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3 **Weight and height at 4 and 7 years of age in children born to**
4 **mothers with a high intake of fish contaminated with**
5 **persistent organochlorine pollutants**

6

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24 **Abstract**

25 In Sweden the main exposure route for persistent organochlorine pollutants (POP) is
26 through consumption of fatty fish from the Baltic Sea (off the eastern coast).
27 The present study aimed to investigate whether intrauterine exposure for POP may have
28 negative impact on children's weight and height at 4 and 7 years of age, respectively.
29 The study included 174 fishermen's wives from the Swedish east coast who had given
30 birth to an infant with either low (n=55) or normal (n=119) birth weight, and 88 and 206
31 corresponding women from the Swedish west coast (where the fish is less polluted).
32 Comparisons between the east and west coast cohorts were performed. In addition,
33 blood samples were collected among the east coast women and the concentrations of
34 2,2',4,4',5,5'-hexachlorobiphenyl (CB-153) in plasma was analyzed and estimated for
35 the year of childbirth. There were no significant differences between the east and west
36 coast cohorts regarding weight and height at 4 and 7 years of age. There were, however,
37 significant negative associations between the estimated plasma concentrations of CB-
38 153 during year of childbirth and weight at 4 and 7 years of age, respectively, among
39 the normal birth weight children. The study gives only very weak support for the
40 hypothesized association.

41

42

43

44 **Key words:** Growth, polychlorinated biphenyls, dioxin, fish

45 **1. Background**

46 Although human exposure to persistent organochlorine pollutants (POP) such as
47 polychlorinated biphenyls (PCB) and dioxins has decreased during recent decades in
48 some regions (Odsjö et al., 1997), low level exposure to these compounds are probably
49 still important from a health perspective. In the general population in the Netherlands,
50 in utero exposure to POP was negatively associated with birth weight and postnatal
51 growth until 3 months of age (Patandin et al., 1998). However, no negative effects of
52 prenatal POP exposure were found on growth rate from 3 to 42 month of age. On the
53 other hand, such long term effects have been observed in studies from the Great Lakes
54 in the US where negative associations between intrauterine PCB exposure through fish
55 consumption and birth weight as well as growth until 4 years of age were seen (Fein et
56 al., 1984; Jacobson et al., 1990). In another study, prenatal PCB exposure was
57 negatively associated with growth among girls (Blanck et al., 2002). There are,
58 however, also epidemiological studies showing no negative effect on growth and even a
59 positive effect on growth after POP exposure (Rogan et al., 1987; Gladen et al., 2000;
60 Hertz-Picciotto et al., 2005).

61

62 Negative effects of prenatal POP exposure has in animal studies been associated with
63 reduced birth weight (Allen et al., 1980; Overman et al., 1987; Brezner et al., 1984) as
64 well as with slower growth later on. In one study prenatal PCB exposed rats gained
65 weight more slowly than controls during the first four month of life (Brezner et al.,
66 1984).

67

68 In Sweden the main exposure route for POP is through consumption of fatty fish from
69 the Baltic Sea, off the eastern coast of Sweden (Asplund et al., 1994; Svensson et al.,

70 1991, 1995). Fishermen's wives from the Swedish east and west coasts have reported
71 that they consume more than twice as much fish as compared with women from the
72 general population (Rylander et al., 1995). During the period 1973-1991 infants born to
73 fishermen's wives from the Swedish east coast had an increased risk for lower birth
74 weight as compared with a corresponding group from the Swedish west coast (Rylander
75 and Hagmar, 1995), where the fish has been much less contaminated (Bergqvist et al.,
76 1989). Case-control studies among the infants born to fishermen's wives from the
77 Swedish east coast indicated an increased risk of lower birth weight among infants born
78 to mothers who reported a relatively high current intake of fish from the Baltic Sea, as
79 well as among infants born to mothers with a relatively high concentration of
80 2,2',4,4',5,5'-hexachlorobiphenyl (CB-153) in plasma (Rylander et al., 1996,1998). CB-
81 153 was selected as a biomarker for POP exposure due to its very high correlations with
82 the total PCB concentration in plasma and serum (Grimvall et al., 1997; Glynn et al.,
83 2000), and the total POP derived 2,3,7,8-tetrachlorodibenzo-p-dioxin equivalent in
84 plasma (Gladen et al., 1999). In addition, the PCB contribution to "dioxin-like"
85 exposure among high consumers of fish from the Baltic Sea had been estimated to be
86 almost 80%, whereas that from polychlorinated dioxins and furans contributed with
87 about 20% (Asplund et al., 1994).

