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Mebane, C.A., Sumpter, J.P., Fairbrother, A. et al. (12 more authors) (2019) Scientific integrity issues in environmental toxicology and chemistry: improving research reproducibility, credibility, and transparency. Integrated Environmental Assessment and Management, 15 (3). pp. 320-344. ISSN 1551-3777

https://doi.org/10.1002/ieam.4119

This is the peer reviewed version of the following article: Mebane, C. A., Sumpter, J. P., Fairbrother, A., Augspurger, T. P., Canfield, T. J., Goodfellow, W. L., Guiney, P. D., LeHuray, A., Maltby, L., Mayfield, D. B., McLaughlin, M. J., Ortego, L., Schlekat, T., Scroggins, R. P. and Verslycke, T. A. (2019), Scientific Integrity Issues in Environmental Toxicology and Chemistry: improving research reproducibility, credibility, and transparency. Integr Environ Assess Manag., which has been published in final form at https://doi.org/10.1002/ieam.4119. This article may be used for non-commercial purposes in accordance with Wiley Terms and Conditions for Use of Self-Archived Versions.

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Critical Review

Integrated Environmental Assessment and Management

DOI 10.1002/ieam.4119

Scientific Integrity Issues in Environmental Toxicology and Chemistry: improving research reproducibility, credibility, and transparency[†]

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⁺This article has been accepted for publication and undergone full peer review but has not been through the copyediting, typesetting, pagination and proofreading process, which may lead to differences between this version and the Version of Record. Please cite this article as doi: [10.1002/ieam.4119]

All Supplemental Data may be found in the online version of this article at the publisher's website.

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Submitted 27 February 2018; Returned for Revision 7 December 2018; Accepted 26

December 2018

Abstract

High profile reports of detrimental scientific practices leading to retractions in the scientific literature contribute to lack of trust in scientific experts. While the bulk of these have been in the literature of other disciplines, environmental toxicology and chemistry are not free from problems. While we believe that egregious misconduct such as fraud, fabrication of data, or plagiarism is rare, scientific integrity is much broader than the absence of misconduct. We are more concerned with more commonly encountered and nuanced issues such as poor reliability and bias. We review a range of topics including conflicts of interests, competing interests, some particularly challenging situations, reproducibility, bias, and other attributes of ecotoxicological studies that enhance or detract from scientific rigor, relevant reproducible research, transparency in competing interests, methods and results, and education. This article is protected by copyright. All rights reserved

Introduction

Large segments of society are distrustful of scientific and other experts. Some have suggested that we are in a culture in which reality is defined by the observer and objective facts do not change peoples' minds, and those that conflict with one's beliefs are justifiably questionable (Campbell and Friesen 2015; Nichols 2017; Vosoughi et al. 2018). Science and scientists have been central to these debates, and the boundaries of science, policy and politics may be indistinct. In a social climate skeptical of science, the easy availability of numerous reports of dubious scientific practices gives fodder to skeptics. Because environmental regulations on use of chemicals and waste management rely heavily on the disciplines of ecotoxicology and chemistry, the integrity of the science is of utmost importance. Here we discuss scientific integrity in the applied environmental sciences, with a focus on ecotoxicology and how the role and culture of the Society of Environmental Toxicology and Chemistry (SETAC) may influence such issues.

Science has long endured questionable science practices and a skeptical public. Galileo's criticisms of prevailing beliefs resulted in his issuing a public retraction of his seminal work. In contrast, purported science "discoveries" such as Piltdown Man, canals on Mars, cold fusion, Archaeoraptor, homeopathic water with memory, arsenic-based life, and many others have not stood the test of time (Gardner 1989; Schiermeier 2012). By 1954, Huff and Geist (1954) illustrated how the presentation of scientific data could be manipulated to become completely misleading yet accurate. Are things worse now? Recent articles in both the scientific literature and popular print and broadcast venues paint a bleak picture of the status of science. One does not have to search hard to find plenty of published concerns about the credibility of science. These include overstated and unreliable results (Harris and Sumpter 2015; Henderson and Thomson 2017; Ioannidis 2005), conflicts of interest (Boone et al. 2014; McGarity and Wagner 2008; Oreskes et al. 2015; Stokstad 2012; Tollefson 2015), profound bias (Atkinson and Macdonald 2010; Bes-Rastrollo et al. 2014; Suter and Cormier 2015a, b), suppression of results to protect financial interests (Wadman 1997; Wise 1997), deliberate misinformation campaigns as a public relations strategy for financial or ideological aims (Baba et al. 2005; Gleick and 252 co-authors 2010; McGarity and Wagner 2008; Oreskes and Conway 2011), political interference with or suppression of results from government scientists (Hutchings 1997; Ogden 2016; Stedeford 2007), self-promotion and sabotage of rivals in hypercompetitive settings (Edwards and Roy 2016; Martinson et al. 2005; Ross 2017), publication bias, peer review and authorship games (Callaway 2015; Fanelli 2012; Young et al. 2008), selective reporting of data or adjusting the questions to fit the data (Fraser et al. 2018), overhyped institutional press releases that are incommensurate with the actual science behind them (Cope and Allison 2009; Sumner et al. 2014), dodgy journals (Bohannon 2013), and dodgy conferences (Van Noorden 2014).

