

The ECOTOXicology Knowledgebase Literature Search and Review Processes for Identifying and Curating Toxicity Data for Ecological Risk Assessments

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The views expressed in this presentation are those of the authors and do not necessarily reflect the views or policies of the U.S. EPA.

The Challenge

- Risk assessors needed a cost-effective means of locating relevant ecological toxicity data for:
 - Prioritizing chemical cleanup at hazardous waste sites
 - Assisting in the assessment of potential hazards of pollutants through the Clean Air Act, the Clean Water Act, the Federal Insecticide, Fungicide and Rodenticide Act and the Toxic Substances Control Act.
- Data must be identified with transparent processes and be accessible
- Duplicative efforts for data gathering wastes resources across state and federal agencies

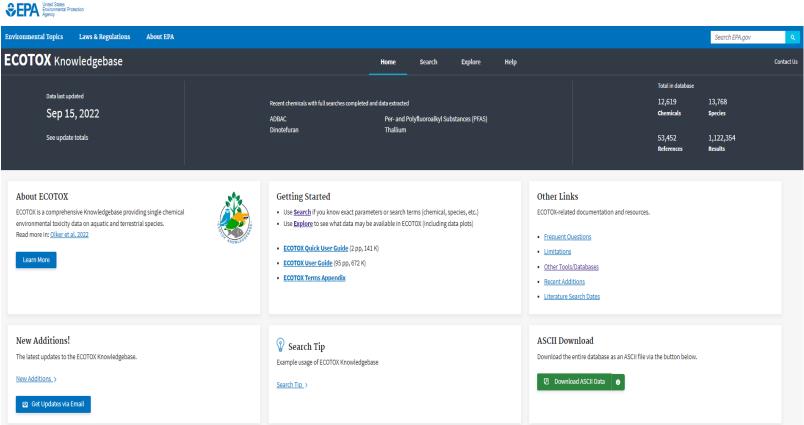
The On-going Challenge and Opportunities

- The accelerated pace of chemical risk assessment for ecological receptors
- Discovery of "new" chemicals of ecotoxicological concern
- Increase in number and diversity of journals
- Changing landscape of toxicity data used for ecological risk assessment
- Decreasing dependency of traditional whole animal testing

- Tool development to:
 - Expedite information gathering of information
 - Rapidly sort through BROAD keyword searches (e.g., chemical name)
- Digital formats of journals allow for use of data analytic applications
- Shift from apical endpoints to NAMs requires transparent development of controlled vocabulary for systematic data curation
- Providing a means to take full advantage of existing data before conducting new toxicity studies

What is the ECOTOXicology Knowledgebase?

- 30+ years of curating single chemical toxicity effects data for aquatic and terrestrial organisms
- Systematic and transparent literature search and review of open and grey literature
- >1 million test results from >52,000 references



www.epa.gov/ecotox

ECOTOX Data Curation Pipeline

chemical verification & development of search terms

Conduct literature searches

Identify and acquire potentially applicable studies

Review literature for applicability

Extract study and toxicity data



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Identify and acquire potentially applicable studies

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Extract study and toxicity data



- Verify CASRN
- Search various sources for chemical term
- Synonyms
- Eliminate poor search terms

Tak(Acilid OR Albrass OR Bexton OR "CP 31393" OR "Kartex A" OR Muharicid OR Niticid OR Propachlor OR Propachlore OR Ramrod OR Satecid OR "US EPA PC Code 019101")

* Web-based tool to identify and document relevant search terms

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Chemical-based Literature Searches

Search Engines

- 1. Scopus/Science Direct
- 2. ProQuest
- 3. Web of Science
- 4. PubAg/AGRICOLA
- 5. PubMed Toxline/TOXNET
- 6. Dissertation Abstracts
- * Semi-automated batch searches with Abstract Sifter Plus

chemical verification & development of search terms

Conduct literature searches

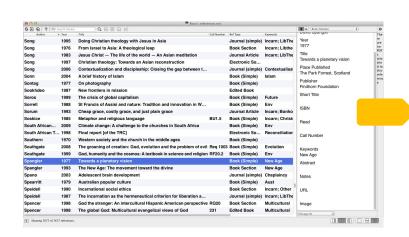
Identify and acquire potentially applicable studies

