Hazard to children: birth defects; developmental, immune and endocrine effects; later in life: cancer, Parkinson's disease, female reproductive problems.





Pesticide Action Network Asia and the Pacific

P.O. Box 1170 10850 Penang, Malaysia Tel: (604) 657 0271 / 656 0381 Fax: (604) 658 3960 Email: panap@panap.net Homepage: www.panap.net

Copyright ©Pesticide Action Network Asia and the Pacific. All

rights reserved.

Pesticide Action Network Asia and the Pacific (PAN AP) encourages the reproduction and use of this publication as long as PAN AP is properly acknowledged as the source and provided with

Meriel Watts, PhD June 2014 **Uses**: dithiocarbamate fungicide; contains zinc, manganese (essential element but neurotoxic in excess); metabolites include ETU.

Residues: drinking water, food; manganese in hair of exposed pregnant women.¹

Acute toxicity: eye

irritation, skin rashes, dermatitis, nausea, dizziness.² Children have been poisoned in Nicaragua.³ Commonly reported cause of poisoning in Tanzania.⁴

Chronic toxicity: liver, brain, kidney (rats).⁵

Neurological: damage to peripheral nerves with abnormal gait and loss of muscle mass (rats);² prenatal exposure alters developing brain (mice);⁶ in other species neurodegeneration and behavioural changes.⁷⁸ Associated with Parkinson's disease.⁹⁻¹¹

Cancer: US EPA probable human carcinogen; in rats tumours of thyroid, liver, pituitary;² mammary glands, ear, pancreas, bones of the head;¹² in mice foetal cells¹³ and skin.¹⁴ Associated with leukaemia,¹⁵ melanoma¹⁶ breast cancer risk.¹⁷

Genotoxicity:

genotoxic in human cells.¹⁸

A PANAP Factsheet Series Highly Hazardous Pesticides Mancozeb

Endocrine disruption:

causes thyroid damage, tumours and altered hormones;² associated with hypothyroidism and hyperthyroidism in women;¹⁹ antiandrogenic.²⁰

Reproduction:

Ovarian toxicant.^{21 22} Birth defects e.g. hydrocephaly, skeletal system defects in rats;² neural tube defects in humans.²³ Reduced female fertility (mice).²¹ Induces a pre-malignant state in ovarian follicles; ^{13 22} increased length of menstrual cycle, missed periods.²⁴

Immune: alters immune system response;²⁵⁻²⁷ allergic sensitisation.²

Environmental effects:

Aquatic: very highly toxic to fish and aquatic invertebrates; risk to freshwater fish and invertebrates; fish kills.²

Terrestrial: chronic risks to birds and mammals, including reproductive and potential endocrine disruption; toxic to some beneficial insects.²²⁸

Environmental fate:

Can cause severe accumulation of manganese in soil, and ETU in surface water.²⁹

References

¹ Mora AM, van Wendel de Joode B, Mergler D, Córdoba L, Cano C, Quesada R, Smith DR, Menezes-Filho JA, Lundh T, Lindh CH, Bradman A, Eskenazi B. 2014. Blood and hair manganese concentrations in pregnant women from the infants' environmental health study (ISA) in Costa Rica. *Environ Sci Technol* 48(6):3467-76.

² US EPA. 2005. Reregistration Eligibility Decision (RED) Mancozeb.

³ Watts MA. 2013. *Poisoning Our Future: Children and Pesticides.* Pesticide Action Network Asia & the Pacific, Penang.

⁴Lekei E, Ngowi AV, London L. 2014. Farmers' knowledge, practices and injuries associated with pesticide exposure in rural farming villages in Tanzania. *BMC Public Health* 14:389.

⁵ Kackar R, Srivastava MK, Raizada RB. 1999. Assessment of toxicological effects of mancozeb in male rats after chronic exposure. *Indian J Exp Biol* 37(6):553-9.

⁶ Miranda-Contreras L, Dávila-Ovalles R, Benítez-Díaz P, Peña-Contreras Z, Palacios-Prü E. 2005. Effects of prenatal paraquat and mancozeb exposure on amino acid synaptic transmission in developing mouse cerebellar cortex. *Brain Res Dev Brain Res* 160(1):19-27.

⁷ Negga R, Rudd DA, Davis NS, Justice AN, Hatfield HE, Valente AL, Fields AS, Fitsanakis VA. 2011. Exposure to Mn/Zn ethylene-bis-dithiocarbamate and glyphosate pesticides leads to neurodegeneration in Caenorhabditis elegans. *Neurotoxicology* 32(3):331-41.

⁸ Harrison Brody A, Chou E, Gray JM, Pokyrwka NJ, Raley-Susman KM. 2013. Mancozeb-induced behavioral deficits precede structural neural degeneration. *Neurotoxicology* 34:74-81.

⁹ US EPA. 2013. *Recognition and Management of Pesticide Poisonings.* 6th Edition.