Such published concerns reasonably raise doubts about science and scientists and could even lead some to conclude that the contemporary system of science is broken. In writing this commentary, we attempt to address some prominent science integrity concerns in the context of environmental toxicology and chemistry. In our view, there is ample room for improvement within our discipline, but the science is not broken, and some criticisms are overstated. In writing this commentary, we do not pretend to have solutions that will overturn insidious pressures on scientists and funders for impressive results, or hold some moral high ground making us immune from such pressures ourselves, or that all of our own works are above reproach. Our recommendations are pragmatic, not dogmatic. Our goal is to nudge practices and pressures on scientists to advance the science, while maintaining and improving credibility through transparency, ongoing review, and self-correction.

Many of the prominent science integrity controversies have been in the high stakes biomedical discipline, and in response that discipline probably has done more self-evaluation and taken more steps toward best practices than most other disciplines. Results of self-reported, anonymous surveys of scientists, mostly in the biomedical fields, have not been reassuring. In a 2002 survey of early and mid-career scientists, 0.3% admitted to falsification of data, 6% to a failure to present conflicting evidence, and 16% to changing of study design, methodology or results in response to funder pressure (Martinson et al. 2005). A subsequent meta-analysis of surveys suggested problems were more common, with close to 2% of scientists admitting to having been involved in serious misconduct, and over 70% reported they personally knew of colleagues committing less severe detrimental research practices (Fanelli 2009). Overt misconduct can occur in ecotoxicology just as with any discipline (Enserink 2017; Keith 2015; Marshall 1983; http://retractiondatabase.org, search term "toxicology") and when exposed, is universally condemned and, in many countries, is career ending. In contrast, the ambiguous, more nuanced issues of science integrity that all of us are likely to experience in our careers require thoughtful consideration, not condemnation. It is toward the latter that we discuss efforts toward remedies from other disciplines to examine similar issues in ecotoxicology, focusing on SETAC.

What is "science" in the context of scientific integrity?

Before we can discuss integrity in ecotoxicology and related environmental science fields, we must first distinguish what is meant by "science" in this context. Broadly speaking, environmental science includes the disciplines of biology, ecology, chemistry, physics, geology, limnology, mineralogy, marine studies, and atmospheric studies; i.e., the study of the natural world and its interconnections. The applications of environmental science extend to agriculture, fisheries management, forestry, natural resource conservation, and chemicals management, all of which have associated multi-billion-dollar industries and vocal environmental advocacy groups.

The subdiscipline of environmental toxicology or ecotoxicology, pursued by SETAC scientists, studies in great detail how the natural world is influenced by chemicals, both natural and synthetic, introduced by human endeavors that are largely in pursuit of the production of desired goods and services (food, clean water, plastic products, metals, etc.). Because exposure to chemicals can have negative and sometimes unexpected consequences for people and the environment, a body of regulation has developed over the past century to control the kinds and amounts of allowable chemical exposures. Such regulations necessarily are based on scientific concepts such as Paracelsus' directive that "the dose makes the poison" and physicochemical properties that influence transport and fate of substances. Because of the complexity, inexactitude, and uncertainty of ecotoxicology and associated sciences, rulemaking often is subject to challenge, leading to accusations of profit over people or the environment or unreasonably restrictive and burdensome requirements. Scientists are called upon to inform disputes based on their knowledge or underlying principles or enter the conversation through self-initiated in-depth literature review and commentary. Only by conscientiously adhering to fundamental principles of the scientific method can environmental scientists maintain their integrity and continue to play a valid role in environmental policy and management.

What is "scientific integrity"?

