

# **Can we maximize nutritional intake while minimizing toxic risk from fish consumption?**

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**An update of our knowledge on mercury and omega-3 fatty acids from marine and fresh-water fish consumption**

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# **The Madison Declaration on Mercury Pollution**

**Henry Anderson, USA**

**Laurie Chan, Canada**

**Kathryn Mahaffey, USA**

**Donna Mergler, Canada**

**Michael Meyer, USA**

**Michael Murray, USA**

**Mineshi Sakamoto, Japan**

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*The Panel on Health Risks and Toxicological Effects of Methylmercury:*  
Donna Mergler, Henry A. Anderson, Laurie Hing Man Chan, Kathryn R. Mahaffey, Michael Murray,  
Mineshi Sakamoto and Alan H. Stern

## **Methylmercury Exposure and Health Effects in Humans: A Worldwide Concern**

- **“Methylmercury is a highly toxic compound that biomagnifies through the aquatic food web, placing at risk humans who consume significant quantities of predatory fish ... or who rely heavily on fish as a food source.”**
- **“..there is growing evidence that current exposures are sufficient to alter normal function of several physiological and developmental systems.. Long-lasting effects of fetal methylmercury exposure have been described in children throughout the world.”**
- **“Current studies suggest that exposure to methylmercury could increase the risk of adverse cardiovascular effects in a significant fraction of the human population.”**

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## **Methylmercury Exposure and Health Effects in Humans: A Worldwide Concern**

- **“Fish can contain both methylmercury and beneficial omega-3 fatty acids. Methylmercury exerts toxicity and can also diminish the beneficial health effects of omega-3 fatty acids....”**
- **“There is some evidence from animal studies showing that selenite protects against inorganic mercury toxicity. However, there is almost no evidence showing protection against methylmercury toxicity by organo-selenium compounds... found in the human diet.”**

# **Nervous system effects in children**

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## **Methylmercury Exposure and Health Effects in Humans: A Worldwide Concern**

- **The panel discussed at length the different findings from the 2 major birth cohort studies in the Faroes and the Seychelles Islands studies:**
  - **The Faroes study has consistently shown associations between cord blood Hg and neurobehavioral deficits and electrophysiological changes up to 14y**
  - **The Seychelles study only showed delayed development in their study among the most highly exposed (mothers' hair Hg) at 9 years old.**

# Recent findings from the Seychelles study

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- **Re-analysis of the data from the Seychelles study of the 9 year olds suggests that susceptibility may not be homogeneous:**
  - **Motor proficiency and activity level improved significantly with increasing MeHg for children who had an average home environment.**
  - **However, motor proficiency significantly decreased with increasing prenatal MeHg exposure in children whose home environment was below average.**

(Huang et al, 2008)

# A new Seychelles cohort

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- In a new cohort study the Seychelles, extensive data was obtained on dietary factors that may positively influence neurobehavioral development.
- An adverse association between MeHg and the mean Psychomotor Developmental Index (PDI) score at 30 months, when nutritional factors were included in the multiple regression model.

(Davidson et al, 2008)

