

Table 2.7.1: Cohort studies on relationship of endocrine disruptors with thyroid functions

| Region and subjects | Number of subjects | Follow-up period | Compound | Measure of effects | Findings | P trend |
|--|---------------------|-------------------|----------|---|---|-------------------------|
| Gocmen, 1989 | | | | | | |
| Southern Anatolia, Turkey | 252 | About 20-30 years | HCB | Frequency of symptom (%) (with symptom/total) | | No comparison attempted |
| About 4000 people exposed (by intake) to HCB in 1955-59. Those who have history of porphyria in the population were investigated in 1977-87. | (162 men, 90 women) | | | Enlarged thyroid gland | 3.49 (88/252) | |
| Average 35.7 years of age, onset at an average 7.6 year of age. | | | | Others | | |
| Retrospective study | | | | | | |
| | | | | | Scars in face or hand 83.7 (211/252) | |
| | | | | | Excessive pigment deposition 65 (164/252) | |
| | | | | | Hypertrichosis 60.6 (131/216) | |
| | | | | | Myotonia 37.9 (82/216) | |
| | | | | | Cogwheeling 41.9 (70/167) | |
| | | | | | Enlarged liver 4.8 (12/252) | |

Table 2.7.2: Case-control study on relationship of endocrine disruptors with thyroid functions

| Region and number of subjects (case/control) | Compound | | | P value |
|--|-------------------|-------------------|---------|---------|
| | | Case | Control | |
| Sukdolova, 2000 | | Serum level (ppb) | | |
| U.S. | 総 PCB | 4.8 | 4.7 | 有意差なし |
| Mohawk women over 30 years of age | PCB8 | 0.015 | 0.03 | <0.05 |
| Hypothyroidism 46, control 75 | PCB31 | 0.025 | 0.045 | <0.05 |
| | PCB28 | 0.04 | 0.08 | <0.05 |
| | PCB49 | 0.015 | 0.035 | <0.05 |
| | PCB47+59 | 0.05 | 0.035 | <0.05 |
| | PCB71 | 0.005 | 0.025 | <0.05 |
| | PCB70 | 0.02 | 0.03 | <0.05 |
| | PCB84 | 0.02 | 0.02 | <0.05 |
| | PCB90+101 | 0.065 | 0.085 | <0.05 |
| | PCB156 | Case > control | | ? |
| | PCB118 | Case > control | | ? |
| | PCB153 | Case = control | | ? |
| | PCB28 | Case < control | | ? |
| | (For information) | | | |
| | Thyroid TEQ (2) | | | |
| | PCB156; 0.03 | | | |
| | PCB118; 0.02 | | | |
| | PCB153; 0.001 | | | |
| | PCB28; 0.0006 | | | |

1) Read from figure

2) Calculated from data in other reports

Table 2.7.3: Synchronic studies on relationship of endocrine disruptors with thyroid functions

| Region and subjects | Compound | Measure of effects | Findings | P trend |
|---|----------------|---|---|--|
| Bahn, 1980 | | | | |
| 35 Factory workers handling PBB 89 Controls | PBB | Hypothyroidism | 4 Exposed/0 control | |
| Murai, 1987 | | | | |
| | PCB | T3 T4 TSH Thyroid adenoma | Significantly higher for cases Significantly higher for cases No difference from controls Serum TCB level not correlated with T3, T4, TSH 11/74 (15%) for women | |
| Emmet, 1988 | | | | |
| U.S. Transformer repair workers 55 Exposed to PCBs (38 currently exposed, 17 exposed in past) 56 Workers without PCB exposure history as controls | | T4 Free T4 index | 8.24 μg/dl for exposed/8.84 μg/dl for control 226 exposed/248 controls | Significant Significant |
| Koopman-Esseboom, 1994 | | | | |
| Holland 105 Mother-newborn pairs | PCB dioxins | Thyroid hormones (TT4, TT3, FT4, TSH) | Mothers whose milk has high PCB levels show low T3 and T4, while the pairing children show high TSH levels 2 weeks after birth. | |
| Tsuji, 1997 | | | | |
| Japan 81 Recognized yusho patients studied 28 years after onset | PCB | Thyroid disorders Disorder found in thyroid check only T4, T3, TSH Anti-thyroglobulin antibody positive Anti-microsome antibody | 3 Graves' disease 2 Chronic thyroiditis 2 Thyroid carcinoma 1 Thyroid tumor 6 Cases (TSH slightly increased in 4, slightly decreased in 2) No difference from controls More frequent in high serum PCB group More frequent in high serum PCB group | |
| Langer, 1998 (The report in 1996 is also similar) | | | | |
| Slovakia Workers in a PCB manufacturing plant and residents around it 238 Plant employees, 572 controls from low pollution area 1419 people 17 years of age (454 from highly polluted urban area and 965 controls) for evaluation of thyroid gland volume | PCB | Thyroid gland volume (ml) Thyroid antibody (%) (PCB plant employees/controls from low pollution area) TPO Ab (all subjects) 26.0/20.7 TPO Ab (women) 28.4/20.5 Tg Ab (women 30-61 years of age) 21.3/14.6 TSHR Ab (238 matched) 10.5/2.5 T4 level (nmol/l) (PCB plant employees/controls from low pollution area) | PCB manufacturing plant workers: 17.3, controls from low-pollution areas: 11.3 17-Year-old residents in polluted cities: 9.0, control youths: 7.7 116.1 / 112.2 | P<0.001 P<0.001 P>0.05 P<0.05 P<0.05 P<0.001 No significant difference |

