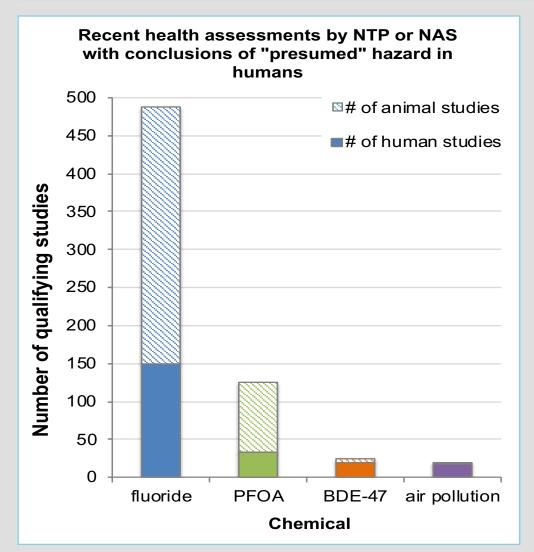
Fluoride Developmental Neurotoxicity:

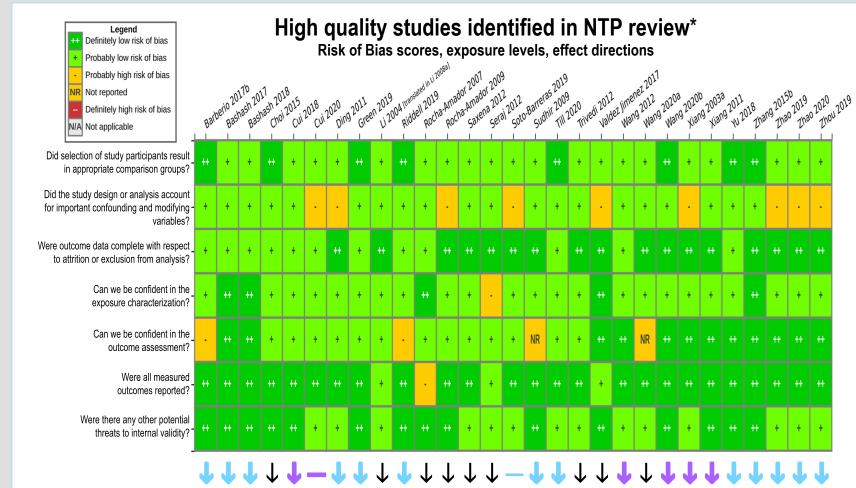
Dose-Response Analyses of Recent High Quality Studies

Chris Neurath^a, Paul Connett^b, Michael Connett^c, Bill Hirzy^b

^a American Environmental Health Studies Project, ^b Fluoride Action Network, ^c Waters Kraus & Paul

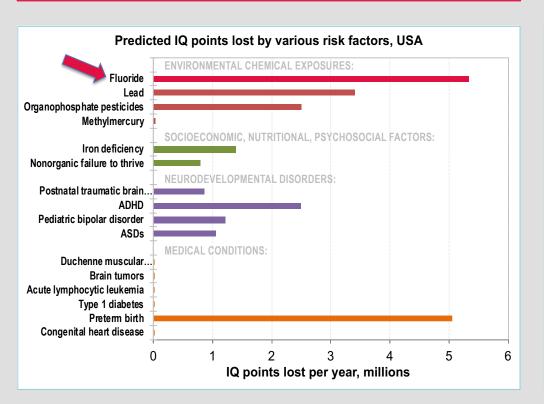






Of 29 high quality studies 27 found statistically significant adverse effects (♣), 2 found no effect (−), and none found beneficial effect (↑)

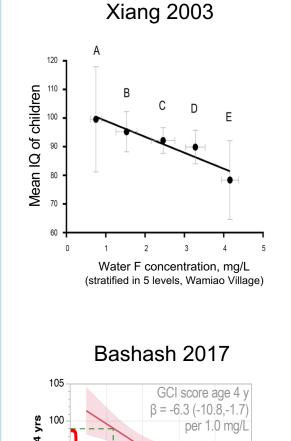
- **↓** = exposures below 1.5 mg/L water F or equivalent (5 studies)
- ↓ = exposures above 1.5 mg/L water F or equivalent (9 studies)

