## NTP Board of Scientific Counselors Meeting December 10, 2014

Title: Assessing NTP's Effectiveness: A Case Study on Hexavalent Chromium

**NTP Scientist:** Mary S. Wolfe, PhD, Yun Xie, PhD, and Danica Andrews, Office of Liaison, Policy, and Review, Division of NTP; Stephanie Holmgren, MLIS, MBA, Office of Scientific Information Management, Office of the Deputy Director, NIEHS

## **Purpose**

For over 35 years, the National Toxicology Program (NTP) has conducted research and analysis activities and disseminated information about potential health hazards in our environment. We sought to assess the effectiveness of NTP's science at advancing toxicology and being translated to public health decision-making. A logic model was developed with defined inputs, outputs (activities and products), and outcomes (proximal, intermediate, distal) and applied retrospectively to NTP's research program on hexavalent chromium (Cr6) as a case study.

## **Background and Significance**

A number of studies, supported by public universities, private organizations, and federal agencies, have evaluated the impact of federally funded research [1–10]. In 2013, the National Institutes of Health (NIH) released a report on the approaches for assessing the value of biomedical research supported by NIH [11], and the National Academies of Science published a report related to measuring the impact of research on society [12]. The literature for assessing the value of research highlights the importance of measuring impact and the challenges involved, such as attribution (finding a connection from research to an impact), lag time (accounting for the potentially long period of time between research and impact), and external factors (research institutions lack direct control over how their work will be applied by federal agencies, industry, and the public) [2,3,7,9–12].

Cr6 was selected for the case study because NTP's work was completed several years earlier with presumably sufficient time to identify its use by stakeholders and evaluate impacts. NTP's products on Cr6 included 1 technical report, 1 toxicity report, 5 journal articles, 3 laboratory reports, and a listing in the *Report on Carcinogens* [13]. Proximal outcomes measured by Webtrends data and external requests showed stakeholders had immediate awareness of NTP's products. Intermediate outcomes of NTP's work to inform science were many. NTP's research on Cr6 was cited in scientific publications to justify further studies, inform study designs, or interpret new data. Stakeholders used NTP's work to identify Cr6 as a hazard in legal and policy documents, science reports, congressional testimonies, proposed legislations or regulations, and non-regulatory actions.

Pinpointing distal outcomes showing NTP's products led to a positive change for public health was challenging because NTP has no regulatory authority, the time lag to impact varies, and external factors may affect use or progress of intended actions. Although proposed federal laws or state regulations cited NTP's products, often they were not enacted or were delayed by external factors. Notably, NTP's research was key to the nation's first-ever drinking water standard for Cr6 adopted by California in 2014.

This case study demonstrated NTP's science on Cr6 had impact in many areas including public health. Furthermore, it identified broad and objective approaches for assessing NTP's effectiveness, along with data and methodological gaps that need to be addressed for more thorough and efficient assessments in the future.

## References

- 1. Battelle Technology Partnership Practice. 2011. Economic Impact of the Human Genome Project. College Park, MD: Battelle Memorial Institute.
- Liebow E, Phelps J, Van Houten B, Rose S, Orians C, Cohen J, Monroe P, and Drew CH. 2009. Toward the assessment of scientific and public health impacts of the National Institute of Environmental Health Sciences Extramural Asthma Research Program using available data. Environ Health Perspect, 117(7), 1147–54.
- 3. Chatterjee A, and DeVol RC. 2012. Estimating Long-Term Economic Returns of NIH Funding on Output in the Biosciences. Santa Monica, CA: Milken Institute.
- 4. Ehrlich EM. 2011. An Economic Engine: NIH Research, Employment, and the Future of the Medical Innovation Sector [online]. Washington, DC: United for Medical Research. Available: <a href="http://www.unitedformedicalresearch.com/wp-content/uploads/2012/07/UMR Economic-Engine.pdf">http://www.unitedformedicalresearch.com/wp-content/uploads/2012/07/UMR Economic-Engine.pdf</a> [accessed 20 October 2014].
- 5. Ehrlich EM. 2012. NIH's Role in Sustaining the U.S. Economy: A 2011 Update Authored by Dr. Everett Ehrlich [online]. Washington, DC: United for Medical Research. Available: <a href="http://www.unitedformedicalresearch.com/wp-content/uploads/2012/07/NIHs-Role-in-Sustaining-the-US-Economy-2011.pdf">http://www.unitedformedicalresearch.com/wp-content/uploads/2012/07/NIHs-Role-in-Sustaining-the-US-Economy-2011.pdf</a> [accessed 20 October 2014].
- 6. Families USA's Global Health Initiative. 2008. In Your Own Backyard: How NIH Funding Helps Your State's Economy. Washington, DC: Families USA.
- 7. Health Economics Research Group, Office of Health Economics, and RAND Europe. 2008. Medical Research: What's it Worth? Estimating the Economic Benefits from Medical Research in the UK. London: UK Evaluation Forum.
- 8. Jacob BA and Lefgren L. 2011. The Impact of Research Grant Funding on Scientific Productivity. J Public Econ, 95(9–10), 1168–77.

