

Dr Tim Chambers

Nitrates – Are they an emerging risk?

University of Otago





Team and funding



































Nitrate contamination in New Zealand

Nitrate in New Zealand

70% increase in dairy cattle (3.8 million to 6.5 million 1994-2017)

Six-fold increase in the use of nitrogen fertilisers (59,000 tonnes to 429,000 tonnes)

Area under irrigation increased 91% from 2002 to 2019

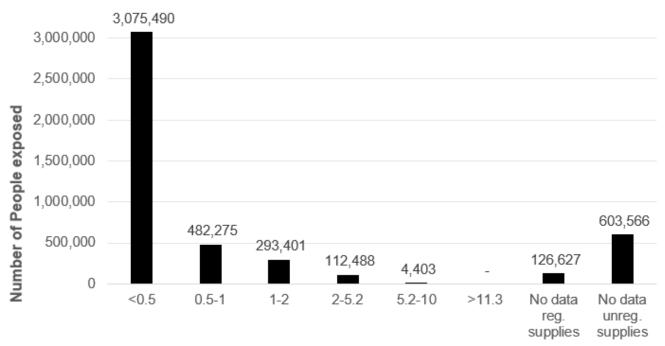






Nitrate in drinking water

2020 – registered supplies

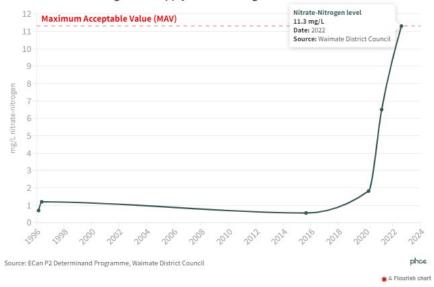


Nitrate Concentration Range (as mg/L NO3-N)

Registered supplies

Contamination events

Lower Waihao Drinking water supply nitrate-nitrogen



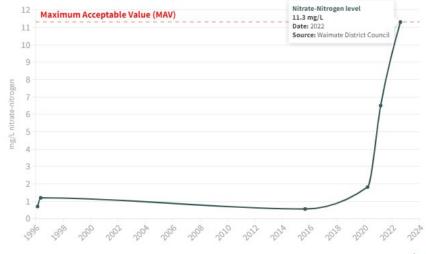
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Registered supplies

Contamination events

Lower Waihao Drinking water supply nitrate-nitrogen

Source: ECan P2 Determinand Programme, Waimate District Council



Water supplies	Population Served	Current NO ₃ -N	5 th to 95 th percentile (mg/L)
Christchurch	342,000	0.7	(3.0- 7.6)
Waimakariri	53,800	1.9	(3.0-8.3)

Unregistered supplies













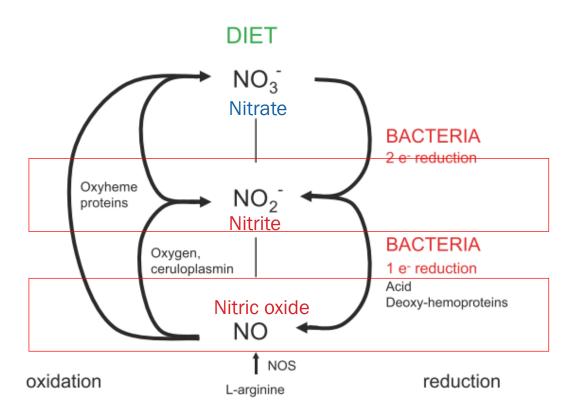
Whangarei





Nitrate and health

Nitrate metabolism



Current nitrate limit

WHO set nitrate limit 11.3 mg/L nitrate-nitrogen (50mg/L as nitrate) based on risk of Blue Baby Syndrome in 1950