¹⁰ Pezzoli G, Cereda E. 2013. Exposure to pesticides or solvents and risk of Parkinson disease. *Neurology* 80(22):2035-41. ¹¹ Erro ME, Muñoz R, Zandio B, Mayor S. 2011. Reversible Parkinsonism after accidental oral intake of mancozeb. *Mov Disord* 26(3):557-8.

¹² Belpoggi F, Soffritti M, Guarino M, Lambertini L, Cevolani D, Maltoni C. 2002. Results of long-term experimental studies on the carcinogenicity of ethylene-bis-dithiocarbamate (Mancozeb) in rats. *Ann N Y Acad Sci* 982:123-36.

¹³Shukla Y, Arora A. 2001. Transplacental carcinogenic potential of the carbamate fungicide mancozeb. *J Environ Pathol Toxicol Oncol* 20(2):127-31.

¹⁴ Shukla Y, Antony M, Kumar S, Mehrotra NK. 1990. Carcinogenic activity of a carbamate fungicide, mancozeb on mouse skin. *Cancer Lett* 53:191–5.

¹⁵ Mills PK, Yang R, Riordan D. 2005. Lymphohematopoietic cancers in the United Farm Workers of America (UFW), 1988-2001. *Cancer Causes Control* 16(7):823-30.

¹⁶ Dennis LK, Lynch CF, Sandler DP, Alavanja MC. 2010. Pesticide use and cutaneous melanoma in pesticide applicators in the agricultural heath study. *Environ Health Perspect* 118(6):812-7.

¹⁷ Watts MA. 2007. *Pesticides & Breast Cancer: A Wakeup Call.* Pesticide Action Network Asia & the Pacific, Penang.

¹⁸ Srivastava AK, Ali W, Singh R, Bhui K, Tyagi S, Al-Khedhairy AA, Srivastava PK, Musarrat J, Shukla Y. 2012. Mancozebinduced genotoxicity and apoptosis in cultured human lymphocytes. *Life Sci* 90(21-22):815-24.

¹⁹ Goldner WS, Sandler DP, Yu F, Hoppin JA, Kamel F, Levan TD. 2010. Pesticide use and thyroid disease among women in the Agricultural Health Study. *Am J Epidemiol* 171(4):455-64.

²⁰ Kjeldsen LS, Ghisari M, Bonefeld-Jørgensen EC. 2013. Currently used pesticides and their mixtures affect the function of sex 2 hormone receptors and aromatase enzyme activity. *Toxicol Appl Pharmacol* 272(2):453-64. ²¹ Paro R, Tiboni GM, Buccione R, Rossi G, Cellini V, Canipari R, Cecconi S. 2012. The fungicide mancozeb induces toxic effects on mammalian granulosa cells. *Toxicol Appl Pharmacol* 260(2):155-61.

²² Cecconi S, Paro R, Rossi G, Macchiarelli G. 2007. The effects of the endocrine disruptors dithiocarbamates on the mammalian ovary with particular regard to mancozeb. *Curr Pharm Des* 13(29):2989-3004.

²³ Nordby KC, Andersen A, Irgens LM, Kristensen P. 2005. Indicators of mancozeb exposure in relation to thyroid cancer and neural tube defects in farmers' families. *Scand J Work Environ Health* 31(2):89-96.

²⁴ Farr SL, Cooper GS, Cai J, Savitz DA, Sandler DP. 2004. Pesticide use and menstrual cycle characteristics among premenopausal women in the Agricultural Health Study. *Am J Epidemiol* 160(12):1194-204.

²⁵ Corsini E, Birindelli S, Fustinoni S, De Paschale G, Mammone T, Visentin S, Galli CL, Marinovich M, Colosio C. 2005. Immunomodulatory effects of the fungicide Mancozeb in agricultural workers. *Toxicol Appl Pharmacol* 208(2):178-85.

²⁶ Colosio C, Barcellini W, Maroni M, Alcini D, Bersani M, Cavallo D, Galli A, Meroni P, Pastorelli R, Rizzardi GP, Soleo L, Foà V. 1996. Immunomodulatory effects of occupational exposure to mancozeb. *Arch Environ Health* 51(6):445-51.

²⁷ Chung AH, Pyo MY. 2005. Effects of mancozeb on the activities of murine peritoneal macrophages in vitro and ex vivo. *Arch Pharm Res* 28(1):100-5.

²⁸ European Commission. 2009. Review Report of the active substance mancozeb. SANCO/4058/2001 - rev. 4.4.

²⁹ Geissen V, Ramos FQ, de J Bastidas-Bastidas P, Díaz-González G, Bello-Mendoza R, Huerta-Lwanga E, Ruiz-Suárez LE. 2010. Soil and water pollution in a banana production region in tropical Mexico. *Bull Environ Contam Toxicol* 85(4):407-13.