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Manganism

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Manganism or **manganese poisoning** is a **toxic** condition resulting from chronic exposure to **manganese**.^[1] It was first identified in 1837 by **James Couper**.^[2]

Signs and symptoms

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Chronic exposure to excessive manganese levels can lead to a variety of psychiatric and motor disturbances, termed manganism. Generally, exposure to ambient manganese air concentrations in excess of 5 micrograms Mn/m³ can lead to manganese-induced symptoms.^[3]

In initial stages of manganism, neurological symptoms consist of reduced response speed, irritability, mood changes, and compulsive behaviors.^[4] Upon protracted exposure symptoms are more prominent and resemble those of **idiopathic Parkinson's disease**, as which it is often misdiagnosed, although there are particular differences in both the symptoms; for example, the nature of the tremors, response to drugs such as **levodopa**, and affected portion of the **basal ganglia**. Symptoms are also similar to **Lou Gehrig's disease** and **multiple sclerosis**.

Causes

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Manganism has become an active issue in **workplace safety** as it has been the subject of numerous **product liability** lawsuits against manufacturers of **arc welding** supplies. In these lawsuits, welders have accused the manufacturers of failing to provide adequate warning that their products could cause welding fumes to contain dangerously high manganese concentrations that could lead welders to develop manganism. Companies employing welders are also being sued, for what colloquially is known as "welders' disease." However, studies fail to show any link between employment as a welder and manganism (or other neurological problems).^{[5][6][7]}

Manganism is also documented in reports of illicit **methcathinone** manufacturing.^[8] This is due to manganese being a byproduct of methcathinone synthesis if **potassium permanganate** is used as an oxidiser.^[9] Symptoms include apathy, bradykinesia, gait disorder with postural instability, and spastic-hypokinetic **dysarthria**. Another street drug sometimes contaminated with manganese is the so-called "Bazooka", prepared by **free-base** methods from **cocaine** using manganese **carbonate**.^[10]

Reports also mention such sources as contaminated drinking water,^[11] and fuel additive **methylcyclopentadienyl manganese tricarbonyl** (MMT),^[12] which on **combustion** becomes partially converted into manganese **phosphates** and **sulfate** that go airborne with the exhaust,^{[13][14][15]} and **manganese ethylene-bis-dithiocarbamate** (**Maneb**), a pesticide.^[16]

And in very rare cases it can be caused by a defect of the gene **SLC30A10**.

Pathophysiology

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Manganese may affect **liver** function, but the threshold of acute toxicity is very high. On the other hand, more than 95 percent of manganese is eliminated by biliary excretion. Any existing liver damage may slow this process, increasing its concentration in blood plasma.^[17] The exact **neurotoxic** mechanism of manganese is uncertain but there are clues pointing at the interaction of manganese with **iron**,^{[18][19][20][21]} **zinc**,^[22] **aluminum**,^{[18][22]} and **copper**.^[22] Based on a number of studies, disturbed iron **metabolism** could underlie the neurotoxic action of manganese.^[23]

It participates in **Fenton reactions** and could thus induce **oxidative damage**, a hypothesis corroborated by the evidence from studies of affected welders.^[24] A study of the exposed workers showed that they have significantly fewer children.^[25] This may indicate that long-term **accumulation** of manganese affects **fertility**. Pregnant animals repeatedly receiving high doses of manganese bore malformed offspring significantly more often compared to controls.^[26] It is found in large quantities in paint and steelmaking.

Diagnosis

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Treatment

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The current mainstay of manganism treatment is **levodopa** and **chelation** with **EDTA**. Both have limited and at best transient efficacy. Replenishing the deficit of **dopamine** with levodopa has been shown to initially improve **extrapyramidal symptoms**.^{[27][28][29]} but the response to treatment goes down after 2 or 3 years,^[30] with worsening condition of the same patients noted even after 10 years since last exposure to manganese.^[31] Enhanced excretion of manganese prompted by **chelation therapy** brings its blood levels down but the symptoms remain largely unchanged, raising questions about efficacy of this form of treatment.^{[32][33]}

Increased **ferroportin** protein expression in human embryonic kidney (**HEK293**) cells is associated with decreased intracellular manganese concentration and attenuated cytotoxicity, characterized by the reversal of manganese-reduced **glutamate** uptake and diminished **lactate dehydrogenase** (LDH) leakage.^[3]

Epidemiology

[edit]

The **Red River Delta** near **Hanoi** has high levels of manganese and **arsenic** in the water. Approximately 65 percent of the region's wells contain high levels of arsenic, manganese, **selenium**, and **barium**.^[34]

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- Welding Rod Litigation Information Network* — Viewpoint of defense on welding rod litigation

External links

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Classification	ICD-10: T57.2 · ICD-9-CM: 985.2 · MeSH: D020149 · SNOMED CT: 88687001 	 D
V · T · E	Poisoning · Toxicity · Overdose	[show]
Categories: Manganese Neurological disorders Toxic effects of metals		

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