Same limit adopted in NZ drinking water standards



Nitrate is reduced to nitrite

nitrite
binds to haemoglobin

formation methaemoglobin
less oxygen carried in blood
+ oxidative stress

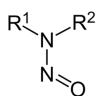
Methaemoglobinemia

Nitrate and preterm births

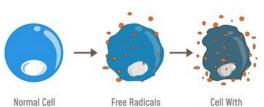
Births <37 weeks

Leading cause of infant mortality (23% of deaths)

N-nitroso compounds (NOC) OXIDATIVE STRESS e.g. N-nitrosamines

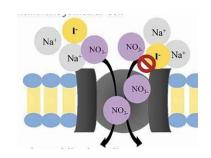






Attacking Cell

Thyroid dysfunction

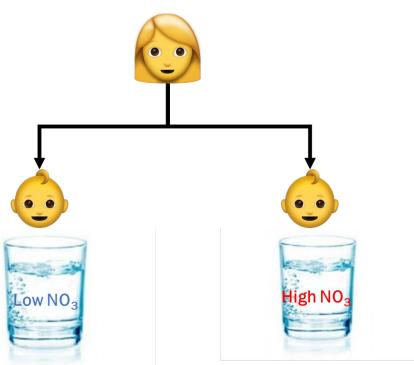


Nitrate and preterm births

Stanford Study (2022)

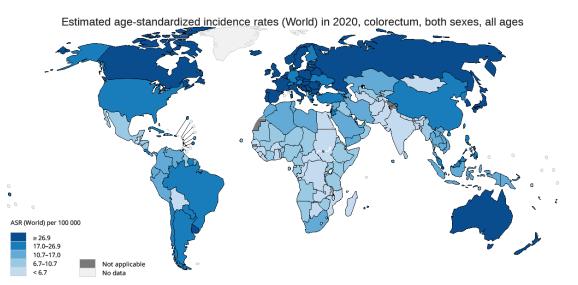
4.6 million births from 2000-2011

 47% increase in risk of preterm birth (<32 weeks) where nitrate in drinking water 5 to 10 mg/L NO3-N compared to <5mg/L



Bowel cancer

Each year approximately 1,200 people in NZ die of bowel

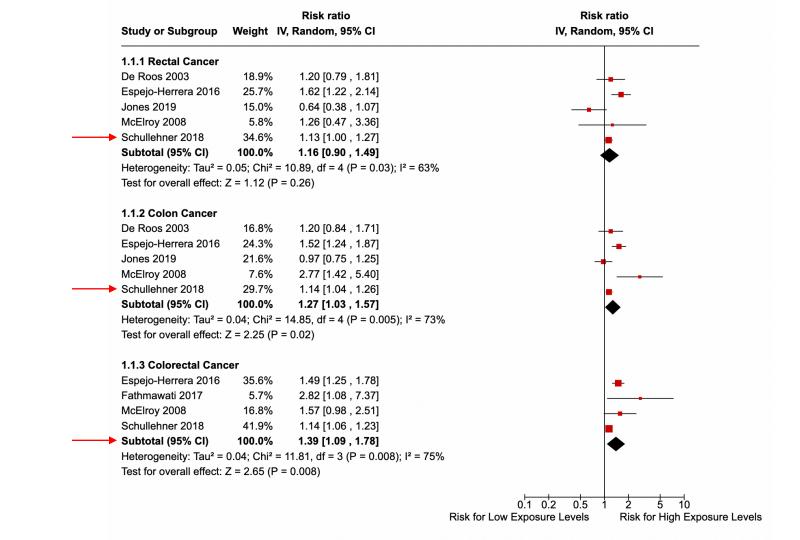




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Data source: GLOBOCAN 2020 Map production: IARC (http://gco.iarc.fr/today) World Health Organization





Agency responses

International Agency for Research on Cancer (IARC, 2010)

Studies up until 2006

"Ingested nitrate or nitrite under conditions that result in endogenous nitrosation is **probably** carcinogenic to humans (group 2A)"

