



Beispiel: Metall-Ionen induzierte toxische Histamin Freisetzung von menschlichen Basophilen und Mast-Zellen.

Metal ion-induced toxic histamine release from human basophils and mast cells. J Biomed Mater Res 1998 Mar 15;39(4):560-7 (ISSN: 0021-9304) Schedle A; Samorapoompitchit P; Fureder W; Rausch-Fan XH; Franz A; Sperr WR; Sperr W; Slavicek R; Simak S; Klepetko W; Ellinger A; Ghannadan M; Baghestanian M; Valent P. School of Dentistry, University of Vienna, Austria.

Abstract: Recent data suggest that distinct metal ions can be released from dental alloys or other biomaterials, and may cause **toxic** effects on various cells. In this study, the effects of 14 metal ions on histamine release from human blood basophils ($n = 4$), isolated tissue mast cells (lung $n = 8$, uterus $n = 2$, skin $n = 1$, gingiva $n = 1$), the basophil cell line KU-812, and the mast cell line HMC-1 were analyzed. Of the 14 metal ions, Ag^+ (0.33 mM) and Hg^{2+} (0.33 mM) were found to induce release of histamine in blood basophils, KU-812, mast cells, and HMC-1. The effects of Ag^+ and Hg^{2+} were dose dependent and were observed within 60 min of incubation. In primary mast cells and basophils, Au^{3+} (0.33 mM) also induced histamine release, whereas no effects of Au^{3+} on HMC-1 or KU-812 cells were seen. The other metal ions showed no effects on primary or immortal cells within 60 min. However, Pt^{4+} (0.33 mM) induced histamine liberation in HMC-1 and lung mast cells after 12 h. The Ag^{+-} and Hg^{2+-} -induced rapid release of histamine from HMC-1 was associated with ultrastructural signs of necrosis, but not apoptosis. In contrast, prolonged exposure to Pt^{4+} (0.33 mM, 14 h) induced apoptotic cell death in HMC-1 cells, as assessed by electron microscopy and DNA analysis. Together, certain metal ions induce distinct cytopathogenic effects in mast cells and basophils. Whereas Ag^+ , Hg^{2+} , and Au^{3+} cause direct **toxicity**, Pt^{4+} causes cell death through induction of apoptosis. Whether such effects contribute to local adverse reactions to metal-containing biomaterials *in vivo* remains to be determined.

Zusammenfassende Übersetzung der Ergebnisse:

Metall-Ionen aus zahnärztlichen Werkstoffen setzen in menschlichen Blutzellen Histamin frei, bei Silber- und Quecksilber-Ionen bereits nach 60 Minuten. Bei Platin-Ionen trat dieser Effekt erst nach 12 Stunden ein. Die Freisetzung von Histamin war bei den Blutzellen verbunden mit Veränderungen der Ultrastrukturen und bei längerer Einwirkung (Platin-Ionen nach 14 Stunden) mit Zelltod. Silber-, Quecksilber- und Gold-Ionen lösen direkte Toxizität aus, Platin-Ionen verursachen Zelltod durch Auflösung der Zelle.



Gibt es noch weitere Literatur zur **Toxizität der Zahnmetalle?**

Arendash GW: Metals and free radicals in neurodegeneration. Curr Opin Neurol 7 (1994) 548-558

Pedersen LM, Permin H: Rheumatic disease heavy metal pigments and the Great Masters. Lancet 1 (1988) 1267-1269

Belghiti D et al.: Lipoid nephrosis of toxic origin. 2 cases. Presse Med 15 (1986) 1953-1955

Joyce DA, Wade DN: Assay for D-penicillamine-protein conjugate in human plasma utilizing chemical reduction followed by high performance liquid chromatography with gold/mercury electrochemical detection. J Chromatogr 430 (1988) 319-327

Halliwell B et al.: Metalions and oxygen radical reactions in human inflammatory joint disease. Philos Trans R Soc Lond B Biol Sci 311 (1985) 659-671

Halliwell B, Gutteridge JM: Oxygen toxicity oxygen radicals transition metals and disease. Biochem J 219 (1984) 1-14

Stejskal J, Stejskal V: The role of metals in autoimmunity and the link to neuroendocrinology. Neuroendocrinology Letters 20 (1999) 351.-364

Zheng W et al.: Choroid plexus protects cerebrospinal fluid against toxic metals. FASEB J 5 (1991) 2188-2193

Walum E et al.: Use of primary cultures and continuous celllines to study on astrocytic regulatory functions. Clin Exp Pharmacol Physiol 22 (1995) 284-287

Vitetta ES, Thorpe PE, Uhr JW: Immunotoxins: Magic bullets or misguided missiles? Immunol Today 1993; 14: 252-259.