Review literature for applicability

Extract study and toxicity data



Literature Search Results



Title and Abstract Screening

 J Econ Entomol. 2016 Jul 18. pii: towl46. [Epub ahead of print]
 Sulfur Dust Bag: A Novel Technique for Ectoparasite Control in Poultry Systems Murillo AC(1), Mullens BA(2).

uuthor information: (]Department of Entomology, University of California, Riverside, CA 92521 (alock0030ucr.edu; bradley.mullens0ucr.edu) alock0030ucr.edu, (2)oepartment of mirodology, University of California, Riverside, CA 92521 (alock0030ucr.edu;

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0 The Authors 2016. Published by Oxford University Press on behalf of Entomological Society of America. All rights reserved. For Permissions, pleenall: journals.permissionsGoup.com.

DDI: 10.1093/jee/tow146

Full Text Screening

otoxicology 2, 93-120 (1993)

The impact of the Cyanamid Canada Co. discharges to benthic invertebrates in the Welland River in Niagara Falls, Canada

MIKE DICKMAN and GRAZYNA RYGIEL

Biological Sciences Department, Brock University, St. Catharines, Ontario, Canada L2S 3A

Received 15 July 1992; accepted 6 December 1992

In 1986, the International Joint Commission (IIC) recommended that the Niagara River watershed should be desired an Area of Concern (AGO, This III Cercommendation was ratified by the 4 signatures of the Great Lakes Water Quality Agreement. In order to delite an AGO, it as projects that german use that were previously impaired. To this and one waterspace of the other projects that german uses that were previously impaired. To the fine due stengende to determine whether or not the sediments at 3 study sites near the Cyanamid Canada (Concisial) Co. were contaminated as the with the world previously internative the natural blook which inhabit the contamination of such that would resid in the impairment of the natural blook which inhabit the

The Cyanamid Canada (Chemical) Co. discharges ammonia watest, cyanide, arsenic and a variety of beavy metals into treatment systems which ultimately duscharge to the Welland River, the major Canadian tributary to the Nagara River. This portion of the Welland River near the factory was designed at Provisidally significant (Class only wellands by the Orizant Ministry of any control of the Control of any control of the Control of any 2500 ml or dor to MCOE. 1997. Smilar discharge volumes occurred in 1999. In 1991, the total discharge was 2500 ml or doe for MCOE. 1991.

The majority of the benthic invertebrates collected from the study area were pollution toteran tax (e.g., shings some constituted 60% of all the organism collected). The lowest featherman tax (e.g., shings some constituted 60% of all the organism collected). The lowest featherman tax (e.g., shings some constituted of the collected of the collec

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0963-9292 © 1993 Chapman & Hall



Applicability Requirements

Identify and acquire potentially applicable studies

Review literature for applicability

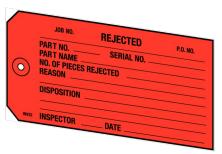
Key Area	Data Requirement	applicable studies	
Species	Taxonomically verifiable, ecologically-relevant organisms (including cells, organs, gametes, embryos, plant cuttings) [NOT bacteria, humans, monkeys, viruses, or yeast]		
Chemical	 Single, verifiable chemical toxicants, ad an acceptable route. 	Single, verifiable chemical toxicants, administered through an acceptable route.	
Exposure Amount (Concentration)	Exposure amount is quantified, either as a concentration in the environment when administered via soil or water, or as a dosage when introduced directly into or on the organism, via injection, orally, or topically.		
Exposure Duration	Known duration from the time of initial exposure to the time of measurement.		
Control	Must have a control treatment		
Effect	Biological effect measuredEffect concurrent with associated chem	Biological effect measured Effect concurrent with associated chemical exposure	
Publication Type	Primary source of the data [NOT a Review] Study must be a full article in English		