88

89 The aim of the present study was to investigate whether intrauterine exposure for POP
90 may have negative impact on children's weight and height at 4 and 7 years of age,
91 respectively.

92 **2. Material and methods**

93 *2.1 Study base*

94 Cohorts of fishermen's wives from the Swedish east and west coasts have previously
95 been established (Rylander and Hagmar., 1995). These women were linked to the
96 Swedish Medical Birth Register, which includes almost every infant born in Sweden
97 since 1973 (Cnattingius et al., 1990). During the period 1973-1991, 757 women in the
98 east coast cohort gave birth to 1501 children and 1834 women in the west coast cohort
99 gave birth to 3553 children (Table 1)(Rylander et al., 1995). In the original cohort study
100 we used 2500 g as well as 3000 g as the cut points for LBW (Rylander et al. 1995). In
101 the following case-control study within the east coast cohort we had the intention to
102 contrast the cases from the controls, and accordingly infants with a birth weight in the
103 span 2750 g and 3250 g were excluded.

104

105 *2.2 Selected normal and low birth weight children*

106 In the cohorts, 89 east and 149 west coast mothers had given birth to an infant who
107 fulfilled the following criteria: singleton, birth weight within the interval 1500-2750 g
108 (in the present study defined as low birth weight, LBW), and without major
109 malformation. If a mother had given birth to more than one infant with LBW, only the
110 first infant was selected.

111

112 For each selected child with LBW two children from the cohorts were randomly
113 selected. These children fulfilled the following criteria: singleton, birth weight within
114 the interval 3250-4500 g (in the present study defined as normal birth weight, NBW),
115 and without major malformation. In addition, they were matched to the LBW child
116 according to gender, parity (1, 2, or ≥ 3), and calendar year of birth (± 5 years).

117

118 Accordingly, the results of this selection process were four groups: 1) 89 children from
119 the east coast cohort with LBW, 2) 149 children from the west coast cohort with LBW,
120 3) 178 children from the east coast cohort with NBW, and 4) 298 children from the west
121 coast cohort with NBW (Table 1). In the study one woman could only contribute with
122 one infant.

123

124 The selected east coast children were the same as in the ones in the former case-control
125 study that investigated the hypothesized association between POP exposure and low
126 birth weight (Rylander et al., 1996, 1998).

127

128 *2.3 Data on weight and height and potential confounders*

129 Information about the children's weight and height in the four groups at about 4 and 7
130 years of age (exact ages for the measurements were always obtained) was collected in
131 two ways. First, the mothers were contacted by telephone and asked if they could
132 provide this information. Second, child health centers (CHS) and school health services
133 (SHS) were contacted and asked if they could provide the requested information.

134 Informed consents were obtained from the mothers or the children (if they were at least
135 15 years of age) before such contacts were made. In Sweden, CHS are responsible for
136 the children's health until the year when the children start the primary school (the year
137 when the child will be 7 years of age). After that the SHS has the health responsibility.

138 When information was received from both sources, *i.e.* the mothers report and

139 CHS/SHS, the data from the CHS/SHS was considered more trustworthy and was

140 therefore primarily used in the statistical analyses. At 4 years of age, CHS data was

141 obtained for about 75% of the participating west coast children and for slightly less than

142 60% of the east coast children, but there were very high correlations with data obtained
143 from the mothers (Pearsons correlation coefficients (r) 0.95 for weight and 0.89 for
144 height). In addition, there were no systematic differences between the two sources. At 7
145 years of age, SHS data were obtained for more than 90% of the children in both the east
146 and west coast cohorts. At this age there were, however, lower correlations between the
147 SHS data given by the mothers (0.82 for weight and 0.70 for height). In addition, the
148 weight and height data obtained from the SHS were systematically somewhat higher,
149 due to that the SHS measurements normally were obtained during autumn semester of
150 the school-year, whereas the data given by the mothers were closer to the child's 7 years
151 birthday.

