂-CB149 rather than to 3-MeSO₂-CB149 (Table 1) because the MeSO₂functional group is present in position 5 rather

than in position 3. This abbreviation indicates only one distinct congener regardless of its chlorine substitution pattern, which eliminates the possibility of misidentifying MeSO₂-PCB structures with the same chlorine substitution pattern. Because the MeSO₂-group is not assigned the lowest possible number, the substitution position of the chlorine atoms is also maintained. Consequently, the identity of the MeSO₂-PCB congener is easily related to the parent PCB structure (Table 1). However, the implementation of this nomenclature approach for methyl sulfonyl-PCB metabolites poses a problem with respect to congener-specific comparisons in earlier publications where alternate nomenclature has been used. For example, some environmentally relevant 3-MeSO2-CBs would have to be renamed as 5-MeSO₂-CBs (Table 2).

Proposed Nomenclature

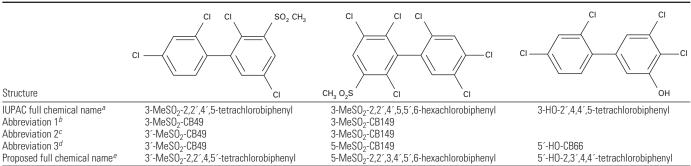
On closer examination of the full chemical name of, for example, 3-MeSO2-2,2',3,4',5',6hexachlorobiphenyl, it is striking that the MeSO₂-group receives the same number (3) as a chlorine substituent present on the same ring. Therefore, it is evident from the full chemical name of PCB metabolites that it is not possible to combine both the fundamental IUPAC approach for naming aromatic compounds (i.e., giving the substituent numbering priority and numbering the chlorine atoms thereafter) and the IUPAC-accepted BZ rules for the naming of PCBs (i.e., a PCB-BZ number is clearly associated with the chlorine substitution pattern). Because the BZ system is so widely adopted, it would be logical to base the PCB metabolite nomenclature on the BZ system. Even though this approach requires exceptions to the IUPAC guidelines of nomenclature, we would suggest that the full chemical name/shorthand notation of the metabolite should be made by using the IUPAC full name/original BZ number of the parent congener and then assigning the MeSO₂- or HO-substituent a ring position

number thereafter. For example, 5-MeSO₂-CB149 is the abbreviation for 5-MeSO₂-2,2',3,4',5',6-hexachlorobiphenyl (Table 1), where the 5-position of the substituent remains the same as in the abbreviation, and the chlorine pattern clearly indicates PCB number 149 (BZ). Table 2 shows how the proposed standardized nomenclature applies to the current list of identified and environmentally relevant HO- and MeSO2-PCB congeners. It may be observed that, for MeSO₂-PCB metabolites, a 2,5-ring substitution of the PCB congener leads to a 3- or a 3'substitution for the MeSO₂-group, whereas a 2,3,6-ring substitution leads to a 5- or a 5'-substitution (Table 2).

Table 2 can be used as a reference to unambiguously label the names of PCB metabolites. This nomenclature for both the abbreviation and full chemical name of a given PCB metabolite provides a clear, unequivocal structure and delivers a unified technique that can be used for both classes of PCB metabolites.

Although HO-PCBs and MeSO₂-PCBs are related structures, different rules for abbreviations of both classes have emerged. A standardized nomenclature that is presently suggested for the naming of a PCB metabolite should be universally adopted and sanctioned by IUPAC to facilitate unambiguous comparison of congener-specific data among published studies. Additionally, this nomenclature may be applicable to similar metabolites of other persistent aromatic organics, such as polybrominated diphenyl ethers (PBDEs) and polybrominated biphenyls (PBBs). Studies on the formation of metabolites from the PBB and PBDE classes of pollutants are becoming more numerous in the literature (Burreau et al. 2000; Haglund et al. 1997; Hakk and Letcher 2003; Meerts et al. 2001; Valters et al. 2003). Therefore, it is important to adopt a general, unified nomenclature system for the full chemical name and shorthand notation of possible metabolites.

