DECISIONS

A young woman concerned about mercury

Stefanos N. Kales MD MPH, Aaron M.S. Thompson MD MPH

A healthy 30-year-old woman is planning to conceive. She seeks advice on having her dental amalgam fillings removed because of concerns about mercury toxicity. She also asks to be tested for mercury.

What questions should be asked to assess the patient's exposure to mercury?

The patient should be asked about potential sources of elemental, inorganic and organic mercury (Appendix 1, available at www.cmaj.ca/lookup/suppl/doi:10.1503/cmaj.150669/-/DC1).¹ A dietary history documenting the types and amount of fish consumed (especially large predatory fish, such as tuna, swordfish and shark) will provide information on potential exposure to organic mercury (methylmercury). Dental amalgam releases small amounts of elemental mercury vapour. Therefore, the physician should inquire about the number and integrity of the patient's fillings, her chewing habits and bruxism. In the occupational history, the physician should inquire about working with or around processes involving mercury.¹ Inadvertent inhalation of mercury vapour from broken instrumentation or fluorescent light bulbs is another potential, albeit infrequent, source of exposure to elemental mercury.¹ Patients may also come into contact with inorganic mercury salts through topical antiseptics on disrupted skin or other tissues.

Although not a clinically relevant source of mercury, patients may have concerns about the vaccine preservative thimerosal, which is partly metabolized to ethylmercury. Ethylmercury is less neurotoxic than methylmercury and is more rapidly excreted.² Thimerosal is not used in pediatric vaccines in Canada other than those for influenza; similarly, it is generally not used in adult vaccines, with only a few exceptions, such as some hepatitis B preparations. A case-control study indicated that levels of mercury in infants who received routine immunization with thimerosal-containing vaccines did not exceed guidelines for methylmercury.³ Moreover, prenatal and early-life exposure to ethylmercury from thimerosal-containing vaccines does not increase the risk of

autism, according to the results of case-control and retrospective cohort studies.^{4,5}

What should be included in the history and physical examination for this patient?

Symptom history and physical examination generally do not contribute to a clinical assessment of mercury because levels are almost always well below toxicity thresholds. In rare instances of high intake of large predatory fish, the focus of the clinical evaluation is the central nervous system. Signs and symptoms documented in historical poisoning outbreaks include perioral paresthesia, dysarthria, visual field defects and ataxia.¹ In cases of occupational or inadvertent inhalation of mercury vapour, the clinical assessment should focus on the central nervous system (tremor, ataxia, emotional instability), the peripheral nervous system (distal sensory loss) and the renal system.¹

Should mercury testing be ordered for this patient?

Testing is generally not indicated but may be considered in cases of high consumption of large predatory fish or exposure to mercury vapour (inadvertent or occupational) (Box 1).⁶ If such testing is clinically indicated, it is prudent to order measurement of both blood and urine mercury levels. Blood mercury primarily reflects methylmercury (from consumption of fish), whereas urine mercury corrected for creatinine concentration primarily reflects exposure to elemental and inorganic mercury.⁷ Unconventional testing methods (e.g., commercial hair analysis, urine mercury challenge or "provoked" testing using a chelator) should be avoided.⁸

Before ordering a mercury test, the physician should explain to the patient that laboratory reference ranges are population averages and do not reflect toxicity thresholds. Blood mercury levels are usually less than 25 nmol/L (5 µg/L) among those who eat little fish, whereas levels up to 100 nmol/L (20 µg/L) may be seen in people who eat fish four to seven times a week. Urine mercury levels are also generally below 25 nmol/L (5 µg/L).⁹ In con-

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The clinical scenario is fictional.

Correspondence to:

Stefanos Kales,
skales@hsph.harvard.edu

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Box 1: Choosing Wisely Canada recommendation on mercury testing⁶

Don't order blood mercury testing unless dietary history suggests risk, the patient is pregnant or planning to become pregnant, and/or the patient is occupationally exposed to organomercury compounds.

- For adults, Health Canada's guidance value for total blood mercury concentrations is 40 nmol/L (8 µg/L) for women of child-bearing age and 100 nmol/L (20 µg/L) for women ≥ 50 years and men > 18 years.
- Although clinically significant exposures may still occur in Canada, less than 1% of Canadian adults have total blood mercury concentrations above Health Canada's guidance value. As such, the large majority of individuals who present with concerns about metal toxicity do not actually have toxicity, and testing results in false positives (values above the reference range but not in the range of toxicity).
- Occupationally exposed workers and women of child-bearing age are susceptible subgroups; therefore, testing in these populations is warranted in cases where a careful occupational and/or environmental history suggests significant exposure.
- In the absence of clinical presentation and history indicating a risk of toxicity, testing should be avoided because it may lead to misinterpretation and unnecessary concern or interventions (dietary restriction, chelation) that may cause harm.

trast, documented clinical toxicity is associated with levels greater than 500 nmol/L (100 µg/L) in urine and 1000 nmol/L (200 µg/L) in blood.¹⁰ For prenatal exposure, prospective cohort studies have suggested that the risk of subtle neurocognitive effects may start at maternal blood mercury levels as low as 200 nmol/L (40 µg/L),⁹ although guidance values are much lower because of the application of safety factors.

How should the patient be counselled about dental amalgam?

During counselling, the patient should be advised that, according to evidence from randomized clinical trials,¹¹ dental amalgam does not result in toxic effects and that replacement of amalgam fillings with nonmercury materials is not recommended. A precautionary approach could include avoiding dental amalgam work during pregnancy unless absolutely necessary, although no clinical evidence supports this recommendation.

How should the patient be counselled about fish consumption?

Women of child-bearing age should be encouraged to include fish in their diet and to preferentially choose species low in mercury and high in n-3 polyunsaturated fatty acids (e.g., salmon and char). This recommendation is based on evidence from randomized trials of maternal supplementation with n-3 polyunsaturated fatty acids, as well as observational cohort studies of neurodevelopmental outcomes and maternal fish consumption.¹²

The case revisited

The patient was counselled that current evidence does not support any risk from dental amalgam

fillings and was given information from the Canadian Dental Association (https://www.cda-adc.ca/_files/position_statements/amalgam.pdf). The patient reported eating two to three fish meals per week and was reassured that, on the basis of her exposure history, mercury testing was not indicated. She was counselled to continue to include fish in her diet and was given information on relative amounts of methylmercury in different fish species, along with Health Canada's advice on fish consumption (www.hc-sc.gc.ca/fn-an/securit/chem-chim/envIRON/mercur/cons-adv-etud-eng.php).

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Affiliations: Environmental and Occupational Medicine and Epidemiology (Kales), Harvard T.H. Chan School of Public Health, Harvard University, Boston, Mass.; The Cambridge Health Alliance (Kales), Harvard Medical School, Cambridge, Mass.; Faculty of Medicine (Thompson) and Dalla Lana School of Public Health (Thompson), University of Toronto, Toronto, Ont.; Department of Occupational and Environmental Health (Thompson), St. Michael's Hospital, Toronto, Ont.

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CMAJ is collaborating with Choosing Wisely Canada (www.choosingwiselycanada.org), with support from Health Canada, to publish a series of articles describing how to apply the Choosing Wisely Canada recommendations in clinical practice.