Real or Not?

2024

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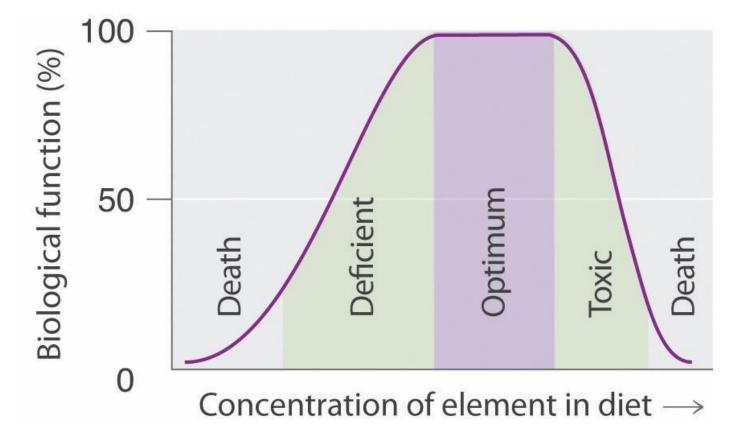
What Is the Idea of Hormesis?

From Saha, p. 17

- A little bit of a bad thing can be good
- What are some examples?

Selenium: Helps or Hurts?

https://saylordotorg.github.io/text_general-chemistry-principles-patterns-and-applications-v1.0/



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Selenium: Helps or Hurts?

K. L. Nuttall. Annals of Clinical & Laboratory Science, vol. 36, no. 4, 2006

Table 3. Published reports discussed in text (with emphasis on serum selenium)

| Report | Age & Gender (yr, M/F) | Dosing pattern | Serum Se (µg/L) | Notes |
|--------|---------------------------|------------------|-----------------------|---------------------------------|
| 3 | 24 M | acute | 30000 | ingestion - 1 hr, fatal - 4 hr |
| 7 | 52 F | acute | 2435 | ingestion at 4 d, fatal at 6 d |
| 11 | 15 F | acute | 3100 | ingestion - 1 hr |
| 12 | 56 M | acute | 2716 | ingestion - 3 hr |
| 13 | 48 F | acute | 2400 | ingestion - 2 hr |
| 14 | 2 F | acute | 1580 ^a | ingestion - 5 hr |
| 15 | 29 M | acute | 931 ^a | ingestion - 3 hr |
| 16 | 2 M | acute | 420 | ingestion at 1 d, fatal at 18 d |
| 17 | adults | chronic | 650-1400 ^b | no toxicity in 54 of 86 |
| 18 | 36 M | chronic | 680 | vitamin therapy for fatigue |
| 19 | 57 F | chronic | 528 | nutritional supplement |
| 20 | adults | 1 - 1 | 123-363° | without toxicity |
| 21 | ≥14 M | - | 101-151 ^d | NHANES III reference interva |

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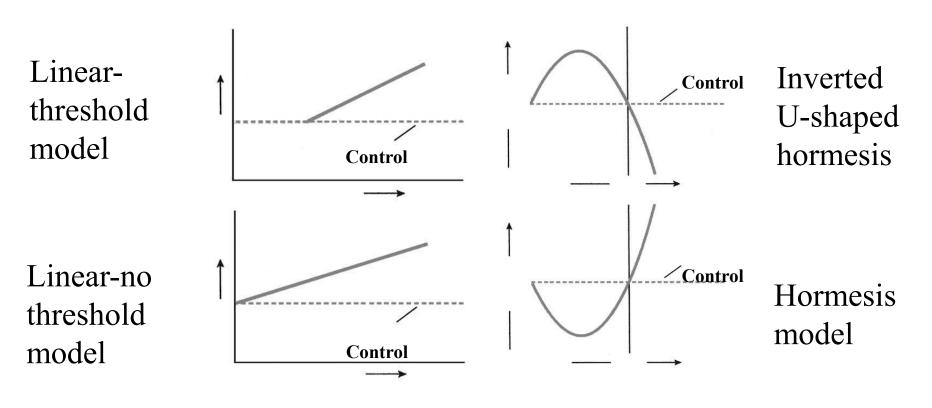
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Arguments For/Against Selenium Hormesis