Table 1. Structure, abbreviations, and full chemical names of three PCB metabolites



^aMeSO₂-PCBs (Larsson et al. 2002; Weistrand and Norén 1997); HO-PCBs (Bergman et al. 1994; Hovander et al. 2002). ^bMeSO₂-PCBs (Letcher et al. 1995; Weistrand and Norén 1997). ^cMeSO₂-PCBs (Guvenius et al. 2002; Hoekstra et al. 2003; Letcher et al. 2000a, 2000b). ^dMeSO₂-PCBs (Larsson et al. 2002); HO-PCBs (Bennett et al. 2002; Campbell et al. 2003; Hovander et al. 2002; Li et al. 2003; Sandala et al., in press; Sandau et al. 2000a; Sjödin et al. 1998). ^eHO-PCBs not previously used for MeSO₂-PCBs (Campbell et al. 2003; Sandala et al., in press; Sandau et al. 2000a; Sjödin et al. 2000a; Sjödin et al. 2000a; Sjödin et al. 2000a; Sjödin et al. 2000a; Sigidin et al. 2000a; Sjödin et al. 2000a; Sjödin

Table 2. Current and proposed PCB metabolite nomenclature.

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JPAC full chemical name ^a	Abbreviation ^b	Proposed full chemical name ^c
-MeSO ₂ -2,4,5-trichlorobiphenyl	3-MeSO ₂ -CB31	3-MeSO ₂ -2,4´,5-trichlorobiphenyl
-MeSO ₂ -2,4 [′] ,5-trichlorobiphenyl	4-MeSO ₂ -CB31	4-MeSO ₂ -2,4 ² ,5-trichlorobiphenyl
MeSO ₂ -2,2',4',5-tetrachlorobiphenyl	3'-MeSO ₂ -CB49	3'-MeSO ₂ -2,2',4,5'-tetrachlorobiphenyl
MeSO ₂ -2,2',4',5-tetrachlorobiphenyl	4'-MeSO ₂ -CB49	4'-MeSO ₂ -2,2',4,5'-tetrachlorobiphenyl
MeSO ₂ -2,2',5,5'-tetrachlorobiphenyl	3-MeSO ₂ -CB52	3-MeSO ₂ -2,2´,5,5´-tetrachlorobiphenyl
MeSO ₂ -2,2',5,5'-tetrachlorobiphenyl	4-MeSO ₂ -CB52	4-MeSO ₂ -2,2´,5,5´-tetrachlorobiphenyl
MeSO ₂ -2,4',5,6-tetrachlorobiphenyl	5-MeSO ₂ -CB64	5-MeSO ₂ -2,3,4 ² ,6-tetrachlorobiphenyl
MeSO ₂ -2,3,4´,6-tetrachlorobiphenyl	4-MeSO ₂ -CB64	4-MeSO ₂ -2,3,4 ⁻ ,6-tetrachlorobiphenyl
MeSO ₂ -2,3 [°] ,4 [°] ,5-tetrachlorobiphenyl	3-MeSO ₂ -CB70	3-MeSO ₂ -2,3 [°] ,4 [°] ,5-tetrachlorobiphenyl
MeSO ₂ -2,3´,4´,5-tetrachlorobiphenyl	4-MeSO ₂ -CB70	4-MeSO ₂ -2,3 [°] ,4 [°] ,5-tetrachlorobiphenyl
MeSO ₂ -2,2´,3´,4´,5-pentachlorobiphenyl	3'-MeSO ₂ -CB87	3 ⁻ MeSO ₂ -2,2 ⁻ ,3,4,5 ⁻ -pentachlorobiphenyl