152

153 At the telephone interview the women were also asked about smoking habits, education,
154 and their own height. One person conducted all the telephone interviews. Fifty-five east
155 coast LBW mother-child pairs participated. The corresponding figure for west coast
156 LBW was 88, for east coast NBW 119, and for west coast NBW 206 (Table 1).

157 Background characteristics of the participants are shown in Table 2. The study was
158 approved by the Ethic's Committee of Lund University.

159

160 *2.4 Non-participants*

161 The age distributions were very similar among participants and non-participants.
162 Among the east coast women the median birth year of the mothers was 1955 (range
163 1938, 1972) among the participating women and 1954 (1933, 1967) among the non-
164 participants. The corresponding figures among the west coast women were 1952 (1930,
165 1971) and 1951 (1929, 1965), respectively. Other characteristics than birth year were
166 unfortunately not available for the non-participants.

167

168 *2.5 Exposure assessments*

169 The cohort affiliation (east and west coast) was treated as a proxy for POP exposure.

170 Due to the decreased levels over time of PCB in fish from the Baltic Sea (Odesjö et al.,

171 1997), the results are also presented separately for two periods (children born 1973-

172 1980 and 1981-1991, respectively).

173

174 For 157 (48 LBW and 109 NBW) out of the 174 (55 LBW and 119 NBW) participating

175 east coast mothers blood samples were drawn in 1995, and the concentration of

176 2,2',4,4',5,5'-hexachlorobiphenyl (CB-153) in plasma was analyzed using gas

177 chromatography with an electron capture detector (GC-ECD). The methodology for

178 plasma extraction, clean-up, identification and quantification is given in detail

179 elsewhere (Grimvall et al., 1997). The total lipid concentration in plasma was calculated

180 by summation of the amounts of triglycerides, cholesterol, and phospholipids using

181 enzymatic methods (Grimvall et al., 1997). Again, the infants had been born during the

182 period 1973-1991, and the plasma levels of CB-153 in 1995 were probably not fully

183 relevant for the intrauterine exposure. We did, therefore, estimate the concentrations of

184 CB-153 in the year when the children were born with a back-calculation model where

185 the impact of reduction of body burden at lactation, biologic half-lives during non-

186 lactating periods, and the decrease of PCB contamination in fish over the calendar years

187 was taken into account (Rylander et al., 1998). We made the following assumptions: 3%

188 yearly reduction of CB-153 in the fish, 33% reduction in body burden of CB-153 at

189 each period of lactation, and 5 years biological half-life for CB-153 during non-

190 lactating periods. The model and a detailed discussion about the chosen assumptions

191 has been described elsewhere (Rylander et al., 1998).

192

193 *2.6 Statistics*

194 Four outcome variables were considered: weight and height at 4 and 7 years of age,

195 respectively. For comparisons between children with LBW from the east and west coast

196 cohorts linear regression models were employed. We confirmed that the regression

197 model assumption was fulfilled by residual analysis. Maternal height (three categories:

198 ≤ 159 , 160-169, and ≥ 170 cm) and exact age (months) of the children at the time of

199 measurement were both strongly associated with the outcome variables and were

200 therefore always included in the multivariate models. In addition, education (9-year

201 compulsory school, senior high school, and university), smoking habits during

202 pregnancy (non-smokers and smokers), maternal age (≤ 24 , 25-29 and ≥ 30 years),203 gender and parity (1, 2, ≥ 3) were considered as potential confounders. These variables

204 were included in the multivariate models, one at a time, and did persist in the

205 multivariate models if they changed the point estimate with at least 15 %. In addition, to

206 investigate possible effect modification we also performed gender specific analyses.