Exclusion Documentation

All Excluded and Non-Applicable studies are Tagged with the reason for rejection

- Abstract published as an abstract
- Bacteria only test organism is a bacteria
- CAS # Unavailable could not verify/locate chemical CAS Registry number
- Chemical method description of chemical analysis procedures
- Fate only report chemical distribution in media
- Human Health data on human subjects of surrogate animal subjects for human health risk assessment
- Incident reports death of animal by poison, but does not provide concentration/duration of exposure
- Method paper only reports methods for conducting a toxicity test or other aspect of an experiment
- Mixture paper reports results from mixture of chemicals; no single chemical exposure results
- Modeling results of the development of a model; no primary data available

- No Conc the authors report a response in an organism but do not provide conc/dose/app rate
- No Duration duration of exposure is not presented
- No Effect paper does not report observed responses adverse of otherwise
- No Toxicant (ozone, CO2)
- Non-English
- Nutrient in situ chemical tested as nutrient
- PUBL AS duplicate data published elsewhere
- Retracted paper retracted by Journal
- Review primary data published elsewhere
- Sediment only sediment concentration presented
- Survey chemical measured in organism, but lack quantification of exposure (dose/duration)
- Virus virus is only test organism
- Yeast yeast is only test organism



Chemical verification & development of search terms

Conduct literature searches

Identify and acquire potentially applicable studies

Review literature for applicability

Extract study and toxicity data

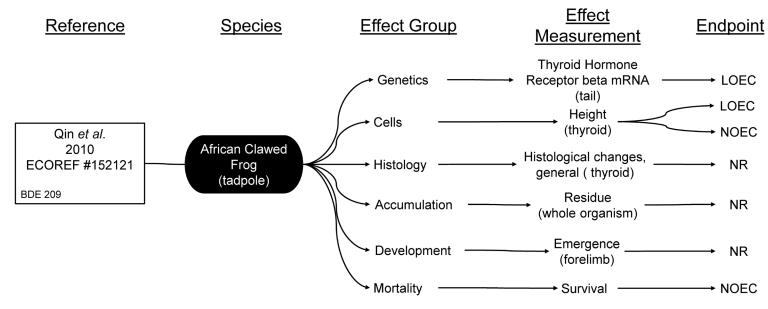




Journal of Environmental Sciences
Volume 22, Issue 5, 2010, Pages 744-751

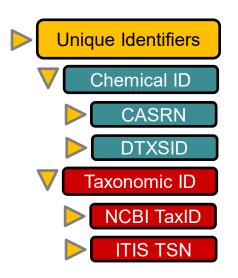


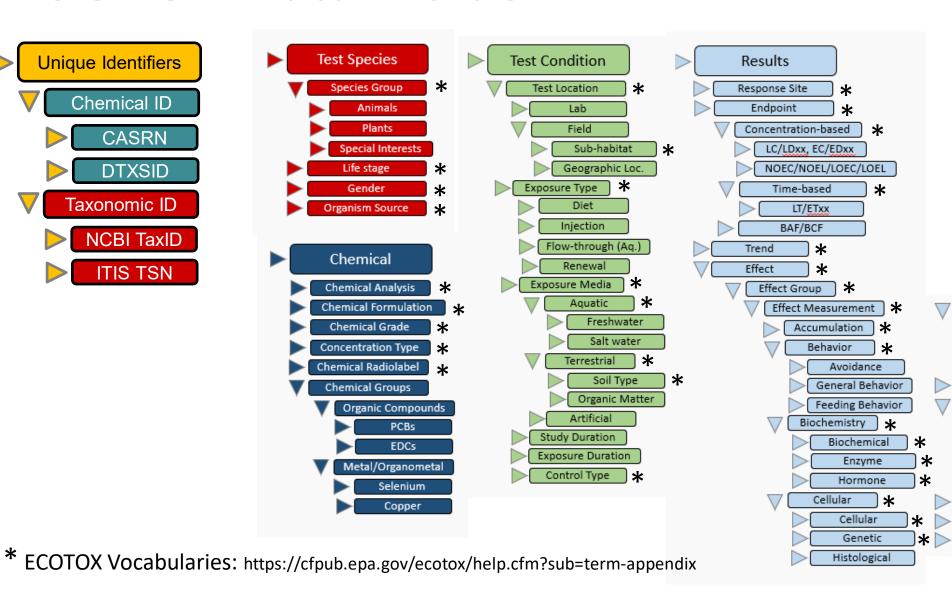
Thyroid disruption by technical decabromodiphenyl ether (DE-83R) at low concentrations in *Xenopus laevis*