MeSO ₂ -2,2',3',4',5-pentachlorobiphenyl	4'-MeSO ₂ -CB87	4 ⁻ -MeSO ₂ -2,2 ⁻ ,3,4,5 ⁻ -pentachlorobiphenyl
MeSO ₂ -2,2',4',5,6-pentachlorobiphenyl	5-MeSO ₂ -CB91	5-MeSO ₂ -2,2 [°] ,3,4 [°] ,6-pentachlorobiphenyl
MeSO ₂ -2,2',3,4',6-pentachlorobiphenyl	4-MeSO ₂ -CB91	4-MeSO ₂ -2,2',3,4',6-pentachlorobiphenyl
MeSO ₂ -2,2',3',5,6'-pentachlorobiphenyl	3'-MeSO ₂ -CB95	3 ⁻ MeSO ₂ -2,2 ⁻ ,3,5 ⁻ ,6-pentachlorobiphenyl
MeSO ₂ -2,2',3',5,6'-pentachlorobiphenyl	4'-MeSO ₂ -CB95	4'-MeSO ₂ -2,2',3,5',6-pentachlorobiphenyl
MeSO ₂ -2,2',4',5,5'-pentachlorobiphenyl	3'-MeSO ₂ -CB101	3'-MeSO ₂ -2,2',4,5,5'-pentachlorobiphenyl
MeSO ₂ -2,2 ⁽⁴⁾ ,5,5 ⁽⁻⁾ pentachlorobiphenyl	4'-MeSO ₂ -CB101	4 ⁻ -MeSO ₂ -2,2 ⁻ ,4,5,5 ⁻ -pentachlorobiphenyl
MeSO ₂ -2,3´,4´,5,6-pentachlorobiphenyl	5-MeSO ₂ -CB110	5-MeSO ₂ -2,3,3 [°] ,4 [°] ,6-pentachlorobiphenyl
MeSO ₂ -2,3,3',4',6-pentachlorobiphenyl	4-MeSO ₂ -CB110	4-MeSO ₂ -2,3,3 [°] ,4 [°] ,6-pentachlorobiphenyl
MeSO ₂ -2,2 [°] ,3 [°] ,4 [°] ,5,6-hexachlorobiphenyl	5'-MeSO ₂ -CB132	5 ⁻ -MeSO ₂ -2,2 ⁻ ,3,3 ⁻ ,4,6 ⁻ -hexachlorobiphenyl
MeSO ₂ -2,2',3,3',4',6-hexachlorobiphenyl	4 ⁻ -MeSO ₂ -CB132	4 [°] -MeSO ₂ -2,2 [°] ,3,3 [°] ,4,6 [°] -hexachlorobiphenyl
MeSO ₂ -2,2',3',4',5,5'-hexachlorobiphenyl	3'-MeSO ₂ -CB141	3 ⁻ MeSO ₂ -2,2 ⁻ ,3,4,5,5 ⁻ hexachlorobiphenyl
MeSO ₂ -2,2 [°] ,3 [°] ,4 [°] ,5,5 [°] -hexachlorobiphenyl	4'-MeSO ₂ -CB141	4 ⁻ -MeSO ₂ -2,2 ⁻ ,3,4,5,5 ⁻ -hexachlorobiphenyl
-MeSO ₂ -2,2 [°] ,4 [°] ,5,5 [°] ,6-hexachlorobiphenyl	5-MeSO ₂ -CB149	5-MeSO ₂ -2,2´,3,4´,5´,6-hexachlorobiphenyl
MeSO ₂ -2,2',3,4',5',6-hexachlorobiphenyl	4-MeSO ₂ -CB149	4-MeSO ₂ -2,2´,3,4´,5´,6-hexachlorobiphenyl
MeSO ₂ -2,2',3',4',5,5',6-heptachlorobiphenyl	5'-MeSO ₂ -CB174	5'-MeSO ₂ -2,2',3,3',4,5,6'-heptachlorobipheny
MeSO ₂ -2,2',3,3',4',5',6-heptachlorobiphenyl	4'-MeSO ₂ -CB174	4 ⁻ -MeSO ₂ -2,2 ⁻ ,3,3 ⁻ ,4,5,6 ⁻ -heptachlorobipheny
HO-2,2',4',6'-tetrachlorobiphenyl	4'-HO-CB50	4´-HO-2,2´,4,6-tetrachlorobiphenyl
HO-2,4,4,5-tetrachlorobiphenyl	5'-HO-CB66	5 ⁻ HO-2,3 ⁻ ,4,4 ⁻ tetrachlorobiphenyl