207 Corresponding comparisons were performed between children with NBW from the east

208 and west coast cohorts. Moreover, separate analyses were made within the east coast

209 cohort for the other exposure variable, *i.e.* the mother's estimated concentration of lipid

210 adjusted CB-153 in the year when the child was born. The CB-153 variable was

211 analysed as a continuous variable as well as dichotomized (at the median concentration

212 [250 ng/g lipid] and at the upper quartil 350 ng/g lipid, respectively). Due to the low

213 number of subjects, especially for LBW (varied between 38-46 subjects), we did in the

214 multivariate models only consider maternal height and exact age of the children at the

215 time of measurement as potential confounders.

216

217 3. Results

218 Regarding weight and height at 4 and 7 years of age, respectively, there were no
219 significant differences between children from the east and west coast cohorts (Tables 3
220 and 4). However, after adjustment for the confounders weight and height at 7 years of
221 age was nearly significant lower for NBW children from the east coast born in 1973-
222 1980, as compared with NBW children from the west coast. The results for 4 years of
223 age were in same direction.

224

225 When the estimated plasma concentrations of CB-153 during year of childbirth among
226 the east coast cohort women were used as exposure biomarker, no significant
227 associations on the children's growth were observed among the LBW children.

228 However, when concentrations of CB-153 was analyzed as a continuous variable there
229 were, among the NBW children, significant associations with weight at 4 and 7 years,
230 respectively. An increase of 100 ng/g lipid of CB-153 in plasma corresponded to a
231 decrease in weight at 4 years of age of 0.4 kg (95% CI 0.01-0.7, $p=0.04$). The
232 corresponded figure at 7 years of age was a decrease of 1.2 kg (95% CI 0.5-1.9,
233 $p=0.001$). Similar associations were seen when concentrations of CB-153 were
234 dichotomized. The NBW children whose mothers had CB-153 concentrations above
235 250 ng/g lipid had, however, significantly lower weight at 7 years of age than the NBW
236 children whose mothers had lower concentrations of CB-153 (Table 5). The pattern was
237 very similar when 350 ng/g lipid was used as cut-off point (data not shown).

238

239 Gender-specific analyses did not result in any obvious difference as compared to when
240 the whole data set was analyzed (data not shown).

241 **4. Discussion**

242 The results from the present study did not give any strong support for the hypothesis of
243 a negative effect of intrauterine exposure for POP on children's growth up to 4 and 7
244 years, respectively. However, although the associations were non-significant, it is
245 noteworthy that seven out of the eight adjusted estimates for the comparisons between
246 the cohorts for the early period were negative, *i.e.* showed impaired growth among the
247 east coast children. These tendencies were more obvious among the NBW children than
248 among the LBW children. It is important to be aware of that the outcome measurements
249 at 4 and 7 years of age were highly correlated (r for height 0.69 and r for weight 0.74)
250 and, accordingly, this would most probably lead to that the results for the outcome
251 measures at 4 and 7 years of age will go in the same direction. In addition, the outcome
252 measurements at 4 years ($r=0.74$ between height and weight) and 7 years ($r=0.69$) were
253 also highly correlated.

254

255 Due to the relatively high proportion of non-participants in the present study, possible
256 selection bias has to be considered. The participation rates in the two cohorts were,
257 however, very similar, and the age distributions among participating and non-
258 participating mothers were also very similar. Moreover, in former studies of women
259 from the cohort of east coast fishermen's wives, where the response rate was only
260 slightly higher, selection bias was not considered to be an issue of major concern, due to
261 similar distributions of age, educational level and smoking habits among participants
262 and non-participants (Rylander et al., 1996, 1998).

263

264 The information about the weight and height of the children were collected in two ways.
265 The somewhat lower correlation between data sources at 7 years of age and the slight

266 systematic underestimation of the outcome measures by the mothers could be explained
267 by that the data given by the mothers were closer to the child's 7 years birthday than the
268 SHS data during the autumn school semester. On the other hand, this was a minor
269 problem as SHS data was available for more than 90% of the children in both cohorts.
270 Anyhow, this validation stresses the importance to adjust for the child's exact age in the
271 statistical analyses.