NOEC = No Observed Effect Level LOEC = Lowest Observed Effect Level NR = Not Reported

ECOTOX Data Fields







Extract study

and toxicity

data

Growth

Mortality

Physiology

Reproduction

Population

Ecosystem

Growth

Development

Morphology

General Phys.

Injury

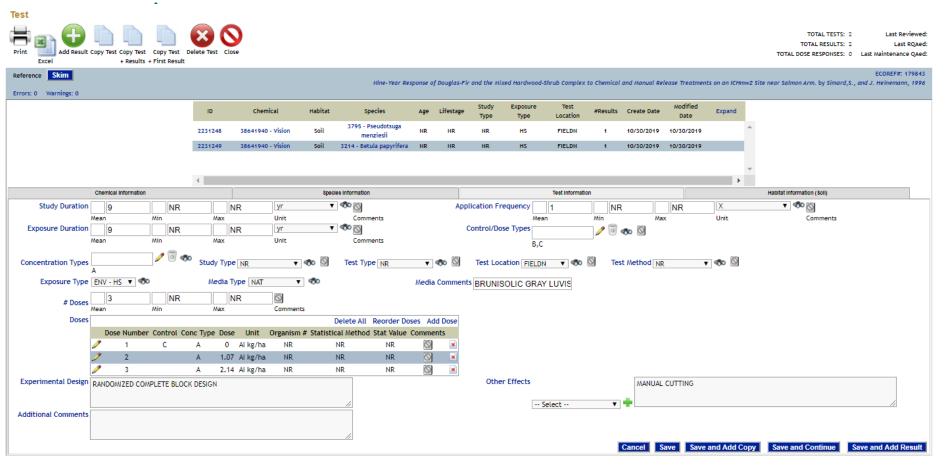
Immunity

Intoxication

]*

*

ECOTOX Data Fields





GUI for data curation

- Up to 90 entities (300-400 data fields) per study
- Following controlled vocabularies
- Computationally assisted
- Consistent extraction

* ECOTOX Vocabularies:

https://cfpub.epa.gov/ecotox/help.cfm?sub=term-appendix

Informing Study Evaluation

Regulatory Toxicology and Pharmacology 88 (2017) 227-237



Contents lists available at ScienceDirect

Regulatory Toxicology and Pharmacology

journal homepage: www.elsevier.com/locate/yrtph

Relevance and reliability of experimental data in human health risk assessment of pesticides



Toxicology Letters

Volume 189, Issue 2, 10 September 2009, Pages 138-144



"ToxRTool", a new tool to assess the reliability of toxicological data



Environmental Toxicology and Chemistry

Hazard/Risk Assessment 🗈 Open Access 😥 📵 🥏

CRED: Criteria for reporting and evaluating ecotoxicity

First published: 24 September 2015 | https://doi.org/10.1002/etc.3259 | Citations: 85

Select study evaluation questions with relevant ECOTOX field(s)

Chemical

- Is test substance identified? Required for inclusion in ECOTOX
- Is the purity of test substance reported? **Chemical Purity**
- Were chemical concentrations verified? <u>Chemical Analysis</u> (e.g., nominal versus measured concentrations)

Species

- Is the species given? Verifiable species (Scientific Name, etc.) required for inclusion in ECOTOX
- Are the organisms well described? <u>Organism Source</u>, <u>Lifestage</u>,
 Age, <u>Gender</u>, <u>Initial</u> and <u>Final Weight</u>

Test Conditions

- Are appropriate controls performed? A control is required for inclusion in ECOTOX, type described in Control
- Is a guideline method (e.g., OECD) used? <u>Test Method</u>
- Are the experimental conditions appropriate and acceptable for the test substance and organism? <u>Test Method</u>, <u>Media Type</u>, <u>Test Location</u>, <u>Experimental Design</u>, Physical and Chemical Soil and Water Parameters (e.g., <u>pH</u>, <u>Temperature</u>, <u>Dissolved</u> <u>Oxygen</u>)