HO-3,3 [°] ,4 [°] ,5-tetrachlorobiphenyl	4'-HO-CB79	4´-HO-3,3´,4,5´-tetrachlorobiphenyl
HO-2,2´,3´,4,4´-pentachlorobiphenyl	3'-HO-CB85	3'-HO-2,2',3,4,4'-pentachlorobiphenyl
4'-diHO-2,2',3',4,5'-pentachlorobiphenyl	3´,4-diHO-CB90	3´,4-diHO-2,2´,3,4´,5-pentachlorobiphenyl
HO-2,2´,3,5,6-pentachlorobiphenyl	4-HO-CB93	4-H0-2,2 ² ,3,5,6-pentachlorobiphenyl
HO-2,2´,3,4´,5´-pentachlorobiphenyl	4-HO-CB97	4-H0-2,2´,3,4,'5´-pentachlorobiphenyl
HO-2,2',4',5,5'-pentachlorobiphenyl	4'-HO-CB101	4´-HO-2,2´,4,5,5´-pentachlorobiphenyl
HO-2,2 ² ,4 ² ,6,6 ² -pentachlorobiphenyl	4'-HO-CB104	4´-HO-2,2´,4,6,6´-pentachlorobiphenyl
HO-2,3,3´,4´,5-pentachlorobiphenyl	4-HO-CB107	4-H0-2,3,3´,4´,5-pentachlorobiphenyl
4´-diHO-2´,3,3´,4,5´-pentachlorobiphenyl	2´,4-diHO-CB107	2 [°] ,4-diHO-2,3,3 [°] ,4 [°] ,5-pentachlorobiphenyl
HO-2',3,3',4',5-pentachlorobiphenyl	4'-HO-CB108	4'-HO-2,3,3',4,5'-pentachlorobiphenyl
HO-2,3,3',5,6-pentachlorobiphenyl	4-HO-CB112	4-HO-2,3,3´,5,6-pentachlorobiphenyl
HO-2,3´,4,4´,5-pentachlorobiphenyl	3-HO-CB118	3-H0-2,3',4,4',5-pentachlorobiphenyl
-HO-2´,3,4´,5,5´-pentachlorobiphenyl	4'-HO-CB120	4´-HO-2,3´,4,5,5´-pentachlorobiphenyl
HO-2',3,4',5,6'-pentachlorobiphenyl	4'-HO-CB121	4´-HO-2,3´,4,5´,6-pentachlorobiphenyl
-HO-3,3',4',5,5'-pentachlorobiphenyl	4'-HO-CB127	4´-HO-3,3´,4,5,5´-pentachlorobiphenyl
HO-2,2´,3,3´,4´,5-hexachlorobiphenyl	4'-HO-CB130	4´-HO-2,2´,3,3´,4,5´-hexachlorobiphenyl
HO-2,2 [°] ,3,3 [°] ,5,6-hexachlorobiphenyl	4-HO-CB134	4-H0-2,2´,3,3´,5,6-hexachlorobiphenyl
HO-2,2',3',4,4',5-hexachlorobiphenyl	3'-HO-CB138	3´-HO-2,2´,3,4,4´,5´-hexachlorobiphenyl
HO-2,2´,3,4´,5,5´-hexachlorobiphenyl	4-HO-CB146	4-HO-2,2´,3,4´,5,5´-hexachlorobiphenyl
HO-2,2',4,4',5,5'-hexachlorobiphenyl	3-HO-CB153	3-HO-2,2´,4,4´,5,5´-hexachlorobiphenyl
HO-2´,3,3´,4´,5,5´-hexachlorobiphenyl	4'-HO-CB159	4´-HO-2,3,3´,4,5,5´-hexachlorobiphenyl
HO-2,3,3 [°] ,4 [°] ,5,5 [°] -hexachlorobiphenyl	4-HO-CB162	4-H0-2,3,3 [°] ,4 [°] ,5,5 [°] -hexachlorobiphenyl
HO-2,3,3 [°] ,4 [°] ,5,6-hexachlorobiphenyl	4-HO-CB163	4-HO-2,3,3',4',5,6-hexachlorobiphenyl
