

Nitrous Oxide (Laughing Gas)



Background





N₂O is a colorless gas with a variety of diverse uses:

- Mild anesthetic in surgery and dentistry
- Recreational drug
- Oxidant in rocket and vehicle engines (NOS)
- Aerosol propellant used to dispense whipped cream as a foam





Safety

- N₂O is generally non-toxic, but it is still a compressed, liquefied gas
- Major hazards include asphyxia, mental and physical impairment, and vitamin B₁₂ deficiency in cases of chronic abuse

Historical Use

- First inhaled drug used for surgical anesthesia in 1844
- Analgesic use first reported by the British chemist Humphry Davy in 1798 while studying "medical airs"
- Popular among British aristocracy as early as 1799



Pharmacology and Mechanism of Action

- N₂O has a broad range of pharmacological effects, many of which are not well understood
- Clinically relevant effects include:

Opioid Receptor Agonist (Analgesic)

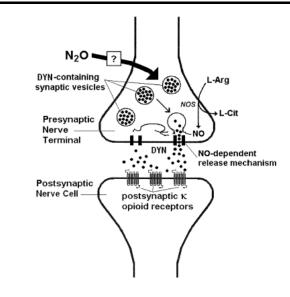
- N₂O-induced analgesia is antagonized by naloxone
- Cross tolerance observed between N₂O and morphine
- N₂O mediates release of endogenous opioid peptides

GABA/Benzodiazepine Receptor Agonist (Anxiolytic)

- Cross tolerance observed between N₂O and benzodiazepines
- Molecular mechanism unknown, potentially analogous to opioid agonism

NMDA Antagonist / Ligand-Gated Ion Channel Binding (Anesthetic)

- Molecular mechanism of general anesthesia is hotly debated
- N₂O up-regulates NMDA ligand in the cerebral cortex



Emmanouil, D. E.; Quock, R. M. Anesth Prog. 2007, 54, 9. https://doi.org/10.2344/0003-3006(2007)54[9:AIUTAO]2.0.CO;2



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Industrial Production and Use

N₂O is commercially synthesized through controlled pyrolysis of ammonium nitrate

$$NH_4NO_3 = 270 \degree C$$
 $2 H_2O + N_2O$

N₂O is used in the industrial synthesis of azide salts as well

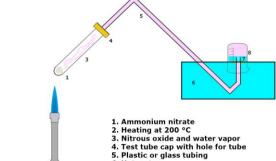
$$2 \text{ NaNH}_2 + \text{N}_2\text{O}$$
 \longrightarrow $\text{NaN}_3 + \text{NaOH} + \text{NH}_3$

Alternate synthesis at lower temperature:

$$2 (NH_2)_2CO + 2 HNO_3 + H_2SO_4$$
 \rightarrow $2 N_2O + 2 CO_2 + (NH_4)_2SO_4 + 2 H_2O$

Volkova, A. V. Russ. J. Gen. Chem. 2019, 89, 1338. https://doi.org/10.1134/S107036321906032X

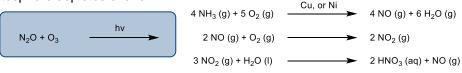
Laboratory Production



- 6. Hot water 7. Plastic or metal with hole to hold tubing
- 8. Pure nitrous oxide gas

N₂O in the stratosphere depletes ozone

Nitrous Oxide as a Pollutant



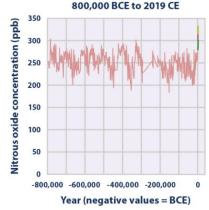
 $2 \text{ NH}_3 (g) + 4 \text{ O}_2 (g) + \text{H}_2 \text{O} (l)$ $3 H_2O (g) + 2 HNO_3 (aq)$ Overall:

The Ostwald Process

cat. Pt.

- N₂O is roughly 300x more potent as a greenhouse gas than CO₂
- Since it's less abundant, it's only the third largest contributor to atmospheric warming
- The majority of anthropogenic nitrous oxide emissions are from the use of fertilizers in agriculture

U.S. Environmental Protection Agency (2010), "Methane and Nitrous Oxide Emissions from Natural Sources". Report EPA 430-R-10-001.



N₂O is a waste byproduct of industrial nitric acid production