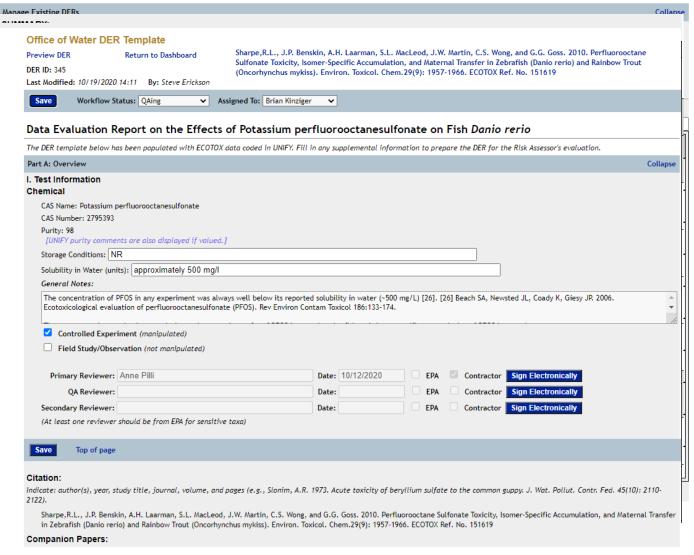
Test Results

- Are the reported effects and endpoints appropriate for the purpose, test substance and organism? <u>Effect Measurement</u>, <u>Endpoint</u>
- Is the response/effect statistically significant? **Statistical**Significance, Significance Level

Developing Data Evaluation Tools



Collaboration with USEPA's Office of Water

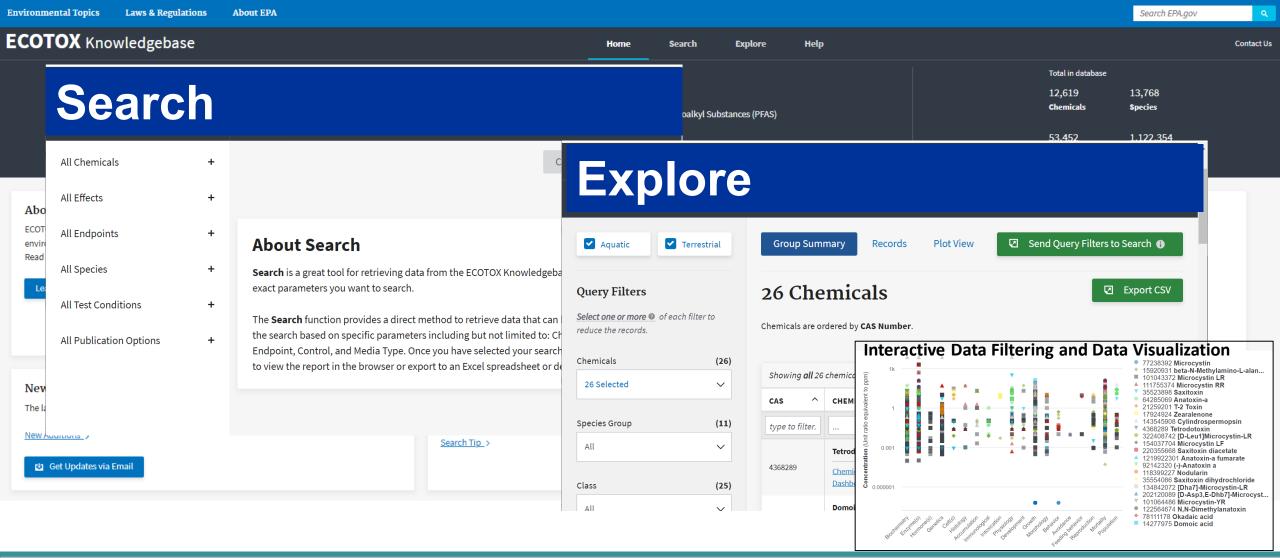


- Application pulls objective data extracted from existing records
- Platform now has ability for curators to add subjective observations
- Allows multiple reviewers
 - Primary
 - QA
 - Secondary