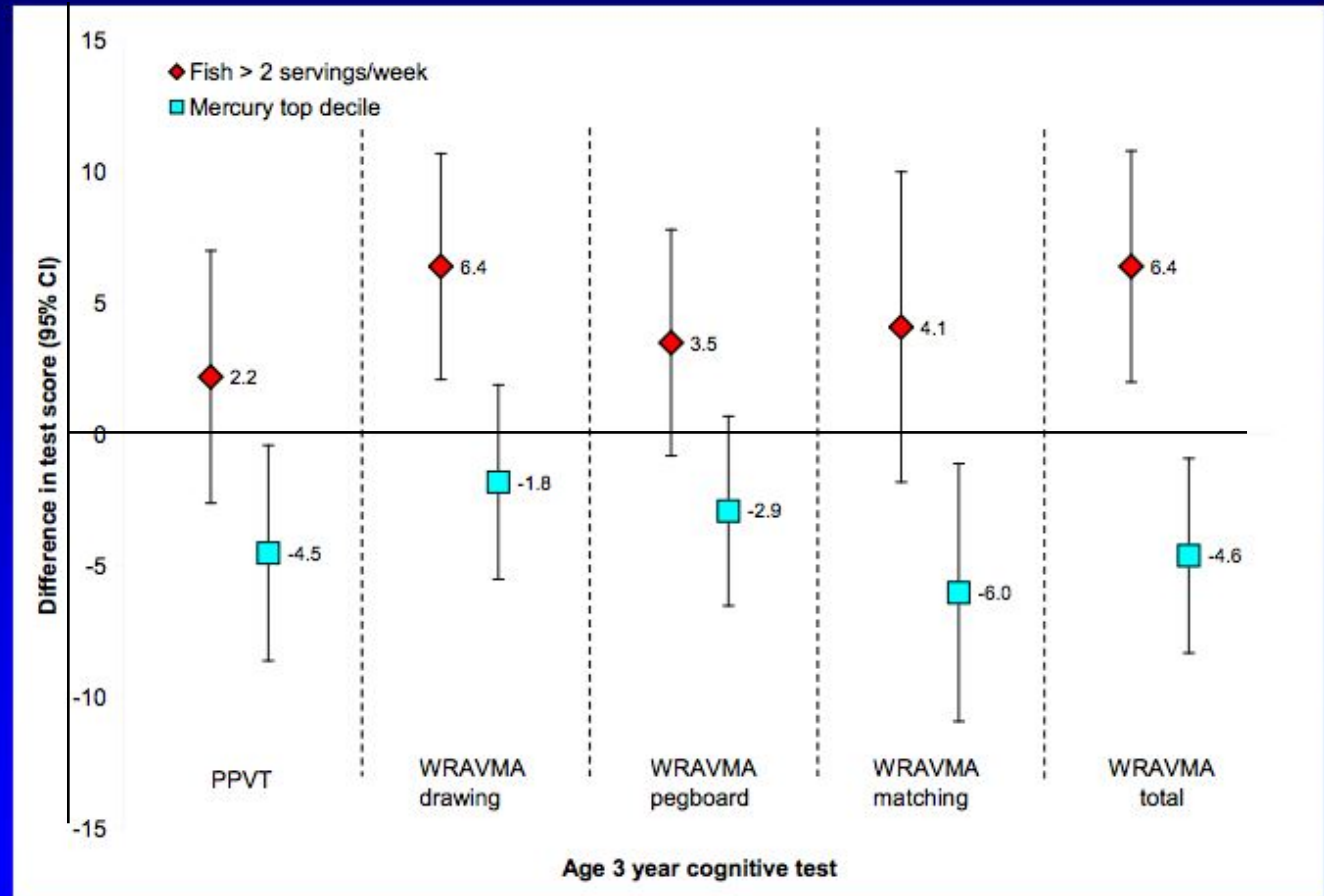
# Opposing effects of Hg and fish consumption on neurobehavioral performance in 3 year olds

N = 341  
mother/child pairs  
(Massachusetts)

Mean maternal  
total fish intake  
: $1.5 \pm 1.4$   
servings/week

40 (12%) mothers  
consumed >2  
servings/week

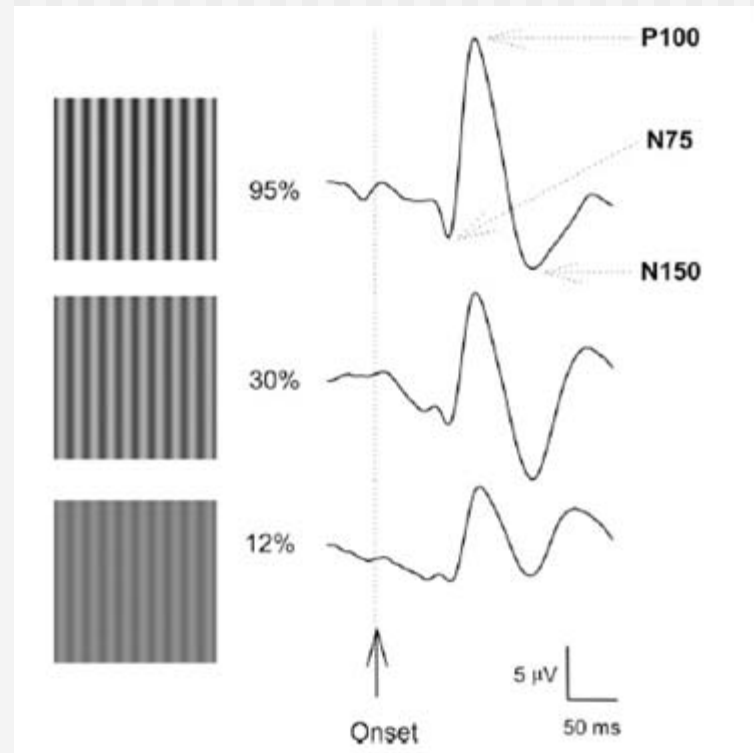
Mean maternal  
hair mercury level:  
3.8 ng/g. Top  
decile >1.2 $\mu$ g/g