Impeccable honesty is a fundamental tenet of science. When we read a paper, we might not agree with the conclusions, authors' interpretations of its implications, importance, or many other things, but we have to be confident that the procedures described were indeed followed and all relevant data were shown, not just those fitting the hypothesis. As Goodstein (1995) put it: *"There are, to be sure, minor deceptions in virtually all scientific papers, as there are in all other aspects of human life. For example, scientific papers typically describe investigations as they logically should have been done rather than as they actually were done. False steps, blind alleys and outright mistakes are usually omitted once the results are in and the whole experiment can be seen in proper perspective." Indeed, no one wants to read the chronology of a study. However, should for example, the omissions include unfruitful statistical fishing trips or anomalous data that were assumed to be in error because they didn't fit expectations, such little omissions may bias the story and the body of literature.*

Various professional and governmental organizations have established policies and definitions prescribing research integrity, responsible conduct of science, or scientific integrity. These may include broad statements of attributes such as the U.S. National Academy of Science's (NAS) six values that they considered most influential in shaping the norms that constitute research practices and relationships and the integrity of science: objectivity, honesty, openness, accountability, fairness, and stewardship (NAS 2017). More specific "research integrity" guidelines define appropriate expectations of individual researchers and their institutions and

may be highly procedural. Protecting the privacy, rights and safety of human research participants and animal welfare with institutional review board clearance requirements is a common element of research integrity guidelines. Academic research integrity guidelines have been established individually or in aggregate by research funders and individual institutions (ARC 2007; Goodstein 1995; NRC-CNRC 2013; NRC 2002; Resnik and Shamoo 2011; Steneck 2006). In most countries, research institutions are usually responsible for investigating potential breaches of research integrity by its scientists, although this can create difficult conflicts of interest for the institution (Glanz and Armendariz 2017). A prominent recent exception is China, which announced reforms that no longer allow institutions to handle their own misconduct investigations (Cyranoski 2018).

Whether research integrity guidelines should best be defined narrowly or broadly has been an area of controversy. As of 2015, 22 of the world's top 40 research countries had national research conduct policies, all of which included fabrication, falsification, and plagiarism (FFP), with some going further. In this context, "fabrication" is making up data; "falsification" includes manipulating studies or changing or omitting data such that the record does not accurately reflect the actual research; and "plagiarism" includes the appropriation of another person's ideas, methods, results, or words without giving appropriate credit (https://ori.hhs.gov/definitionmisconduct). The Research Councils of the UK has a lengthy list of misdeeds including FFP, misrepresentation, breach of duty of care, and improper dealing with allegations of misconduct, with many subcategories (NAS 2017). In contrast, from the 1980s to 2000, the National Science Foundation (US) had defined serious science misconduct broadly to include, "...fabrication, falsification, plagiarism, or other practices that seriously deviate from those that are commonly accepted within the scientific community for proposing, conducting and reporting research" (Goodstein 1995). The controversial part was the catchall phrase "practices that seriously deviate from those commonly accepted ... " To the stewards of public science funds, such a catchall phrase was preferable to an itemized lists of all potential avenues of mischief, yet it raised the specter of penalizing scientists who strayed too far from orthodox thought (Goodstein 1995). In 2000, this definition of disbarring research misconduct was narrowed to just "fabrication, falsification, or plagiarism in proposing, performing, or reporting research" with lesser offenses classified as questionable research practices. Other misconduct was defined as forms of unacceptable behavior that are clearly not unique to the conduct of science, although they may occur in the laboratory or research environment." Yet only FFP research misconduct findings were subject to reporting requirements to federal science funding agencies, with questionable science practices or other misconduct handled locally (NAS 2017; Resnik et al. 2015).

In many countries, there is an active debate about whether a legal definition is appropriate for something that is really an academic judgment rather than a legal one. Denmark recently similarly narrowed its broad definitions of research misconduct to only FFP following high profile cases in which scientists succeeded in having their academic misconduct findings overturned in the courts. Yet if research conduct policies are considered "academic" without legal weight, institutions may have difficulty enforcing polices, such as when deliberate intent is required to be shown and the researcher claims "honest mistake." For instance, the U.S. Office of Research Integrity found that a tenured professor had committed research misconduct by inappropriately altering data in five images from three papers. Yet when the university sought to terminate her, she fought back contesting the university's procedures, and the university ultimately paid her \$100,000 USD to leave (Stern 2017). In private research, it is not obvious which scientific integrity concepts have the force of law. In an example from the U.S., testimony of egregious breaches of scientific integrity norms (including faking credentials and selective publication of only favorable results) was disallowed in a court dispute between two private companies because there was no federal law on scientific integrity (Krimsky 2003).