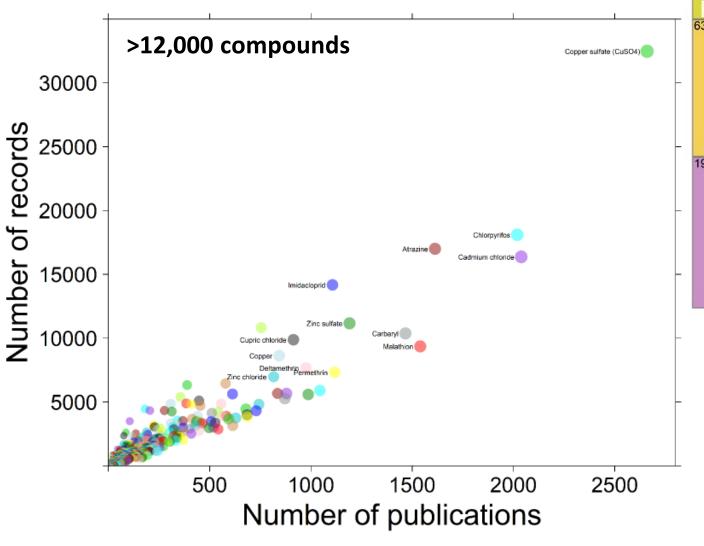
ECOTOX Version 5

www.epa.gov/ecotox

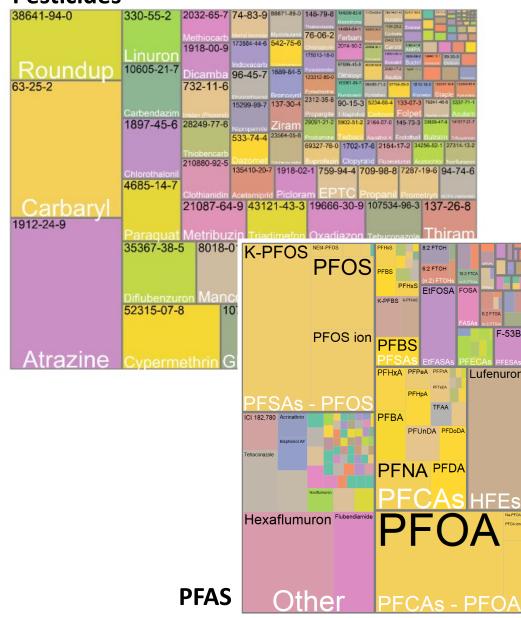




Chemicals

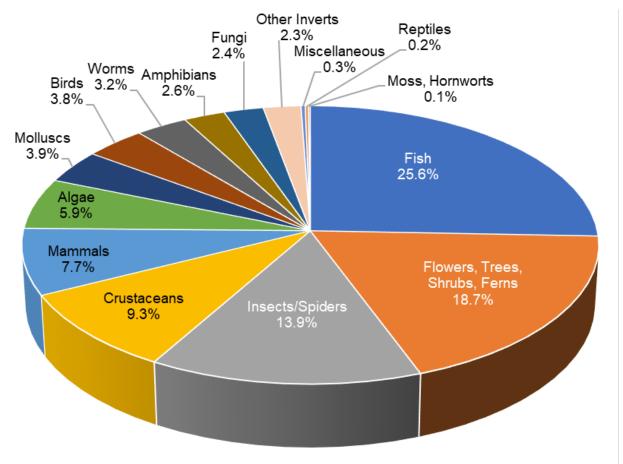


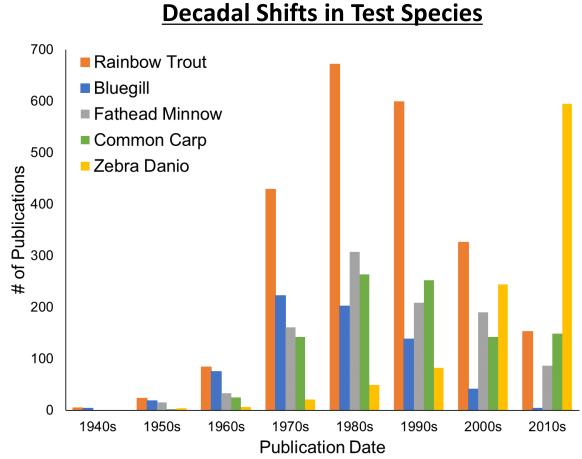
Pesticides



Species (n = 13,621)

