



LEARNING TOXICOLOGY THROUGH OPEN EDUCATIONAL RESOURCES

Chromium

Marie Vopršalová

Department of Pharmacology and Toxicology
Faculty of Pharmacy in Hradec Králové, Charles University,
Heyrovského 1203, 500 05 Hradec Králové, Czech Republic

e-mail: marie.voprsalova@faf.cuni.cz



Erasmus+

This work is licensed under a Creative
commons attribution – non commercial 4.0
international license



CHROMIUM (*Lat. Chromium, Cr*)

= durable metal

Cr occurs naturally in the Earth's crust, predominantly in the trivalent Cr^{3+} form.

It is ubiquitous in air, water, soil and biological materials.

In foodstuffs, Cr is generally considered to be present as Cr^{3+}

Cr^{6+} compounds are essentially anthropogenically produced and do not occur naturally in the environment.

In biological systems, the oxidation of Cr^{3+} to Cr^{6+} never occurs.

The toxicity of chromium depends on the oxidation state:

Cr^{6+} compounds (chromic acid, potassium and ammonium dichromates) are more toxic than Cr^{3+} .

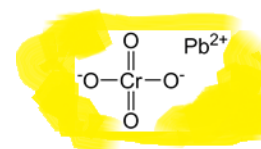
Cr^{3+} is considered an essential element in humans (is a constituent of the glucose tolerance factor, acts as a cofactor in insulin function), is required in glucose and lipid metabolism.

The daily requirement for Cr in humans is approximately 60 μg .

Elemental Cr is nontoxic.

1. Sources and uses:

- ❖ Manufacturing batteries, stainless steel
- ❖ Electroplating other metals to increase hardness and corrosion resistance
- ❖ Paint pigments: chrome yellow = lead chromate = PbCrO_4
- ❖ Wood preservatives
- ❖ Leather tanning: potassium dichromate = $\text{K}_2\text{Cr}_2\text{O}_7$
- ❖ Combustion of coal and oil
- ❖ Cement works
- ❖ Cigarette smoke



2. Fate in the organism:

Absorption:

Cr^{6+} is more efficiently absorbed in the lungs, GIT and through the skin than Cr^{3+} . Cr^{3+} is poorly absorbed from the GIT. Inhaled chromium particles can remain in the lung for a long time. Water soluble Cr^{3+} salts penetrate the skin but they don't reach the systemic circulation.

Distribution:

Cr^{3+} : in the blood, 95% is bound to large molecular mass proteins (transferrin).

Cr^{6+} : greater tendency to cross membranes, barriers → greater distribution to all tissues.

Highest Cr concentrations: in the kidney, liver and brain.

Cr^{6+} : unstable in the body reduction to more stable Cr^{3+} (by endogenous reducing agents, e.g. ascorbate and glutathione).

Excretion:

Cr is mainly excreted by the kidneys (about 80% of a chromium dose), elimination through the intestine and in breast milk may also occur.

3. Mechanism of toxicity:

Chromium toxicity results from its ability to penetrate cell membranes, inciting events resulting in cell death.

Cr^{6+} compounds are more toxic:

- ❖ oxidizing agents (→ corrosive to the mucous membranes in airways, GIT and skin)
- ❖ generation of free radicals → cellular damage
- ❖ depletes cellular antioxidants

4. Intoxication:

Cr^{3+} : Chromium dietary supplement (chromium picolinate) ingestions are generally nontoxic

Cr^{6+} : is associated with widespread organ toxicity.

Acute toxicity:

The lethal dose in humans is 2g $K_2Cr_2O_7$ by mouth.

$Na_2Cr_2O_7$ is also extremely toxic, the estimated oral lethal dose is between 1 to 10 g in an adult.

Corrosive nature can cause serious damage to mucous membranes of the respiratory and the gastrointestinal tract and the skin:

- ❖ **Ingestion** → esophageal and gastric necrosis, renal tubular necrosis
- ❖ **Inhalation** → upper respiratory tract irritation, nasal septum ulceration and perforation, bronchitis, pulmonary edema
- ❖ **Eye** → corneal injury
- ❖ **Skin** → dermal ulcers (known as „chrome holes“) skin burns may enhance systemic absorption



For more see:

<http://www.cdc.gov/niosh/topics/skin/occderm-slides/ocderm8.html> - slide 38

Chronic toxicity:

Cr^{6+} Cr^{3+} → allergic responses in sensitized individuals (e.g. asthma and contact dermatitis)

e.g. in construction workers – Cr containing cement



Blackjack disease = contact dermatitis

in card players or gamblers caused by exposure to chromium salts used for dyeing green felt or baize, which covers the gambling tables



Cr^{6+} → human carcinogen (lung cancer, cancer of the larynx, nasal cavity and sinuses). Workers who have been exposed to Cr fumes: an increased incidence of pulmonary cancer, cancer of the larynx, nasal cavity and sinuses with a latent period of about 20 years.



5. Laboratory determination:

Detection in the urine may confirm exposure.

Normal serum chromium concentration is $1\mu\text{g/l}$.

Normal urine levels are less than $40\mu\text{g/l}$.

6. Treatment: specific drugs and antidotes

- ❖ Skin contamination with soluble Cr salts: the area should be immediately rinsed with water, and CaNa_2EDTA in polyglycol solution.
- ❖ Cr^{6+} can be transformed into less toxic and poorly absorbed Cr^{3+} by administering ascorbic acid (oral doses of 2 to 4g per 1g of ingested chromium have been recommended if there are no symptoms of severe gastroesophageal injury). Larger doses could be harmful because ascorbic acid is a metabolic precursor of oxalate that can cause nephropathy.
- ❖ DMPS is recommended as an antidote for Cr poisoning.
Chelation therapy with BAL is not effective.



Share video:

<https://www.youtube.com/watch?v=fXF69p2UkVw>



Additional reading:

<http://www.dovemed.com/diseases-conditions/chromium-toxicity/>



References:

- Sun H, Brocato J, Costa M.: Oral chromium exposure and toxicity. *Curr Environ Health Rep.* 2015, 2(3), 295-303
- Klaassen, C D.: Casarett and Doull's toxicology: The Basic Science of Poisons, 7th ed., McGraw-Hill: New York, 2008, 931-980
- Shannon, M.W., Borron, S.W., Burns, M. J.: Haddad and Winchester's Clinical Management of Poisoning and Drug Overdose, 4th ed., Saunders/Elsevier: Philadelphia, 2007, 1111-1170
- Bryson, P.D.: Comprehensive Review in Toxicology for Emergency Clinicians, 3rd edition, Taylor and Francis: London, 1997, 579-642
- Olson, K. R. at al.: Poisoning & Drug Overdose, 5th Edition, McGraw-Hill, New York, 2006, 139-140
- Reichel, F-X., Ritter, L.: Illustrated Handbook of Toxicology, 4th edition. Thieme, Stuttgart, 2011, 160-182
- Timbrell, J.: The Poison Paradox: Chemicals as Friends and Foes, 1st edition, Oxford University Press, New York, 2005, 348



**VNiVERSiDAD
D SALAMANCA**

CAMPUS OF INTERNATIONAL EXCELLENCE



ALMA MATER STUDIORUM
UNIVERSITA DI BOLOGNA



South-Eastern Finland
University of Applied Sciences



UNIVERZITA
KARLOVA



Universitatea
TRANSILVANIA
din Braşov



<https://toxoeer.com>

Project coordinator: Ana I. Morales
Headquarters office in Salamanca.
Dept. Building, Campus Miguel de Unamuno, 37007.
Contact Phone: +34 663 056 665