

Sources of Exposure

Toxicokinetics and Normal Human Levels

Biomarkers/Environmental Levels

ToxGuide™

for

Nitrate (NO₃⁻)

CAS# 14797-55-8

and

Nitrite (NO₂⁻)

CAS# 14797-65-0

September 2015

U.S. Department of Health and
Human Services
Public Health Service
Agency for Toxic Substances
and Disease Registry
www.atsdr.cdc.gov

Contact Information:
Division of Toxicology
and Human Health Sciences
Environmental Toxicology Branch

1600 Clifton Road NE, F-57
Atlanta, GA 30329-4027
1-800-CDC-INFO
1-800-232-4636



General Populations

- Most dietary intake of nitrate comes from the consumption of certain vegetables; those particularly high in nitrate include celery, lettuce, and spinach.
- Selected meats and meat products contain sodium nitrate and/or sodium nitrite as preservatives and can be a source of overexposure to sodium nitrate and/or sodium nitrite.
- Drinking water from water supplies contaminated by nitrate from fertilizer runoff, animal waste, and/or nitrate-containing substances at waste disposal sites may result in overexposure to nitrate.
- Release of nitrate and/or nitrite to soil from fertilizer runoff, animal waste, and/or waste disposal sites could result in increased uptake by plants used for human consumption.

Occupational Populations

- Workers who are employed in occupations where fertilizer use is common (e.g., farming, greenhouse operations) may be exposed to nitrate and nitrite through dermal routes and inhalation of dust particles.

Toxicokinetics

- There is no information regarding the toxicokinetics of nitrate or nitrite following inhalation or dermal exposure.
- Gastrointestinal absorption of nitrate and nitrite occurs after drinking water or eating food that contain these compounds.
- Some of the nitrate in your body moves from blood to the salivary glands where some of it is changed to nitrite. Both nitrate and nitrite can be produced inside the body.
- Nitrate and nitrite are widely distributed in the body.
- Most of the nitrate that you take into your body each day leaves in the urine the same day it enters your body.
- Some nitrite in the stomach forms other substances called nitrosamines, some of which may be harmful.

Normal Human Levels

- National surveys of the U.S. general population conducted in 2005-2008 showed that the geometric mean level of nitrate in urine of nearly 18,000 participants was 42.7 or 46.3 mg/L.

Biomarkers

- There are no biomarkers specific to nitrate or nitrite exposure because these substances are also produced in the body. Methemoglobinemia has been used as a biomarker of nitrate and nitrite toxicity; however this effect is not specific to nitrate and nitrite.

Environmental Levels

Air

- Nitrate has been reported at up to a few parts per billion in air.

Sediment and Soil

- Nitrate levels in humid temperate U.S. soils typically range from 25 to 150 kg/hectare (as nitrogen), although much higher levels have been reported.

Water

- Naturally occurring levels of nitrate in streams and ground water in the U.S. were estimated at 0.24-1.0 mg/L; anthropogenic activity may result in much higher levels

Reference

Agency for Toxic Substances and Disease Registry (ATSDR). 2015. Toxicological Profile for Nitrate and Nitrite (Draft for public comment). Atlanta, GA: U.S. Department of Health and Human Services, Public Health Service.

NO₃⁻/NO₂⁻

Chemical and Physical Information

- Nitrate and nitrite are chemicals that combine with other chemicals such as sodium or potassium; these combinations are known as salts.
- Nitrate and nitrite salts are highly water-soluble and completely dissociate in aqueous environments.
- Nitrate and nitrite are part of the earth's nitrogen cycle whereby some nitrogen gas in air is converted to nitrite and nitrate, which can be used by plants and animals and returned to the air as nitrogen gas.

Routes of Exposure

- Inhalation exposure is not a likely route of concern for the general population.
- Oral exposure is the dominant route of exposure for the general population through ingestion of a normal diet. Over-exposure is possible by consumption of nitrate-rich diets and/or nitrate-contaminated drinking water.
- Dermal exposure is not a likely route of concern for the general population.

Nitrate and nitrite in the Environment

- Human-made and natural sources contribute to nitrate and nitrite aerosols in the atmosphere.
- Nitrite combines readily with oxygen to form nitrate. Nitrate is generally stable in the environment; however, it may be changed to nitrite through biological processes involving plants, microbes, etc.
- In nature, nitrate and nitrite can be found in igneous and volcanic rocks.
- Nitrogen exists naturally in soils, typically bound to organic matter and mineral soil material.

Relevance to Public Health (Health Effects)

Health effects are determined by the dose (how much), the duration (how long), and the route of exposure.

Minimal Risk Levels (MRLs)

Inhalation

- No acute-, intermediate- or chronic duration inhalation MRLs were derived for nitrate or nitrite

Oral

- MRLs of 4 mg/kg/day have been derived for acute-, intermediate-, and chronic duration oral exposure to nitrate.
- MRLs of 0.1 mg/kg/day have been derived for acute-, intermediate-, and chronic-duration oral exposure to nitrite.

Health Effects

- Exposure to high levels of nitrite can cause methemoglobinemia, a change to hemoglobin that decreases its ability to transport oxygen to tissues and related symptoms such as decreased blood pressure, increased heart rate, headaches, abdominal cramps, vomiting, and even death.
- The International Agency for Research on Cancer (IARC) noted that the presence of nitrite and some types of amines or amides in the acid environment of the stomach may result in the production of some cancer-causing N-nitroso compounds; under these conditions, IARC determined that ingested nitrate and nitrite is probably carcinogenic to humans. The EPA has not classified nitrate or nitrite for carcinogenicity.

Children's Health

- Children can experience the same effects as adults from overexposure to nitrate or nitrite.
- Young infants (<6 months of age) appeared to be particularly sensitive to the effects of nitrite on hemoglobin after consuming formula prepared with drinking water that contained nitrate at levels higher than recommended limits; some of these infants died.
- It is not known whether nitrate or nitrite can cause birth defects.