Science is a human endeavor and the "other misconduct" that scientists may commit is diverse and may be horrific, such as bullying and abuse of power; taking advantage of students or subordinates; sexual coercion, assault, or harassment; misuse of funds; sabotage; specious whistleblowing or retaliation against valid whistleblowers; and poisoning coworkers (e.g., Else 2018; Extance 2018; Ghorayshi 2016; Gibbons 2014; http://retractionwatch.com). The exclusion of such malfeasance from "research misconduct" has been questioned. For example, a researcher who failed to meet her study objectives after being sabotaged by a rival argued that she was further penalized by being instructed not to divulge the reason for her study failures to her funders (Enserink 2014). In contrast, institutions often do go beyond the minimum "FFP" definition in their policies (Resnik et al. 2015), which has led to objections of conflation of egregious misconduct such as fraud with failure to comply with administrative requirements that did not compromise data validity (Couzin-Frankel 2017).

The U.S. National Academy of Science (NAS 2017) recently argued that the definitions of research misconduct as fabrication, falsification, or plagiarism were too narrow. In particular, questionable research practices were more than just "questionable," but were clear violations of the fundamental tenets of research and were given a less ambiguous label of "detrimental." Consensus detrimental research practices were:

 Detrimental authorship practices that may not be considered misconduct, such as honorary authorship, demanding authorship in return for access to previously collected data or materials or denying authorship to those who deserve to be designated as authors. [Here we think it is important to distinguish between pairing a data reuse with

an invitation to collaborate and share authorship versus demanding authorship as a condition of data access (Duke and Porter 2013)].

- 2. Not retaining or making data, code, or other information/materials underlying research results available as specified in institutional or sponsor policies, or standard practices in the field.
- 3. Neglectful or exploitative supervision in research.
- 4. Misleading statistical analysis that falls short of falsification.
- 5. Inadequate institutional policies, procedures, or capacity to foster research integrity and address research misconduct allegations, and deficient implementation of policies and procedures, and
- 6. Abusive or irresponsible publication practices by journal editors and peer reviewers (NAS 2017).

The term "scientific integrity" is sometimes used synonymously with research integrity. However in recent usage, the term has included insulation of science from political interference, manipulation, or suppression of science (Doremus 2007; Douglas 2014). The term "scientific integrity" has been used in government science policy in the United States. There, scientific integrity guidelines were developed in an overarching sense that includes research integrity at the individual and institutional level but were also intended to protect federal scientists from political interferences. Political officials were not to alter or suppress scientific findings, and transparency was encouraged in the preparation of the government-supported scientific research (Obama 2009; Stein and Eilperin 2010). The scientific integrity guidelines in the US were followed by derivative policies intended to put substance to the transparency provisions, requiring openaccess to federally funded research articles and more importantly, requiring archiving and public availability of the underlying raw data (Holdren 2013). These broad policies become more specific and procedural in government science agencies, and expanded to codes of scholarly and scientific conduct such as a list of 19 principles for the U.S. Department of Interior (U.S. Department of Interior 2014).

We expect the vast majority of scientists consider themselves to hold science integrity, as selfdefined in terms of honesty, transparency, and objectivity, sticking to the research question and avoiding bias in data interpretation (e.g., Shaw and Satalkar 2018). Yet most scientists will encounter ethically ambiguous situations. For instance, some may feel that they struggle to advance science against a rising tide of administrative requirements accompanied by declining support for science and increasing competition for funding. When does cutting through bureaucratic institutional requirements cross the line from being commendable efficiency to violating research integrity rules? Using grant/project funds for unrelated purchases or

conference travel? Should minor misbehaviors such as posting ones' article on a website after signing a publication and copyright transfer agreement with the publisher agreeing not to do so still be considered misbehaviors when done by many? When does cleaning data become cooking data when, for example, anomalous values are suppressed? There are many ethically ambiguous situations in which scientists may consider doing the "right thing" (compliance with all rules) might need to be balanced with doing the "good thing," especially when the welfare of others such as students or subordinates is involved (Johnson and Ecklund 2016).