- Taking megadoses of 200 mcg/day (4x the RDA) of selenium may have acute toxic effects, and showed no decreased incidence of prostate cancer mortality for low doses, and increased high-grade prostate cancer rates (~35,533 men)
 - Refs: Brasky TM, Kristal AR. J Natl Cancer Inst (2015) 107(1): dju375, Kenfield SA, et al. *J Natl Cancer Inst* (2015) 107 (1): dju360
- Selenium supplements of 200 mcg/day (4x the RDA) greatly reduced (63%) secondary prostate cancer evolution (974 men, 13 Se vs. 35 placebo cases, 11 years follow-up)
 - Ref: L. C. Clark et al. British Journal of Urology [1998, 81(5):730-734]

Models of Dose-Response

D. P. Hayes. European Journal of Clinical Nutrition (2007) 61, 147–159

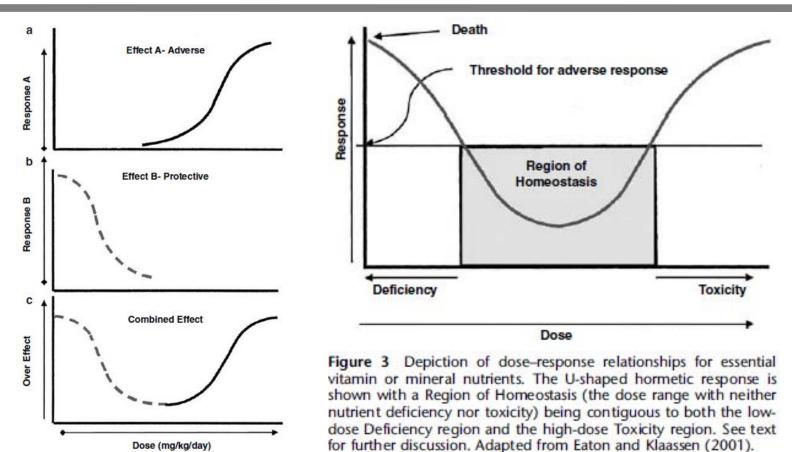


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Is Radiation Hormetic?



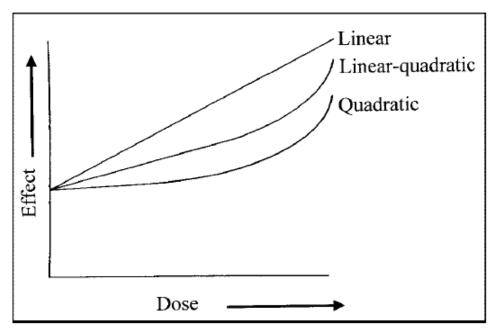


Courtesy of Macmillan Publishers Ltd.

Dose (mg/kg/day)

History of Non-Hormetic Models

J. L. Prekeges, J. Nucl. Med. Technol., 31:11-17 (2003)



Original image source: Mettler, F.A., and A.C. Upton. *Medical Effects of Ionizing Radiation*. 2nd ed. W.B. Saunders, 1995.

FIGURE 1. Three hypotheses of radiation dose-response relationship, as proposed by the United Nations Scientific Committee on the Effects of Atomic Radiation in 1958. (Reprinted with permission of (3).)

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History: The LNT Model

J. L. Prekeges, J. Nucl. Med. Technol., 31:11-17 (2003)

Linear No-Threshold model states that:

- Radiation exposure is harmful;
- Radiation exposure is harmful at all exposure levels;
- Each increment of exposure adds to the overall risk;
- The rate of accumulation of radiation exposure has no bearing on risk.