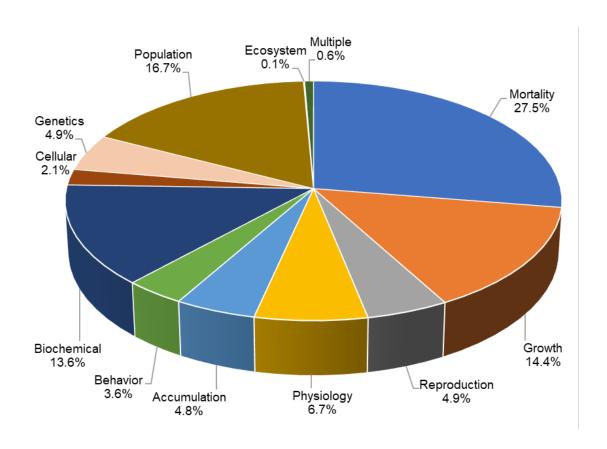
% of Records by Species Group (Sept 2021)



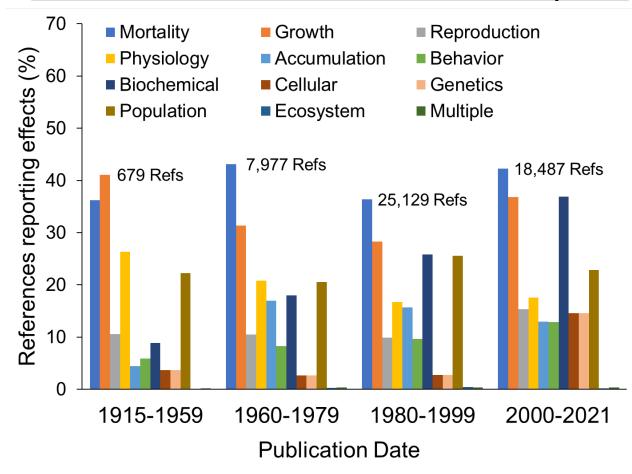


Diversity of Effects

% of Records by Effect Group (Sept 2021)



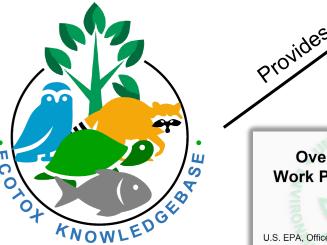
Increases in Biochemical and Genetic Effects Reported



18

Applications

Chemical environmental toxicity data for aquatic life, terrestrial plants and wildlife



EPA Program Offices and Regions,
States, Tribes, Other Federal Agencies
and International Entities

<u>Ecological Risk Assessment</u> for Office of Pesticides for chemical registration and re-registration.

<u>Ambient Water Quality Criteria</u> for Aquatic Life

Ecological hazard data for the <u>Prioritization</u> and <u>Assessment of Chemicals</u> for TSCA/Lautenberg Act.

Overview of TSCA
Work Plan Methodology

Maria Doa
U.S. EPA, Office of Pollution Prevention and Toxics
December 11, 2017

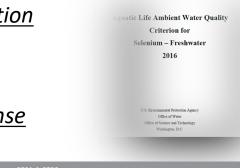
TEXAS COMMISSION ON ENVIRONMENTAL QUALITY

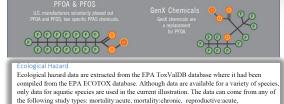
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Ecological Site Assessments and in Emergency Response by Office of Land and Emergency Management (Superfund and ORCR), HQ, Regions and States.