WORLD HEALTH ORGANIZATION
INTERNATIONAL AGENCY FOR RESEARCH ON CANCER



IARC Monographs on the Evaluation of Carcinogenic Risks to Humans

VOLUME 94

Ingested Nitrate and Nitrite, and Cyanobacterial Peptide Toxins

20, AVENUE APPIA - CH-1211 GENEVA 27 - SWITZERLAND - TEL CENTRAL +41 22 791 2111 - FAX CENTRAL +41 22 791 3111 - WWW.WHO.INT

Agency responses

World Health Organization (2019)

Tel. direct:

t: +41 22 791 1540 defrancei@who.int

In reply please

Your reference:

Dr Caroline McElnay Director of Public Health Ministry of Health New Zealand 133 Molesworth Street PO Box 5013 Wellington 6140 New Zealand

"There is a biological plausibility of endogenous nitrosation of ingested nitrate and nitrite which **could potentially** cause cancer in humans"

However

"the weight of evidence does not support an association between cancer and exposure to nitrate or nitrite **per se**."

Agency responses

French Agency for Food, Environmental and Occupational Health & Safety (2022)

"There is an association between the risk of bowel cancer and exposure to nitrites and/or nitrates, whether they are ingested via the consumption of processed meat or drinking water."

And recommended

"assessing, in light of new epidemiological and toxicological data, the relevance of the quality limit of 11.3 mg L of nitrates in drinking water."



Uncertainties

- Conversion of nitrate to nitrite (5-20% interpersonal variation)
- Other dietary sources of nitrite/nitrate (processed meats)
- Impact of antioxidants
- Not all NOC are carcinogens
- And much more...

Treatment options

Reverse osmosis and ion exchange

Registered supplies = \$750,000 (Waimate), \$6M (Selwyn), \$610M (Christchurch)

Unregistered supplies = low-end \$500-\$1000 to \$10,000s

Nitrates – Are they an emerging risk YES

- Increasing levels of contamination in environmental monitoring sites, registered and unregistered supplies
- Uncertainties around potential adverse health outcomes
- Cost of treatment and groundwater lags











Current work

Outcome data: IDI back to 2008 – (700,000 births)

Exposure: Regional, district and city council data + modelling + community nitrate testing

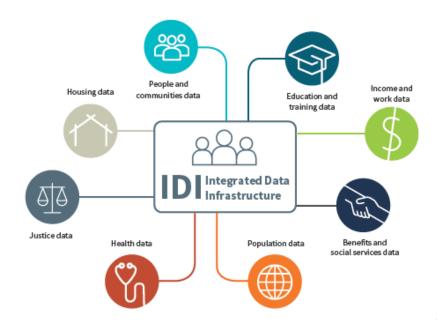


S Stuff.co.nz

Nitrate risk to unborn babies to be measured in \$1.2 million

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The results found the risk of early preterm birth (where an infant is born at least nine weeks early) increased by half when nitrates in...





Thank you tim.chambers@Otago.ac.nz

N-Nitroso Compounds

Form N-Nitroso Compounds – "NOC" (90% tested are known carcinogens)

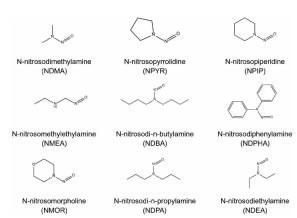
Proteins + nitrosating agents (e.g. nitrite + nitric oxide)

Antioxidants (Vitamin C, Green tea)