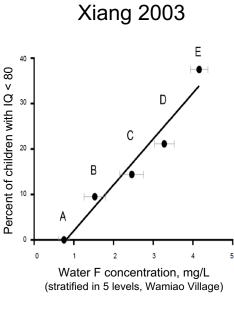


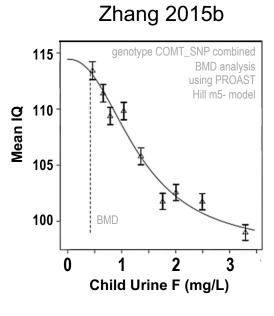
Dose-response analysis summaries Study Effect Magnitude β **BMDL Xiang 2003** -5.7 IQ / 1 mg/L water F 0.27 mg/d +10% IQ<80 / 1 mg/L water F **Xiang 2003** 0.16 mg/L Zhang 2015b **-9.7** IQ / 1 mg/L urine F -12.3 IQ / 1 mg/L urine F 0.00 mg/L Cui 2018 Bashash 2017 -6.3 IQ / 1 mg/L urine F 0.10 mg/L +5.7 pts / 1 mg/L urine F Bashash 2018 Green 2019 -4.5 IQ / 1 mg/L urine F 0.12 mg/L Till 2020 -8.8 IQ / 1 mg/L water F 0.06 mg/L 0.09 mg/L

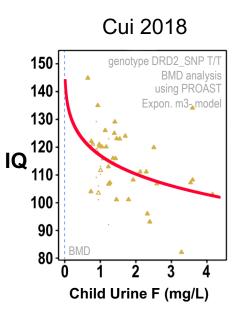
Examples of studies suitable for dose-response analyses

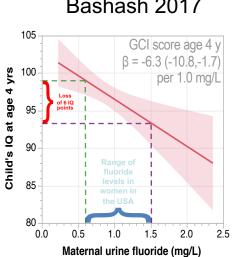
Dose-response curves and BMD analyses based on data or figures in each paper

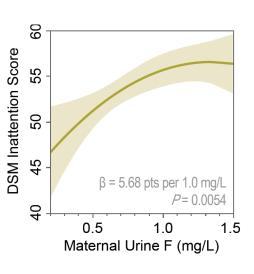




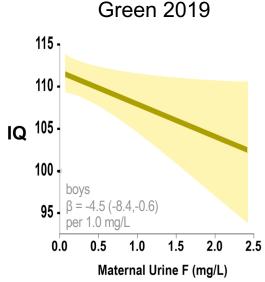


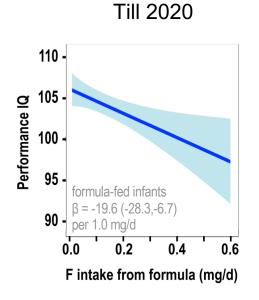






Bashash 2018





Notes

- 1.) Exposures measured as urine F concentrations are considered equivalent to drinking water F concentrations.
- 2.) Community water fluoridation concentration is typically 0.7 1.0 mg/L.
- 3.) For studies with multiple subpopulations, outcomes or exposure measures, the most sensitive significant association was chosen, consistent with standard risk assessment practice.
- 4.) Benchmark Dose analyses (BMD) used response (BMR) of –1 IQ point as adverse effect.
- 5.) No intra-species Uncertainty Factor (UF) applied to BMDLs.
- 6.) BMDLs for Xiang 2003 from Hirzy 2016; for Bashash 2017, Green 2019 from Grandjean 2019; for Zhang 2015b, Cui 2018, Till 2020 by Neurath using PROAST BMD software or linear dose-response method of Grandjean 2019.

References

*Adapted from NTP draft monograph data: https://hawcproject.org/assessment/405/ https://hawcproject.org/summary/visual/524/

Hirzy 2016

https://www.fluorideresearch.org/494Pt1/files/FJ 2016 v49 n4Pt1 p379-400 pq.pdf

Grandjean 2019

https://doi.org/10.1186/s12940-019-0551-x

PROAST BMD software:

https://proastweb.rivm.nl/

additional information:
http://fluoridealert.org/studies/neurath-powerpoint-developmental-neurotoxicity/