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8

9 **Manganese**

10 **Introduction**

11 Chemically, manganese is closely related to iron. It is a catalytic cofactor for arginase,
 12 pyruvate carboxylase and mitochondrial superoxide dismutase (SOD), but also
 13 functions as a specific or unspecific activator for a large number of other enzymes,
 14 some of which participate in the synthesis of proteins, mucopolysaccharides and
 15 cholesterol.

16

17 **Dietary sources and intake**

18 Unrefined cereals, nuts and leafy vegetables have high manganese content. Tea is a
 19 substantial contributor to manganese intake, containing about 2.7 mg/L. Accordingly,
 20 manganese intake varies from very low, <2 mg/day, to high, >8 mg/day, in vegetarian
 21 diets.

22

23 The average content of manganese in the Swedish diet analysed in supermarket baskets
 24 and duplicate portions collected in the late 1980s was 3.6-3.7 mg/day (Becker et al
 25 1997, Jorhem et al 1998). A Danish study where 100 men collected duplicate portions
 26 of their regular diets for 48 hours showed a manganese intake of 3.9 mg/day (Bro et al
 27 1990). The manganese intake of Finnish children 3-18 years of age was in the range of
 28 3-7 mg/day calculated from food consumption data and food contents (Bro et al 1990).
 29 These data indicate that manganese intake is adequate in these countries. Multivitamin-
 30 mineral and mineral supplements for adults may provide 2-5 mg manganese/dose.

31

32 **Physiology and metabolism**

33 The total body content of manganese is estimated to be 10-20 mg. The concentration is
 34 relatively high in bone and in organs rich in mitochondria, such as liver, pancreas and
 35 kidney, while muscle and plasma have low concentrations. Absorption from the diet is
 36 low, approximately 5 %, and excretion is primarily through the bile. Animal studies
 37 have shown that iron, calcium and phytic acid reduce the absorption of manganese
 38 (Hurley et al 1987). A negative effect of calcium has been shown in humans, while the
 39 effect of iron and phytic acid does not seem to be pronounced (Davidsson et al 1991).
 40 High intakes of manganese inhibit iron absorption (Rossander-Hulten 1991), and a
 41 higher absorption of manganese has been reported in iron deficiency (Mena et al 1969,
 42 Meltzer et al 2010).

43

44 Manganese deficiency in experimental animals results in reduced growth, skeletal
 45 abnormalities and defects in lipid and carbohydrate metabolism (Hurley et al 1987). In
 46 humans, only a limited number of possible manganese deficiency symptoms have been
 47 described in experimental studies with a manganese-deficient diet (Friedman et al

48 1987). Dermal changes and hypocholesterolaemia are possible signs of manganese
49 deficiency, as well as diffuse bone demineralization and poor growth in children. Very
50 little information is available concerning the relationship between manganese intake
51 and health endpoints or disease prevention (Brown et al. 2012).

52

53 **Requirement and recommended intake**

54 Our knowledge of manganese metabolism and the consequences of low intakes are
55 insufficient for determining requirements and recommended daily intakes for humans.
56 Balance studies have suggested that an intake of 0.74 mg/day should be sufficient to
57 replace daily losses of manganese (Freeland et al 1988). Intakes over 1 mg/day
58 generally result in a positive manganese balance (Brown et al. 2012).

59

60 The EU Scientific Committee for Food (1993) considered a ‘safe and adequate intake’
61 to be 1-10 mg/person/day.

62

63 The US Food and Nutrition Board (2001) found data to be insufficient for setting an
64 Estimated Average Requirement (EAR) for manganese, but used median intakes
65 reported from the US Total Diet Study 1982-9 as a basis for setting adequate intakes
66 (Pennington and Young 1991). The AI for adult men and women is set at 2.3 and 1.8
67 mg/day, respectively. In 1993, the EU Scientific Committee for Food (1993) suggested
68 1-10 mg/day to be an acceptable intake of manganese.

69

70 The Nordic Recommendations of 2004 did not include recommendations for
71 manganese intake. As very few relevant human studies have been conducted since
72 then, requirements are also difficult to determine this time, and accordingly,
73 recommendations are not given for any age group.

74

75 Data are also too limited to determine requirements for manganese during pregnancy
76 and lactation, and manganese deficiency of pregnant or lactating women has not been
77 observed in humans. Manganese excretion from breast milk is estimated to be below
78 1% of the total manganese excretion. There is no clear correlation between dietary
79 intake and breast milk manganese concentration (Brown et al. 2012). In a systematic
80 review of studies, including studies published from January 1990 to October 2011, 15
81 studies reporting breast milk manganese concentration were retrieved (Brown et al.
82 2012), with levels ranging from 0.8-30 µg/L. Median (SD) manganese concentration
83 of 31 Swedish milk samples was found to be 3.23 (0.27) µg/L (Parr et al. 1991).

84

85 **Upper intake levels and toxicity**

86 Manganese is regarded as one of the least toxic trace elements. Manganese toxicity,
87 which manifests as psychological and neurological changes, has been observed in
88 workers in manganese mines (Mena et al 1969). The neurological symptoms are
89 reminiscent of those seen in Parkinson’s disease. Inhalation of manganese dust may be
90 the explanation, while toxicity caused by a high dietary intake is unknown. The EU
91 Scientific Committee for Food (2006) found that data for setting a Tolerable Upper
92 Intake Level of manganese were too uncertain. The UK Foods Standards Agency
93 (2003) has also found data to be insufficient to establish a Safe Upper Level for
94 manganese.

95

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