Oken et al. 2008

# A study with Inuit children reported Hg-induced electrophysiological changes

- In Canadian Inuit children, prenatal and current Hg exposure were associated with changes in latencies for Visual Evoked Potentials



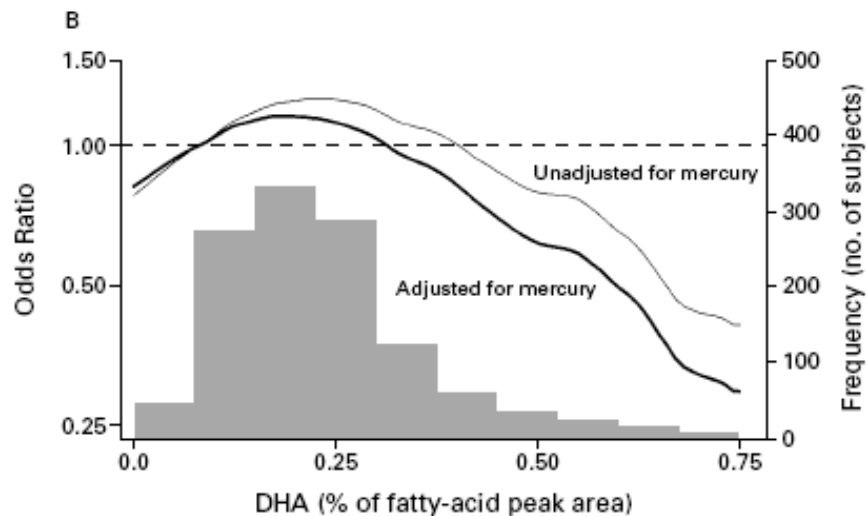
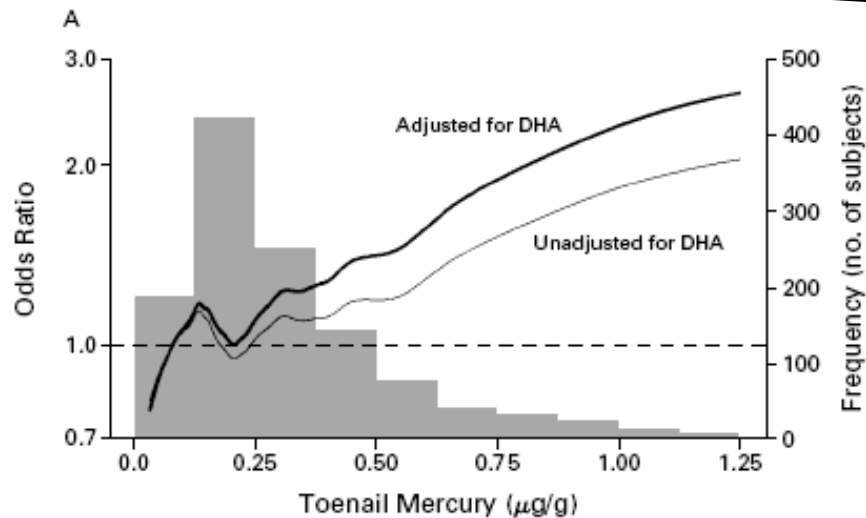
(St-Amour et al, 2006)

# **Cardiovascular system in adults**

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## Methylmercury Exposure and Health Effects in Humans: A Worldwide Concern



**Case control  
study of 684 men  
with a first  
diagnosis of  
myocardial  
infarction and 724  
controls**

**(Guallar et al, 2004)**

# Recent studies on myocardial infarction

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- **At the Mercury as a Global Pollutant Meeting in June 2009:**
  - **Jykri Virtanen presented data showing an increased incidence of myocardial infarction in a Finnish longitudinal cohort study in relation to mercury exposure,**
  - **Bengt Vessby presented data from Sweden showing decreased risk with Hg exposure in Swedish study; the authors consider that at these lower exposures, Hg is a proxy for fish consumption**