272

273 We did, simultaneously with the children's exact age, also include the height of the
274 mothers in the multivariate models. Unfortunately, data on the height of the fathers were
275 only collected for the west coast cohort. The interviews with the west coast women was
276 performed after the interviews with the east coast women and this question was the only
277 one added. When separate analyses were performed within the west coast cohort, the
278 height of the father was associated only with the height of the male children. At 7 years
279 of age were, however, the height of the mothers was of greater importance than the
280 height of the fathers. In an ongoing study, where 2436 west coast fishermen and 1082
281 east coast fishermen have answered a questionnaire, the median height in the two
282 groups differed only by one cm (west coast 179 cm, east coast 178 cm, unpublished
283 data). Accordingly, the lack of information about the fathers height in the east coast
284 cohort did probably not confound the comparisons between the cohorts. It was not
285 possible to link the questionnaire based data to the women in the present study.

286

287 In the present study we used cohort affiliation as a proxy measure of exposure. It is
288 well-known that fishermen's families have a higher intake of locally caught fish as
289 compared with individuals from the general population (Svensson et al., 1995; Rylander
290 and Hagmar, 1995). The fatty fish from the Baltic Sea contains higher levels of POP as

291 compared with the corresponding fish from the Swedish west coast (Bergqvist et al.,
292 1989). This was clearly reflected in plasma collected from fishermen from the Swedish
293 east and west coasts, with a higher concentrations of dioxin-like POP among the east
294 coast fishermen (290 pg/g lipid) as compared with the west coast fishermen (139 pg/g
295 lipid) (Svensson et al., 1995). This taken together with the socio-economic similarities
296 between the cohorts make the cohort affiliation an appropriate proxy variable for POP
297 exposure.

298

299 We do believe that the estimated CB-153 concentrations in plasma during year of
300 childbirth is more relevant as compared with the CB-153 concentrations in 1995 (the
301 year when the samples were collected). The back-calculation model we used had
302 previously been validated (Rylander et al., 1998). In certain geographic regions in
303 Sweden, blood sera have been collected at antenatal clinics in a rubella screening
304 program. The CB-153 concentrations in 1995 and the CB-153 concentrations estimated
305 during year of rubella screening, respectively, were compared with the measured CB-
306 153 concentrations during year of rubella screening. The use of the back-calculation
307 model did clearly improve the agreement. In addition, the back-calculation model was
308 recently evaluated by Karmaus and colleagues who recommended the use of a complex
309 decay model following our strategies when repeated measurements are not available
310 (Karmaus et al., 2004). In the present study there was a very good correlation between
311 the CB-153 concentrations in 1995 and the estimated concentrations during year of
312 childbirth ($r_s=0.94$), but the concentrations during year of childbirth were generally
313 higher than in samples drawn in 1995.

314

315 Boys born in 1973-1975 to fishermen's wives and fishermen's sisters from the Swedish
316 east and west coast have in a previous study been examined at 18 years of age
317 (Rylander et al., 2000). Although the height significantly differed between the cohorts
318 (mean height: west coast 180.1 cm and east coast 178.8 cm) the absolute difference
319 were judged to be of little consequence. In the Lake Michigan studies, where the
320 maternal exposure levels have been estimated to be similar to the exposure levels
321 among fishermen's wives from the Swedish east coast (Grimvall et al., 1997), the
322 children's growth were negatively affected until 4 years of age (Jacobson et al., 1990).
323 There are, however, also studies which showed no negative long-term growth effects
324 (Rogan et al., 1987; Patandin et al., 1998; Gladen et al., 2000; Hertz-Picciotto et al.,
325 2005). Thus, including the present study, the epidemiological studies performed so far,
326 do not give any clear evidence that dietary POP exposure may result in long term
327 growth retardation.
328

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337

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419 *Work. Environ. Health.* 21, 96-105.

420 **Table 1.** Number of children in the original cohort and in the present study.
 421

	West coast		East coast	
	N	%	N	%
The original cohorts 1973-1991				
Number of children born	3553		1501	
Women given birth	1834		757	
Selected children for the present study				
LBW ^a	149		89	
NBW ^b	298		178	
Participants in the present study ^c				
LBW ^a	88	59	55	62
NBW ^b	206	69	119	67

422 ^a Children with a birth weight between 1500 and 2750 g,
 423 defined as low birth weight (LBW).