Checking the LNT Model

J. L. Prekeges, J. Nucl. Med. Technol., 31:11-17 (2003)

Linear No-Threshold model states that:

• Radiation exposure is harmful;

We know this from LOTS of accidental exposure data

- Radiation exposure is harmful at all exposure levels;
- Each increment of exposure adds to the overall risk;
- The rate of accumulation of radiation exposure has no bearing on risk.

Checking the LNT Model

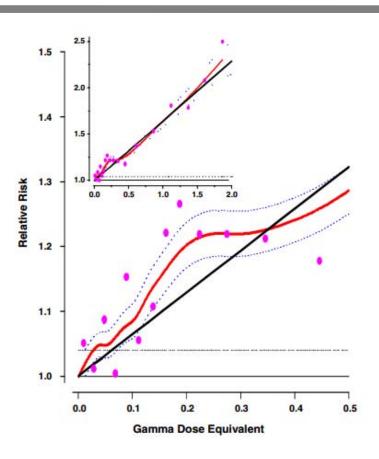
J. L. Prekeges, J. Nucl. Med. Technol., 31:11-17 (2003)

ICRP Publication 99, p. 32

Linear No-Threshold model states that:

- Radiation exposure is harmful;
- Radiation exposure is harmful at all exposure levels;
- Each increment of exposure adds to the overall risk;
- The rate of accumulation of radiation exposure has no bearing on risk.

The ICRP-99 study showed this to be false



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Checking the LNT Model

J. L. Prekeges, J. Nucl. Med. Technol., 31:11-17 (2003)

Linear No-Threshold model states that:

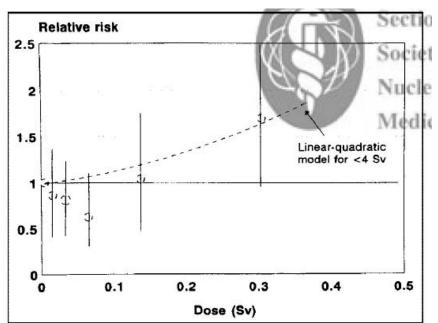
- Radiation exposure is harmful;
- Radiation exposure is harmful at all exposure levels;
- Each increment of exposure adds to the overall risk;

These are the open questions today!

• The rate of accumulation of radiation exposure has no bearing on risk.

Evidence for Hormesis

- Ask yourself:
 - What study size was used?
 - How were the error bars defined?
 - Can hormesis be conclusively shown from this study?



J. L. Prekeges, J. Nucl. Med. Technol., 31:11-17 (2003)

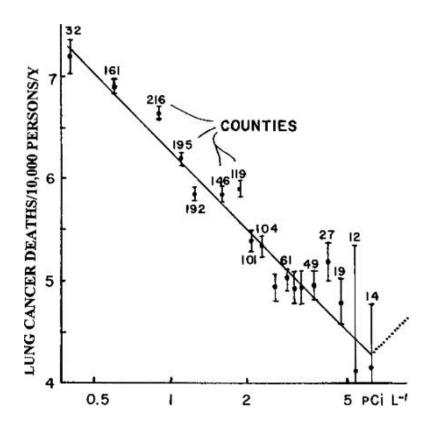
FIGURE 3. Incidence of leukemia as function of radiation dose in atomic bomb survivors in Japan. (Reprinted with permission of (3).)

Original image source: Mettler, F.A., and A.C. Upton. *Medical Effects of Ionizing Radiation*. 2nd ed. W.B. Saunders, 1995.