# Cardiovascular function

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- Blood Hg was associated with changes in heart rate variability (HRV) and increased systolic pressure in Canadian Inuit men and women (Valera et al, 2008)
- In Faroese whalers, Hg exposure was associated with increased blood pressure and common carotid intima-media thickness, but HRV was equivocal (Choi et al, 2009)

# Systolic Blood Pressure in a Canadian Inuit Population (n =731)

	Beta estimate	p
Blood Hg (log)	+1.27	0.05
Serum %EPA	- 1.75	0.05
Blood Se	- 2.80	0.03
Blood Hg (log)	+2.14	0.0004

- EPA and Hg did NOT modify the relation between blood Hg and blood pressure (interaction term not significant)
- Not adjusting for these elements could underestimate effect size

# Since the Madison Declaration in 2006

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- **Mercury developmental neurotoxicity has been confirmed at very low doses**
- **There is more evidence for mercury-induced cardiovascular alterations and illness**
- **There is growing evidence that omega-3 Fatty Acids (FA) and possibly Selenium (Se) can offset some of the toxic effects of mercury**
- **BUT, is increased fish consumption synonymous with increased omega-3 FA and Se?**

# Nutrients from fish consumption

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- There is a lot of information on omega-3 and Se from marine fish
- What about freshwater fish eaters?

# **Omega-3 Fatty Acids in serum of freshwater fisheaters**

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- **Great Lake fish-eaters (Cole et al, 2002)**
  - No relation between GL fish consumption and serum omega-3 FA for Euro-Canadians (n=45)
  - Significant relation between GL fish meals and serum DHA for Asian Canadians (n = 41), but not EPA
  - Significant relation between consumption of “other” fish meals and serum omega-3 FA.
- **Sportfishers (n=112) in the fluvial lakes of the St. Lawrence (Godin et al, 2003)**
  - No relation between St. Lawrence fish consumption and serum omega-3 FA

# Fishermen at James Bay, Quebec (n= 31)

	Prior to fishing season (june)	Post fishing season (November)	difference
Blood Hg (nmol/L)	21.9 ± 3.7	35.6 ± 5.2	+ 63%***
Hair Hg (µg/g)	1.4 ± 0.3	2.8 ± 0.4	+100%***
Blood Se (µg/L)	242.9 ± 6.2	247.7 ± 5.6	-
EPA + DHA (%)	4.92 ± 0.20	5.30 ± 0.60 %	-

# Omega-3 Fatty Acids in serum of freshwater fisheaters

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- A study of 259 persons who eat fish from lakes of the St. Lawrence and Abitibi
- Extensive food frequency questionnaire on fish species from local lakes, other lakes and market (frequency and portion = g/day)
- Measures of multiple contaminants, fatty acids and selenium (Se)
- Using published data, omega-3 FA from each species was estimated

(Philibert et al. 2006)

# Results

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- Highly significant ( $p < 0.0001$ ) relation between fatty fish intake and serum omega-3 F
- Highly significant ( $p < 0.0001$ ) relation between estimated omega-3 FA intake and serum omega-3 FA
- No relation between local catch consumption and serum omega-3 FA
- No relation between estimated omega-3 FA intake from local catch consumption and serum omega-3 FA

(Philibert et al. 2006)