424 ^b Children with a birth weight between 3250 and 4500 g,
 425 defined as normal birth weight (NBW).

426 ^c Children with information of height or weight at 4 or 7 years of age.

427

428 **Table 2.** Background and exposure characteristics of the maternal and child participants in the present study.
429

	West coast				East coast			
	LBW (n=88) ^a		NBW (n=206) ^b		LBW (n=55) ^a		NBW (n=119) ^b	
	%	Median (5, 95 perc)	%	Median (5, 95 perc)	%	Median (5, 95 perc)	%	Median (5, 95 perc)
Maternal								
Age (yr) ^c								
≤ 24	23		28		40		21	
25-29	43		41		24		45	
≥ 30	34		32		36		34	
First parity	52		53		51		45	
Education								
Compulsory school	43		32		33		38	
Senior high school	41		46		45		45	
University	16		22		22		16	
Smoking ^c	53		33		58		50	
Height (cm)		165 (154, 174)		167 (158, 175)		165 (158, 175)		165 (158, 175)
CB-153 (ng/g lipid) ^e								
In 1995		-		-		178 (59, 464)		159 (69, 500)
Estimated for year of childbirth ^f		-		-		295 (102, 599)		243 (97, 719)
Child								
Calendar year of birth								
1973-1980	59		56		45		44	
1981-1991	41		44		55		56	
Male gender	39		42		53		43	

Birth weight (kg)	2.43 (1.55, 2.70)	3.73 (3.29, 4.34)	2.44 (1.74, 2.73)	3.65 (3.30, 4.36)
Exact age at the 4 year examination (months)	48.2 (47.7, 53.0)	48.1 (47.6, 53.1)	48.8 (46.4, 55.1)	48.2 (46.6, 55.3)
Exact age at the 7 year examination (months)	86.7 (73.2, 91.4)	85.7 (73.9, 93.0)	87.2 (80.0, 92.2)	87.1 (78.2, 92.4)

430 ^a Children with a birth weight between 1500 and 2750 g, defined as low birth weight (LBW).

431 ^b Children with a birth weight between 3250 and 4500 g, defined as normal birth weight (NBW).

432 ^c At year of childbirth.

433 ^d Lactation for the index pregnancy.

434 ^e CB-153 is 2,2',4,4',5,5'-hexachlorobiphenyl.

435 ^f See Materials and Metho

436 **Table 3.** The effect of cohort affiliation (east versus west coast) on weight (at 4 and 7 years of age, respectively) among children with a birth
 437 weight between 1500 and 2750 g (defined as LBW). Corresponding effect among children with a birth weight between 3250 and 4500 g (defined
 438 as NBW). Unadjusted, as well as adjusted estimates with 95% confidence intervals (CI) obtained from linear regression models are shown.
 439 Moreover, the results are divided in two calendar year periods.

440

	Weight at 4 years of age (kg)				Weight at 7 years of age (kg)			
	Unadjusted		Adjusted ^b		Unadjusted		Adjusted ^b	
	β^a	95% CI	β^a	95% CI	β^a	95% CI	β^a	95% CI
<i>LBW</i>								
born 1973-1991	0.41	-0.34, 1.16	0.17	-0.56, 0.90	1.07	-0.59, 2.73	0.06	-1.54, 1.66
- Born 1973-1980	-0.13	-1.11, 0.84	-0.12	-1.07, 0.84	1.05	-1.10, 3.21	0.13	-1.86, 2.12
- Born 1981-1991	0.62	-0.53, 1.76	0.20	-0.99, 1.39	0.76	-1.90, 3.41	-0.14	-2.83, 2.54
<i>NBW</i>								
born 1973-1991	-0.27	-0.84, 0.31	-0.27	-0.84, 0.31	0.13	-0.83, 1.10	0.17	-0.79, 1.13
- Born 1973-1980	-0.50	-1.47, 0.46	-0.62	-1.60, 0.36	-1.21	-2.49, 0.07	-1.28	-2.58, 0.01
- Born 1981-1991	-0.12	-0.86, 0.62	-0.06	-0.80, 0.69	1.06	-0.37, 2.50	1.18	-0.22, 2.58

441

^a Mean differences between east and west coast cohort children.