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Evidence for Hormesis

Cohen B.L. Health Phys. 1995;68:157–174



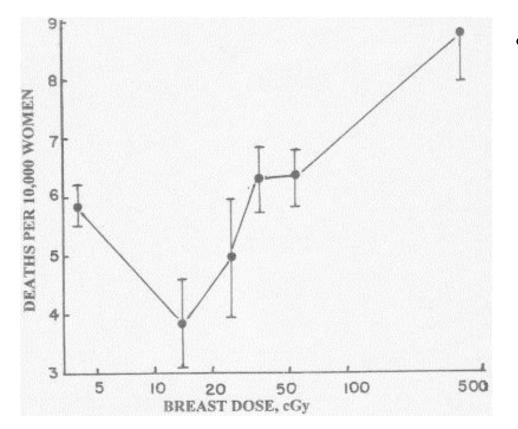
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- To a point, radon levels in homes were found to *decrease* lung cancer deaths
 - 200,000 radon samples
 - 1,600 counties, comprising 90% of the U.S. population

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Evidence for Hormesis

A. B Miller et al. New England J Med. 1989;321:11285–11288



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- Women who received
 multiple x-rays to monitor
 lung collapse during
 tuberculosis treatment
 experienced less breast
 cancer
 - One standard deviation displayed

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CAN Hormesis Be Real?

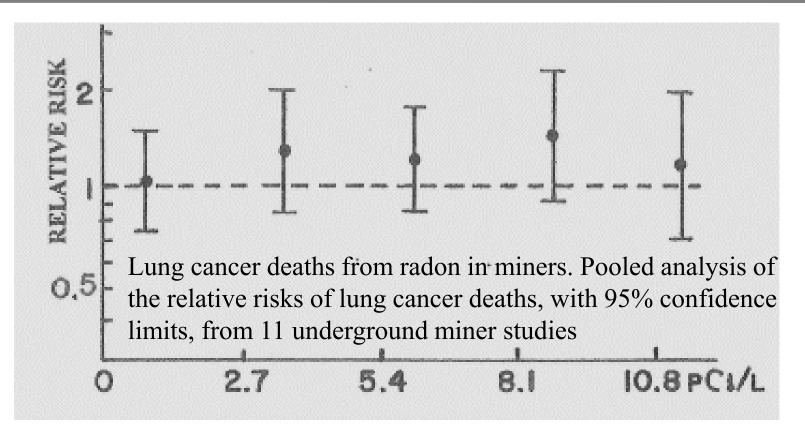
• Certainly at the cellular level

• Maybe not at the multicellular/organism level

• More data needed?

Evidence for a Threshold Model

Refer to BEIR VI Committee. Health Effects of Exposure to Radon. The National Academy Press; 1999.

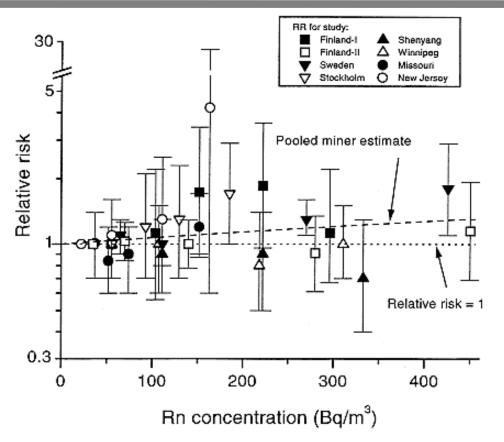


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Evidence for a Threshold Model

BEIR VI Committee . Health Effects of Exposure to Radon. The National Academy Press; 1999.



Lung cancer deaths from radon in homes. Eight case control studies of relative risks for lung cancer death rates of people in homes. One standard deviation shown.

Courtesy of National Academies Press. Used with permission. Source: National Research Council. *Health Effects of Exposure to Radon: BEIR VI*. The National Academies Press, 1999. doi:10.17226/5499.

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The 22.01 Hormesis Debate

Together, we will debate whether hormesis can actually exist

- 1. Split into two groups: Pro and Against
- Spend ~30 minutes to find evidence, papers, and construct arguments for/against hormesis
- 3. Present your findings in 10 minutes each, with 5 minutes each to rebut

22.01 Introduction to Nuclear Engineering and Ionizing Radiation Spring 2024

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