Table 2.7.3: Synchronic studies on relationship of endocrine disruptors with thyroid functions

| Region and subjects | Compound | Measure of effects | Findings | P trend |
|---|--------------------|---------------------------------------|--|---------------------------|
| | | Frequency of normal thyroid gland (%) | | |
| | | | Employed for 21-35 years: 33.6, employed for 11-20 years: 49.3 Employed for 21-35 years: 33.6, matched controls 46.1 | P<0.05 P<0.025 |
| Mazhitova, 1998 | | | | |
| Kazakhstan | PCB | Thyroid hormone | | No significant difference |
| Region near Aral Sea | | TSH | | No significant difference |
| 12 Hospitalized children, 7.5-15 years of age | | | | |
| Schoolchildren in Stockholm as controls | | | | |
| Nagayama, 1998 | | | | |
| Japan | PCDD, PCDF, Co-PCB | Thyroid functions | | |
| 36 Infants 1 year of age | | | | |
| Sala, 1999 | | | | |
| Catalonia, Spain | HCB | Hypothyroidism | | |
| Areas with high atmospheric HCB levels | | | Women: With employment history 1/62, never employed 17/952 Men: With employment history 1/445, never employed 0/341 | |
| 1800 Residents of villages around an electrochemical plant including employees of the plant | | Others | | |
| Average serum HCB level (ng/ml) of 608 residents: | | | Natural miscarriage: With employment history 8/60, never employed 143/396 Low body weight of newborn: With employment history 2/46, never employed 51/719 Congenital malformation: With employment history 1/46, never employed 26/719 | |
| 54.6 for current male employees | | | | |
| 27.1 for male ex-employees | | | | |
| 9.0 for men without employment history | | | | |
| 14.9 for current female employees | | | | |
| 22.2 for female ex-employees | | | | |
| 13.5 for women without employment history | | | | |
| Guo, 1999 | | | | |
| Taiwan | | Thyroid adenoma | High frequency (20%) | |
| Yusho patients studied 13 years after onset | | | | |

Table 2.7.4: Ecological study on relationship of endocrine disruptors with thyroid functions

| | | | | P trend |
|--|-----------------------------------|--------------------|---|---------|
| Region and subjects | Compound | Measure of effects | Findings | |
| Sandau et al. 2002 | | | | |
| Quebec, Canada Cord blood samples obtained in 3 regions in 1993-96 Highly PCB-exposed Inuit people in Nunavik, subsistence fishermen around Lower North Shore of the Gulf of St. Laurence, and a southern Quebec urban center where PCB exposure was at background level | PBB PCP 4-HO-HpCS OH-PCB | FT4 | Principal agent was PCP, no regional difference in concentration (628-7680 pg/g wet weight) OH-PCB concentrations: 553 (238-1750) pg/g wet weight in Lower North Shore, 286 (103-788) pg/g wet weight in Nunavik, 234 (147-464) pg/g wet weight in Southern Quebec Average of total plasma PCB concentration (sum of 49 congeners): 2710 (525-7720) ng/g wet weight in Lower North Shore, 1510 pg/g wet weight in Nunavik, 843 pg/g wet weight in Southern Quebec Total HO-PCB significantly correlated with total PCB (logarithmic transformation) ($r = 0.69$, $p < 0.001$) FT4 level (logarithmic transformation) inversely correlated with total chlorinated phenols in Nunavik and Lower North Shore groups | |

Table 2.7.5: Intervention study on relationship of endocrine disruptors with thyroid functions

| | | | | P trend |
|---|-----------------|-------------------------------|--|---------|
| Region and subjects | Compound | Measure of effects | Findings | |
| Pelletier et al. 2002 | | | | |
| Quebec, Canada | b-HCH | T3 | Significant decrease of T3 level and RMR after weight loss program | |
| 16 Obese men | DDT | Resting metabolism rate (RMR) | Significant increase of concentrations of 13 organochlorines during weight loss program | |
| Nonmacronutrient-specific energy-restricted diet for 15 weeks | HCB | | Changes in organochlorine concentration inversely correlated with changes in serum T3 level (significant for p,p'-DDT, HCB, alochlor-1260, PCB-28, PCB-99, PCB-118 and PcB-170) and RMR (significant for HCB and PCB-156) after correction for weight loss | |
| | mirex | | | |
| | oxychlordane | | | |
| | trans-nonachlor | | | |
| | alroclor-1260 | | | |
| | PCBs | | | |