442

^b Adjusted for the children's exact age (in month) and height of the mother (<160, 160-169, and \geq 170 cm).

443

444 **Table 4.** The effect of cohort affiliation (east versus west coast) on height (at 4 and 7 years of age, respectively) among children with a birth
 445 weight between 1500 and 2750 g (defined as LBW), and corresponding effect among children with a birth weight between 3250 and 4500 g
 446 (defined as NBW). Unadjusted, as well as adjusted estimates with 95% confidence intervals (CI) obtained from linear regression models are
 447 shown. Moreover, the results are divided in two calendar year periods.

448

	Height at 4 years of age (cm)				Height at 7 years of age (cm)			
	Unadjusted		Adjusted ^b		Unadjusted		Adjusted ^b	
	β^a	95% CI	β^a	95% CI	β^a	95% CI	β^a	95% CI
<i>LBW</i>								
born 1973-1991	0.84	-0.74, 2.41	0.09	-1.43, 1.61	2.22	-0.19, 4.63	0.26	-1.80, 2.31
- Born 1973-1980	-0.32	-2.64, 2.00	-0.50	-2.73, 1.74	0.73	-2.64, 4.10	-1.33	-3.61, 0.96
- Born 1981-1991	1.30	-0.93, 3.53	-0.03	-2.32, 2.26	2.73	-0.75, 6.20	1.51	-1.98, 4.99
<i>NBW</i>								
born 1973-1991	-0.40	-1.46, 0.66	-0.55	-1.53, 0.43	-0.10	-1.33, 1.12	-0.33	-1.42, 0.76
- Born 1973-1980	-0.66	-2.49, 1.16	-1.46	-3.13, 0.22	-1.01	-2.76, 0.73	-1.46	-3.06, 0.13
- Born 1981-1991	-0.13	-1.49, 1.22	0.09	-1.17, 1.35	0.72	-1.05, 2.49	0.64	-0.88, 2.17

449

^a Mean differences between east and west coast cohort children.

450

^b Adjusted for the children's exact age (in month) and height of the mother (<160, 160-169, and \geq 170 cm).

451

452 **Table 5.** The effect of maternal concentration of lipid-adjusted CB-153 (2,2',4,4',5,5'-hexachlorobiphenyl) in plasma in the year when the child
 453 was born on growth (height and weight at 4 and 7 years of age, respectively) among children born to fishermen's wives from the Swedish east
 454 coast. Estimates with 95% confidence intervals (CI) obtained from linear regression models are shown. Moreover, the results are divided in
 455 children with low birth weight (LBW) and children with normal birth weight (NBW).

456

	Growth at 4 years of age				Growth at 7 years of age			
	Weight (kg)		Height (cm)		Weight (kg)		Height (cm)	
	β^a	95% CI	β^a	95% CI	β^a	95% CI	β^a	95% CI
<i>LBW</i>								
Crude ^b	-1.34	-2.74, 0.05	-1.16	-4.08, 1.77	0.26	-2.50, 3.02	-2.50	-6.97, 1.96
Adjusted ^c	-1.05	-2.51, 0.42	-0.21	-2.91, 2.49	0.73	-1.93, 3.39	-1.69	-6.06, 2.69
<i>NBW</i>								
Crude ^b	-0.12	-1.02, 0.79	0.92	-0.75, 2.60	-1.67	-3.37, 0.02	0.14	-1.78, 2.06
Adjusted ^c	-0.39	-1.21, 0.44	0.35	-1.08, 1.79	-2.00	-3.63, -0.37	-0.31	-2.05, 1.43

457 ^a Mean differences between children whose mothers had plasma concentrations of CB-153 greater than 250 ng/g lipid compared
 458 with those whose mothers had lower concentrations.

459 ^b Univariate models

460 ^c Adjusted for the children's exact age (in month) and height of the mother (<160, 160-169, and \geq 170 cm).

461

462