- 9. Toole AA. 2007. Does public scientific research complement private investment in research and development in the pharmaceutical industry? J Law Econ, 50(1), 81–104.
- 10. Toole AA. 2012. The impact of public basic research on industrial innovation: Evidence from the pharmaceutical industry. Research Policy, 41(1), 1–12.
- 11. National Institutes of Health. 2013. Scientific Management Review Board Report on the Approaches to Assess the Value of Biomedical Research. Bethesda, MD: National Institutes of Health. Available: <a href="http://smrb.od.nih.gov/documents/reports/VOBR%20SMRB">http://smrb.od.nih.gov/documents/reports/VOBR%20SMRB</a> Report 2014.pdf [accessed 20 October 2014].
- 12. National Research Council. 2014. Furthering America's Research Enterprise. Washington, DC: The National Academies Press. Available: http://www.nap.edu/catalog.php?record\_id=18804 [accessed 20 October 2014].
- 13. (A) National Toxicology Program. 2008. Toxicology and Carcinogenesis Studies of Sodium Dichromate Dihydrate (CAS No. 7789-12-0) in F344/N Rats and B6C3F1 Mice (Drinking Water Studies). NTP Technical Report 546. Research Triangle Park, NC: National Toxicology Program. Available: <a href="http://ntp.niehs.nih.gov/files/546">http://ntp.niehs.nih.gov/files/546</a> web FINAL.pdf [accessed 23 October 2014].
  - (B) Bucher, JR. 2007. NTP Technical Report on the Toxicity Studies of Sodium Dichromate Dihydrate (CAS No. 7789-12-0) Administered in Drinking Water to Male and Female F344/N Rats and B6C3F 1 Mice and Male BALB/c and am3-C57BL/6 Mice. NTP Toxicity Report Series Number 72. Research Triangle Park, NC: National Toxicology Program. Available: <a href="http://ntp.niehs.nih.gov/ntp/htdocs/ST">http://ntp.niehs.nih.gov/ntp/htdocs/ST</a> rpts/tox072.pdf [accessed 23 October 2014].
  - (C) Collins BJ, Stout MD, Levine KE, Kissling GE, Melnick RL, Fennell TR, Walden R, Abdo K, Pritchard JB, Fernando RA, Burka LT, and Hooth MJ. 2010. Exposure to hexavalent chromium resulted in significantly higher tissue chromium burden compared with trivalent chromium following similar oral doses to male F344/N rats and female B6C3F1 mice. Toxicol Sci, 118(2), 368–79.
  - (D) Levine KE, Stout MD, Ross GT, Essader AS, Perlmutter JM, Grohse PM, Fernando RA, Lang M, and Collins BJ. 2009. Validation of a method for the determination of total chromium in rat feces by inductively coupled plasma optical emission spectrometry. Anal. Lett, 42, 2729–2746.
  - (E) Levine KE, Stout MD, Ross GT, Essader AS, Weber FX, Grohse PM, Fernando RA, Milstein LS, Hooth MJ, and Collins BJ. 2010. Validation and application of a method for the determination of total chromium in rat tissues by inductively coupled plasma mass spectrometry. Arch Environ Contam Toxicol, 58(3), 883–91.
  - (F) Stout MD, Herbert RA, Kissling GE, Collins BJ, Travlos GS, Witt KL, Melnick RL, Abdo KM, Malarkey DE, and Hooth MJ. 2009. Hexavalent chromium is carcinogenic to F344/N

- rats and B6C3F1 mice after chronic oral exposure. Environ Health Perspect, 117(5), 716–22.
- (G) Witt KL, Stout MD, Herbert RA, Travlos GS, Kissling GE, Collins BJ, and Hooth MJ. 2013. Mechanistic insights from the NTP studies of chromium. Toxicol Pathol, 41(2), 326–42.
- (H) National Toxicology Program. 1996. Final report on the reproductive toxicity of potassium dichromate (hexavalent) (CAS No. 7778- 50-9) administered in diet to SD rats. Research Triangle Park, NC: National Toxicology Program.
- (I) National Toxicology Program. 1996. Final report on the reproductive toxicity of potassium dichromate (hexavalent) (CAS No. 7778- 50-9) administered in diet to BALB/c mice. Research Triangle Park, NC: National Toxicology Program.
- (J) National Toxicology Program. 1997. Final report on the reproductive toxicity of potassium dichromate (CAS No. 7778-50-9) administered in diet to BALB/c mice. Research Triangle Park, NC: National Toxicology Program.
- (K) National Toxicology Program. 2014. 13<sup>th</sup> Report on Carcinogens. Research Triangle Park, NC: National Toxicology Program. Available: <a href="http://ntp.niehs.nih.gov/go/roc13">http://ntp.niehs.nih.gov/go/roc13</a> [accessed 23 October 2014].