# Fish consumption and biomarkers

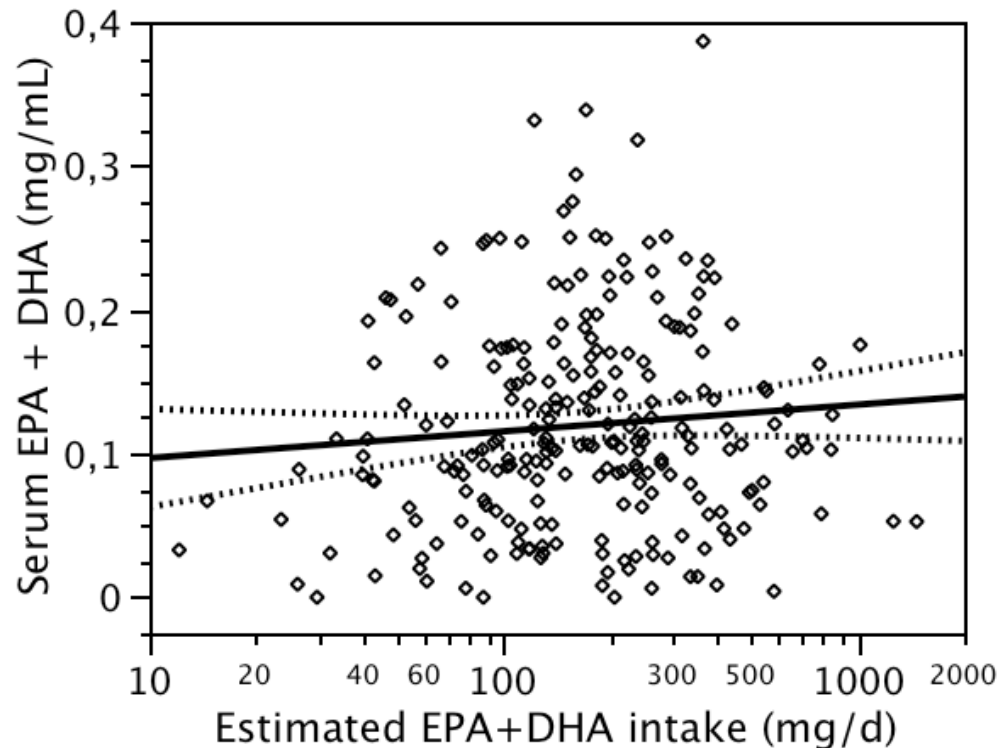
	mean intake (g/day)	% EPA + DHA	Blood Se (µg/L)	Hair Hg (µg/g)
<b>Total fish</b>	<b>57.1 ± 64.9 (3.26 - 641)</b>	<b>ns</b>	<b>ns</b>	<b>↑ **</b>
<b>Local catch</b>	<b>26.7 ± 59.9</b>	<b>ns</b>	<b>ns</b>	<b>↑ ***</b>
<b>Lean marine</b>	<b>18.2 ± 18.4</b>	<b>ns</b>	<b>ns</b>	<b>ns</b>
<b>Fatty marine</b>	<b>6.09 ± 8.70</b>	<b>↑ *</b>	<b>↑ **</b>	<b>ns</b>
<b>Trout</b>	<b>1.63 ± 3.77</b>	<b>↑ *</b>	<b>ns</b>	<b>↑ **</b>

# A conundrum

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- This group ate on average 57 g/day of fish (median: 40 g/day)
- Based on published estimates of EPA and DHA in fish species, mean EPA + DHA intake would be 225 mg/d  $\pm$  202 (median: 171 mg/day)
- But: EPA+ DHA levels were low: 0.12 mg/mL  $\pm$  0.07
- The % EPA + DHA in fatty acids were : 2.1%  $\pm$  1
- Overall, there was no relation between estimated intake and serum EPA+DHA

# No overall relation between estimated EPA + DHA intake and serum EPA + DHA



**$R^2 = 0.002$  ;  $p = 0.45$**

# Conclusions from these studies

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- The results from these studies confirm the relation between estimated omega-3 FA in fish and serum omega-3 FA for fatty fish
- For lean fish, this relation does not appear to hold,
  - Animal studies suggest that fatty fish oils may be necessary for the assimilation of EPA and/or DHA in plasma
  - Seasonal and fish size variations in EPA and DHA may be more important in lean fish
  - Cooking differences? Frying fish reduces omega-3
- Marine fatty fish appear to be the only fish contributor to blood Se.

## ■ Why is this important?

- Because according to current estimates and guidelines, many people who are eating primarily lean fish (which includes many freshwater fish) think that they are getting adequate nutrient intake...
- Because when regulators estimate benefits, they may overestimate for lean fish consumption

## ■ What to do?

# Fishermen at James Bay, Quebec (n= 31)

	Prior to fishing season (june)	Post fishing season (November)	difference
Cholesterol VLDL (mmol/L plasma)	$0.60 \pm 0.04$	$0.55 \pm 0.04$	- 8%*
Cholesterol VLDL (mmol/L plasma)	$0.77 \pm 0.04$	$0.81 \pm 0.05$	+5%*
GPx (U/g Hb)	$75.1 \pm 2.3$	$82.4 \pm 2.8$	+ 9.7%**
Beta-carotene ( $\mu$ mol/L)	$0.37 \pm 0.04$	$0.54 \pm 0.07$	+ 46%*

Bélanger et al, 2008

# Maximize nutrition, minimize risk