HO-2,3,3',5,5',6-hexachlorobiphenyl	4-HO-CB165	4-H0-2,3,3´,5,5´,6-hexachlorobiphenyl
HO-2,2´,3,3´,4´,5,5´-heptachlorobiphenyl	4'-HO-CB172	4'-HO-2,2',3,3',4,5,5'-heptachlorobiphenyl
HO-2,2',3,3',4',5,6'-heptachlorobiphenyl	4'-HO-CB175	4´-HO-2,2´,3,3´,4,5´,6-heptachlorobiphenyl
HO-2,2´,3,3´,5,5´,6-heptachlorobiphenyl	4-HO-CB178	4-HO-2,2´,3,3´,5,5´,6-heptachlorobiphenyl
HO-2,2´,3,3´,5,5´,6´-heptachlorobiphenyl	4'-HO-CB178	4'-HO-2,2',3,3',5,5',6-heptachlorobiphenyl
4 ⁻ -diHO-2,2 ⁻ ,3,3 ⁻ ,5,5 ⁻ ,6-heptachlorobiphenyl	4,4 ⁻ -diHO-CB178	4,4'-diHO-2,2',3,3',5,5',6-heptachlorobipheny
HO-2,2´,3´,4,4´,5,5´-heptachlorobiphenyl	3'-HO-CB180	3´-HO-2,2´,3,4,4´,5,5´-heptachlorobiphenyl
HO-2,2´,3´,4,4´,5´,6-heptachlorobiphenyl	3'-HO-CB182	3 [°] -HO-2,2 [°] ,3,4,4 [°] ,5,6 [°] -heptachlorobiphenyl
HO-2,2´,3´,4,4´,5,6´-heptachlorobiphenyl	3'-HO-CB183	3´-HO-2,2´,3,4,4´,5´,6-heptachlorobiphenyl
HO-2,2´,3´,4,4´,6,6´-heptachlorobiphenyl	3'-HO-CB184	3´-HO-2,2´,3,4,4´,6,6´-heptachlorobiphenyl
HO-2,2´,3´,4,5,5´,6´-heptachlorobiphenyl	3'-HO-CB187	3´-HO-2,2´,3,4´,5,5´,6-heptachlorobiphenyl
HO-2,2´,3,4´,5,5´,6-heptachlorobiphenyl	4-HO-CB187	4-H0-2,2´,3,4´,5,5´,6-heptachlorobiphenyl
4´-diHO-2,2´,3´,4,5,5´,6´-heptachlorobiphenyl	3´,4-diHO-CB187	3´,4-diHO-2,2´,3,4´,5,5´,6-heptachlorobipheny
HO-2,3,3 [°] ,4 [°] ,5,5 [°] ,6-heptachlorobiphenyl	4-HO-CB193	4-H0-2,3,3´,4´,5,5´,6-heptachlorobiphenyl
HO-2,2',3,3',4',5,5',6'-octachlorobiphenyl	4'-HO-CB198	4'-HO-2,2',3,3',4,5,5',6-octachlorobiphenyl
HO-2,2´,3,3´,4´,5,5´,6-octachlorobiphenyl	4'-HO-CB201/199d	4´-HO-2,2´,3,3´,4,5,5´,6´-octachlorobiphenyl
HO-2,2´,3,3´,4´,5,6,6´-octachlorobiphenyl	4 ⁻ H0-CB200/201 ^d	4´-HO-2,2´,3,3´,4,5´,6,6´-octachlorobiphenyl
HO-2,2´,3,3´,5,5´,6,6´-octachlorobiphenyl	4-HO-CB202	4-HO-2,2´,3,3´,5,5´,6,6´-octachlorobiphenyl
4´-diHO-2,2´,3,3´,5,5´,6,6´-octachlorobiphenyl	4,4´-diHO-CB202	4,4'-diHO-2,2',3,3',5,5',6,6'-octachlorobipheny
HO-2,2 [°] ,3 [°] ,4,4 [°] ,5,5 [°] ,6 [°] -octachlorobiphenyl	3'-HO-CB203	3´-HO-2,2´,3,4,4´,5,5´,6-octachlorobiphenyl