Gibt es weitere Literatur zur **immunologischen Wirkung der Zahnmetalle?**

Prochazkova Jet al.: Immunogenetic findings in patients with altered tolerance to heavy metals. Eur J Hum Genet 6 (1998) 175

Saito K: Analysis of a genetic factor of metal allergy -polymorphism of HLA-DR. -DQ gene. Kokubyo Gakkai Zasshi 63 (1996) 53-69

Goldman Met al: TH2 cells in systemic autoimmunity: insights from allogenic diseases and chemically induced autoimmunity. Immunol Today 12 (1991) 223-227

Murdoch RD. Pepys J: Enhancement of antibody production by mercury and platinum group metal halide salts. Kinetics of total and ovalbumin- specific IgE synthesis. Int Arch Allergy Appl Immunol 80 (1986) 405-411

Vassilev TL: Aluminum phosphate but not calcium phosphate stimulates the specific IgE response in guinea pigs to tetanus toxoid. Allergy 33 (1978) 155-159

Biagini RE et al.: The diversity of reaginic immune responses to platinum and palladium metallic salts. J Allergy Clin Immunol 76 (1985) 794-802

Bergman A et al.: Contact urticaria with anaphylactic reactions causes by . occupational exposure to iridium salt. Contact Dermatitis 35 (1995) 14-17

Penz MG et al.: In vitro analysis of lymphocyte reactivity to nickel (II) in patients with nickel contact dermatitis. Eur J Labor Med 7 (1999) 1-8

Stejskal V: MELISA -an in vitro tool for the study of metal allergy. Toxicology in vitro 8 (1994) 991-1000

Stejskal VDM et al.: Metal-specific lymphocytes: biomarkers of sensitivity in man. Neuroendocrinol Lett 20 (1999) 289-298

Casciola-Rosen Let al.. Scleroderma autoantigens are uniquely fragmented by metal-catalyzed oxidation reactions: implications for pathogenesis. J Exp Med 185 (1997) 71-79

Bigazzi P: Autoimmunity induced by metals. In: Chang L (ed.): Toxicology of metals. USA: Lewis Publishers, CRC Press Inc. (1996) 835-852 "



Akademie für Integrative Medizin, Zahnmedizin und Bewusstseinsttechniken

Fritzler MJ: Autoantibodies: diagnostic fingerprints and ethiologic perplexities. Clin Invest Med 20 (1997) 50-66

Monestier Met al.: D-penicillamine- and quinidine-induced antinuclear antibodies in A.SW (H-2s) mice: similarities with autoantibodies in spontaneous and heavy metal induced autoimmunity. Eur J Immunol 24 (1997) 723-730

Hultman P et al.: Selective induction of antifibrillarin autoantibodies by silver nitrate in mice. Clin Exp Immunol 96 (1994)

Kusaka Y: Occupational diseases caused by exposure to sensitizing metals. Sangyo Igaku 35 (1993) 75-87

Kohdera T et al.: Antigen-specific Lymphocyte stimulation test on patients with psoriasis vulgaris. Proceedings of the XVI International Congress of Allergology and Clinical Immunology (1997) Cancun Mexico

Stejskal VDM et al.: Lymphocyte transformation test for diagnosis of isothiazolinone allergy in man. J Invest Dermatol 94 (1990) 798-802

Sanders B: The role of general and metal-specific cellular responses in protection and repair of metal-induced damage: stress proteins and metallothioneins. In: Chang L (ed.): Toxicology of Metals. USA: Lewis Publishers CRC Press Inc. (1996) 835-852