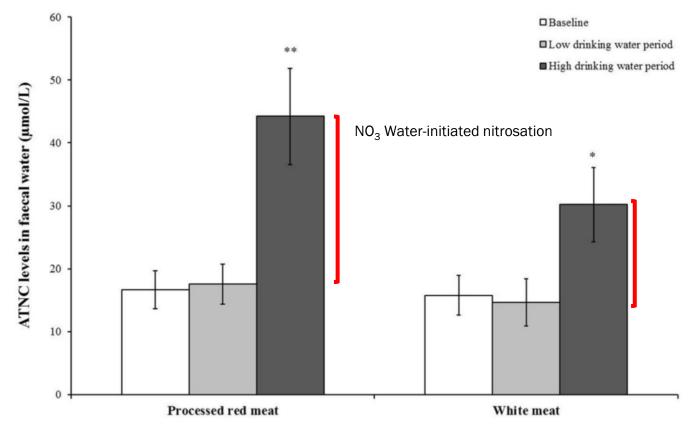












van Breda SG, Mathijs K, Sági-Kiss V, Kuhnle GG, Van der Veer B, Jones RR *et al.* Impact of high drinking water nitrate levels on the endogenous formation of apparent N-nitroso compounds in combination with meat intake in healthy volunteers. *Environmental Health.* 2019;18(1):87. https://doi.org/10.1186/s12940-019-0525-z

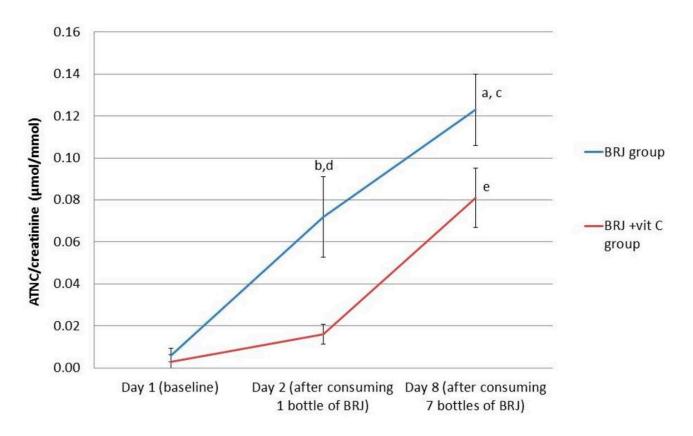


Figure from Randomised control trial by Berends 2019 – 1000mg Vitamin C In NZ - 99mg Vitamin C per day (NZ Nutrition Survey 2008)

Nitrate and bowel cancer



Contents lists available at ScienceDirect

Public Health





Original Research

Do low-fat foods alter risk of colorectal cancer from processed meat?



M.J. Sneyd ¹, B. Cox*

High Adam Cancer Epidemiology Unit, Department of Preventive and Social Medicine, Dunedin School of Medicine, University of Orago, PO Box 56,

Hugh Adam Cancer Epidemiology Unit, Department of Preventive and Social Medicine, Dunedin School of Medicine, University of Otago, PO Box 5 Dunedin, 9054, New Zealand

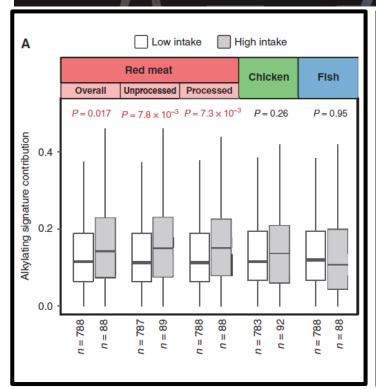
Established case-control study of ~800 bowel cancer cases

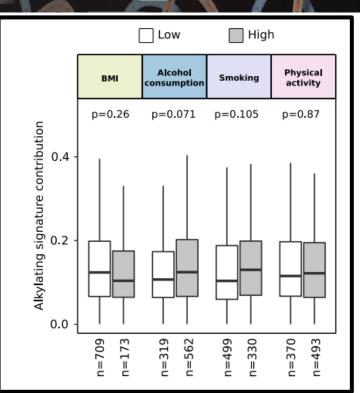
Test the association between nitrate and bowel cancer, adjusting for established risk factors including alcohol and tobacco use, diet, body mass index and physical activity.

Biopsies of cases are still available and could be re-analysed to investigate whether nitrate exposure was associated with a specific type of DNA damage

RESEARCH BRIEF

Discovery and Features of an Alkylating Signature in Colorectal Cancer





Nitrate and preterm births

Meta-analysis 2023 Lin