^aMeS0₂-PCBs (Larsson et al. 2002; Weistrand and Norén 1997); HO-PCBs (Bergman et al. 1994; Hovander et al. 2002). ^bMeS0₂-PCBs (Larsson et al. 2002); HO-PCBs (Bennett et al. 2002; Campbell et al. 2003; Hovander et al. 2002; Li et al. 2003; Sandala et al., in press; Sandau et al. 2000a; Sjödin et al. 1998). ^cHO-PCBs (Campbell et al. 2003; Sandala et al., in press; Sandau et al. 2000a; Sjödin et al. 1998). ^dOriginal BZ/revised PCB number.

REFERENCES

- Ballschmiter K, Bacher R, Mennel A, Fischer R, Riehle U, Swerev M. 1992. The determination of chlorinated biphenyls, chlorinated dibenzodioxins, and chlorinated dibenzofurans by GC-MS. J High Resolut Chromatogr 15:260–270.
- Ballschmiter K, Zell M. 1980. Analysis of polychlorinated biphenyls (PCB) by glass capillary gas chromatography. Fresenius J Anal Chem 302:20–31.
- Bennett ER, Ross PS, Letcher RJ. 2002. Polyhalogenated phenolic contaminants in Pacific killer whale (Orcinus orca). Organohalogen Compounds 58:81–84.
- Bergman Å, Klasson-Wehler E, Kuroki H. 1994. Selective retention of hydroxylated PCB metabolites in blood. Environ Health Perspect 102:464–469.
- Brouwer A, Morse DC, Lans MC, Schuur AG, Murk AJ, Klasson-Wehler E, et al. 1998. Interactions of persistent environmental organohalogens with the thyroid hormone system: mechanisms and possible consequences for animal and human health. Toxicol Ind Health 14:59–84.
- Burreau S, Broman D, Orn U. 2000. Tissue distribution of 2,2',4,4'-tetrabromo[C-14]diphenyl ether ([C-14]-PBDE 47) in pike (*Esox lucius*) after dietary exposure—a time series study using whole body autoradiography. Chemosphere 40:977–985.
- Campbell LM, Muir DCG, Whittle DM, Backus S, Norstrom RJ, Fisk AT. 2003. Hydroxylated PCBs and other chlorinated phenolic compounds in lake trout (*Salvelinus namaycush*) blood plasma from the Great Lakes region. Environ Sci Technol 37:1720–1725.
- Chu S, Covaci A, Haraguchi K, Schepens P. 2002. Optimized separation and determination of methyl sulfone metabolites of polychlorinated biphenyls (PCBs) and p,p ⁻DDE in biota samples. Analyst 127:1621–1626.
- Chu S, Covaci A, Jacobs W, Haraguchi K, Schepens P. 2003. Distribution of methyl sulfone metabolites of PCBs and *p*,*p* '-DDE in human tissues. Environ Health Perspect 111:1222–1227.
- de Voogt P, Wells DE, Reutergårdh L, Brinkman UATh. 1990. Biological activity, determination and occurence of planar, mono- and di-ortho PCBs. Int J Environ Anal Chem 40:1–46.
- Erickson MD. 1997. Analytical Chemistry of PCBs. 2nd ed. Boca Raton, FL:CRC Press/Lewis Publishers.
 - 2001. Introduction: PCB properties, uses, occurrence, and regulatory history. In: PCBs: Recent Advances in Environmental Toxicology and Health Effects (Robertson LW, Hansen LG, eds). Lexington, KY:The University Press of Kentucky, xi–xxx.
- Frame GM, Cochran JW, Bøwadt SS. 1996. Complete PCB congener distribution for 17 Aroclor mixtures determined by 3 HRGC systems optimized for comprehensive, quantitative, congener-specific analysis. J High Resolut Chromatogr 19:657–668.
- Guitart R, Puig P, Gómez-Catalán J. 1993. Requirement for a standardized nomenclature criterium for PCBs: computer assisted assignment of correct congener denomination and number. Chemosphere 27:1451–1459.
- Guvenius DM, Hassanzadeh P, Bergman Å, Norén K. 2002. Metabolites of polychlorinated biphenyls in human liver and adipose tissue. Environ Toxicol Chem 21:2264–2269.
- Haglund PS, Zook DR, Buser HR, Hu JW. 1997. Identification and quantification of polybrominated diphenyl ethers and methoxy-polybrominated diphenyl ethers in Baltic biota. Environ Sci Technol 31:3281–3287.
- Hakk H, Letcher RJ. 2003. Metabolism in the toxicokinetics and fate of brominated flame retardants (BFRs): a review. Environ Int 29:801–828.
- Hoekstra PF, Letcher RJ, O'Hara TM, Backus SM, Solomon KR, Muir DCG. 2003. Hydroxylated and methylsulfone-containing metabolites of polychlorinated biphenyls in the plasma and blubber of bowhead whales (*Balaena mysticetus*). Environ Toxicol Chem 22:2650–2658.
- Hovander L, Malmberg T, Athanasiadou M, Athanassiadis I, Rahm S, Bergman Å, et al. 2002. Identification of hydroxylated PCB metabolites and other phenolic halogenated pollutants in human blood plasma. Arch Environ Contam Toxicol 42:105–117.
- Larsson C, Ellerichmann T, Hühnerfuss H, Bergman Å. 2002. Chiral PCB methyl sulfones in rat tissues after exposure to technical PCBs. Environ Sci Technol 36:2833–2838.
- Letcher RJ, Klasson-Wehler E, Bergman Å. 2000a. Methyl sulfone and hydroxylated metabolites of polychlorinated biphenyls.