<u>Ecological toxicity data for PFAS</u> to researchers, EPA Eco Risk Assessment Forum, DoD Tri-Services ERA Work Group and others.





reproductive:chronic, growth:acute, growth:chronic (all from ECOTOX). The types of effect levels are LDxx/LCxx/ECxx/EDxx where xx can range from 1% to 100%, and LOEL/NOEL/LOEC/NOEC. Values must be in units of mg/L. For each chemical, the lowest toxicity value was separately determined for acute and chronic studies, regardless of species. Th

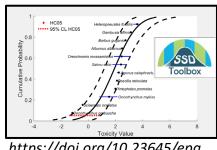
Applications

Chemical environmental toxicity data for aquatic life, terrestrial

plants and wildlife



Data used for



https://doi.org/10.23645/epa comptox.11971392

Linking environmental contaminant concentrations to potential effects

Identify data gaps, Inform study design, Compare to new toxicity studies

Adverse Outcome Pathway (AOP) development

EPA Program Offices and Regions, States, Tribes, Other Federal Agencies and International Entities

Tools and Applications

Toxicity Reference Values (TRVs) and Benchmarks

Species Sensitivity Distributions (e.g., US EPA's SSD Toolbox, Endangered species analyses, U.S. EPA's WebICE, NOAA's CAFE)

Predicted No Effect Concentrations (PNECs) and

Eco-TTCs (e.g., EnviroTox, NORMAN)

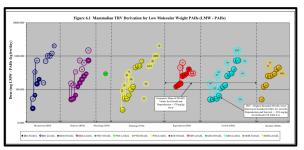
Quantitative Structure-Activity Relationships and other models



https://asartoolbox.org



https://www.epa.gov/tsca-screeningtools/ecological-structure-activityrelationships-ecosar-predictive-model



https://www.epa.gov/chemical-research/interimecological-soil-screening-level-documents





https://www.norman-network.com/nds/ecotox

T.E.S.T. (Toxicity **Estimation Software Tool)**

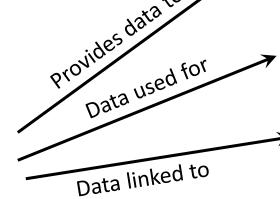
https://www.epa.gov/chemicalresearch/toxicity-estimationsoftware-tool-test

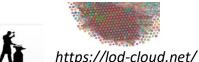


Applications

Chemical environmental toxicity data for aquatic life, terrestrial plants and wildlife







Biological Ontologies*

http://www.obofoundry.org/

Pathways and Disease Databases*







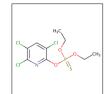
Health & Environmental Research Online (HERO)* (https://hero.epa.gov/)

EPA Program Offices and Regions, States, Tribes, Other Federal Agencies and International Entities

Tools and Applications

Databases/Resources

U.S. EPA CompTox Chemicals Dashboard (https://comptox.epa.gov/dashboard/)
OECD eChemPortal (https://www.echemportal.org/echemportal/



Adverse Outcome Pathway Wiki* (https://aopwiki.org/)



Sequence Alignment to Predict Across Species
Susceptibility (SeqAPASS)*
(https://doi.org/10.1093/toxsci/kfw119)

*Future

Summary

- Systematic and transparent procedures to identify and curate ecological toxicity data
- 30 year plus history, with major recent updates and evolution in the near future
 - Maintain comprehensive and quality review of toxicity data
 - Enhance ease of data access and clarity
 - Meet the demands for increased pace of chemical assessments
 - Expand to reflect shifts in toxicity testing paradigm
- Curated data on public website (<u>www.epa.gov/ecotox</u>), readily available for exploration, querying, and export for risk assessments, risk management and research

Acknowledgements and Contact Info

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US EPA ORD Center for Computational Toxicology and Exposure

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Dale Hoff, GLTED Division Director

Contract staff:

General Dynamics Information Technology (GDIT)
SpecPro Professional Services (SPS)

Senior Environmental Employment (SEE) staff

www.epa.gov/ecotox

ECOTOX Support:

218-529-5225

ecotox.support@epa.gov