In: The Handbook of Environmental Chemistry 3K (Paasivirta J, ed). Berlin:Springer-Verlag, 315–359.

- Letcher RJ, Norstrom RJ, Bergman Å. 1995. Geographical distribution and identification of methyl sulphone PCB and DDE metabolites in pooled polar bear (Ursus maritimus) adipose tissue from Western hemisphere Arctic and subarctic regions. Sci Total Environ 160/161:409–420.
- Letcher RJ, Norstrom RJ, Muir DCG, Sandau CD, Koczanski K, Michaud R, et al. 2000b. Methylsulfone polychlorinated biphenyl and 2,2-bis(chlorophenyl)-1,1-dichloroethylene metabolites in beluga whale (*Delphinapterus leucas*) from the St. Lawrence River Estuary and Western Hudson Bay, Canada. Environ Toxicol Chem 19:1378–1388.
- Letcher RJ, van der Burg B, Brouwer A, Lemmen J, Bergman Å, van den Berg M. 2002. In vitro antiestrogenic effects of aryl methyl sulfone metabolites of polychlorinated biphenyls and 2,2-bis(4-chlorophenyl)-1,1-dichloroethene on 17β-estradiol-induced gene expression in several bioassay systems. Toxicol Sci 69:362–372.
- Li H, Drouillard KG, Bennett E, Haffner GD, Letcher RJ. 2003. Plasma-associated halogenated phenolic contaminants in benthic and pelagic fish species from the Detroit River. Environ Sci Technol 37:832–839.

- Meerts IATM, Letcher RJ, Hoving S, Marsh G, Bergman Å, Lemmen JG, et al. 2001. In vitro estrogenicity of polybrominated diphenyl ethers, hydroxylated PBDEs, and polybrominated bisphenol A compounds. Environ Health Perspect 109:399–407.
- Sandala GM, Sonne-Hansen C, Dietz R, Muir DCG, Valters K, Bennett ER, et al. In press. Methyl sulfone and hydroxylated PCB metabolites in adipose and whole blood of polar bear (*Ursus maritimus*) from East Greenland. Sci Total Environ.
- Sandau CD, Ayotte P, Dewailly E, Duffe J, Norstrom RJ. 2000a. Analysis of hydroxylated metabolites of PCBs (H0-PCBs) and other chlorinated phenolic compounds in whole blood from Canadian Inuit. Environ Health Perspect 108:611–616.
- 2002. Pentachlorophenol and hydroxylated polychlorinated biphenyl metabolites in umbilical cord plasma of neonates from coastal populations in Quebec. Environ Health Perspect 110:411–417.
- Sandau CD, Meerts IATM, Letcher RJ, McAlees AJ, Chittim B, Brouwer A, et al. 2000b. Identification of 4-hydroxyheptachlorostyrene in polar bear plasma and its binding affinity to transthyretin: a metabolite of octachlorostyrene? Environ Sci Technol 34:3871–3877.
- Schulte E, Malisch R. 1983. Berechnung der Vahren PCB-Gehalte

in Umweltproben I. Ermittlung der Zusammensetzung Zweier Technischer PCB-Gemische. Fresenius J Anal Chem 314:545–551.

- Sjödin A, Tullsten AK, Klasson-Wehler E. 1998. Identification of the parent compounds to selectively retained hydroxylated PCB metabolites in rat blood plasma. Organohalogen Compounds 37:365–368.
- Stapleton HM, Letcher RJ, Baker JE. 2001. Metabolism of PCBs by the deepwater sculpin (*Myoxocephalus thompsoni*). Environ Sci Technol 35:4747–4752.
- U.S. Environmental Protection Agency. 2003. PCB ID—Table of PCB Congeners and Other Species. Washington, DC:Environmental Protection Agency. Available: http:// www.epa.gov/toxteam/pcbid/table.htm [accessed 23 January 2004].
- Valters K, Alaee M, Marsh G, D'Sa I, Li H, Bennett ER, et al. 2003. Polybrominated diphenyl ethers and hydroxylated and methoxylated analogues in Detroit River fish. Organohalogen Compounds 61:29–32.
- Weistrand C, Norén K. 1997. Methylsulfonyl metabolites of PCBs and DDE in human tissues. Environ Health Perspect 105:644–649.