To us, scientific integrity can be simplified to cultures of personal integrity plus a few profession-specific provisions of transparency and reproducibility. At their roots, these norms are those children are hopefully acculturated to in primary school. *Tell the truth, and tell the whole truth* (no data sanitizing, selective reporting, and report all conflicts), *tell both sides of the story* (avoid bias), *do your own work* (no plagiarism), *read the book, not just the back cover before writing your report* (properly research and cite primary sources), *show your work for full credit* (transparency), *practice makes perfect* (rigor), *share* (publish your work and data in peerreviewed outlets for collective learning), and *listen* (with humility and collegial fraternity to observations and suggestions of others). Finally, the golden rule "*do unto others as you would have them do unto you*" should resonate throughout the professional interactions of environmental scientists, and especially in peer reviewing and data sharing. When encountering an inevitable science dispute, keep criticisms objective, constructive, and focused on the work and not the worker; do peer reviews of your rivals' work as you would hope to receive reviews of your own, reward and recognize good behavior in science, and so on.

The interested scientist: conflicts of interest, competing interests, and bias

Although we would like to believe that outright fraud or deliberate campaigns to manipulate science are rare in the environmental sciences, at some points in their careers almost every practicing scientist must grapple with questions of conflicting or competing interests and must guard against bias in approaches and interpretation.

The term "conflicts of interest" is commonly narrowly defined to financial conflicts. One definition is "a set of conditions in which professional judgment concerning a primary interest tends to be or could be perceived to be unduly influenced by a secondary interest (such as financial gain). More simply, a conflict of interest is any financial arrangement that compromises, has the capacity to compromise, or has the appearance of compromising trust (Krimsky 2003, 2007). The term "competing interests" is often used where non-financial factors compete with objectivity, such as allegiances, personal friendships or dislikes, career advancement, having taken public stances on an issue, political, academic, ideological, or religious affiliations (Nature Editors 2018b; PLOS Medicine Editors 2008). Bias in study design

or data interpretation may arise from either conflicts or competing interests and can be either overt or unrecognized by the scientist (Suter and Cormier 2015b)

Generally, the concern over conflicting or competing interests in science is that secondary interests such as financial gain or maintaining professional relationships compromise the primary interest of upholding scientific norms such as reporting data accurately and completely, interpreting data appropriately, and acknowledging value judgments or interpretive assumptions (Elliott 2014). Conflict of interest policies may be better developed in the biomedical fields than in the applied environmental sciences because the former often involves human participants, and because of the strong financial ties between academia and the pharmaceutical industry (Tollefson 2015). For instance, if a research team is reporting on the efficacy of a medical device or a pharmaceutical, and they or their employers hold a patent or stand to gain financially from a positive report, then they clearly have a financial conflict of interest (Figure 1). Examples of financial conflicts of interest encountered in common life include the physician who recommends to the patient a medical procedure that would conveniently (and lucratively) be conducted in a private surgery center in which the physician is a part owner, or the financial advisor who receives commissions when clients are steered to financial products offered through their employer. Such arrangements don't mean that the patient's care will suffer or bad financial advice proffered, but these self-interests compete with the interests of those they serve (Cain et al. 2005).

The mere existence of a potential conflict of interest should not alone throw results in doubt where it is disclosed and acknowledged appropriately. However, although most articles in the environmental sciences routinely disclose funding sources that could be perceived as potential conflicts of interest, major omissions have occurred (Krimsky and Gillam 2018; McClellan 2018; Oreskes et al. 2015; Ruff 2015; Tollefson 2015). For instance, the findings of a study on risks of contamination from natural gas extraction from hydraulic fracturing of bedrock were undermined when it came out (apparently unbeknownst to the university) that the research supervisor was being paid 3X his university salary by serving as an advisor to an oil and gas company invested in the practice. The failure to disclose this financial relationship in the publication brought the study's objectivity and credibility into question, independent of its substance (Stokstad 2012). Authors and journals have been criticized for gaming ethical financial disclosure requirements, such as by overly narrow disclosures or disclosing a conflict in the reviewers and readers) but not including it in the actual article (Marcus and Oransky 2016).

It should be noted that the severe conflicts of interest that some academic biomedical researchers have created for themselves by setting up business interests to directly and personally profit from their research outcomes (Krimsky 2003) are probably much less of an issue in the

environmental sciences. Dual affiliations and the resultant potential for divided loyalties for university researchers have certainly come to light in the environmental sciences, such as if the scientist has a public facing, disinterested, researcher identity but privately has set up spin-off personal, business interests (Fellner 2018; Stokstad 2012). While we are not aware of any systematic review, we think these situations are far less pervasive in the environmental sciences than biomedicine. Rather, in ecotoxicology and environmental chemistry, the more common (and less insidious) concern for authors and institutions to be self-aware of the potential for funding bias through unconscious internalization of the interests of their research sponsors. The informative value of conflict of interest or funding disclosures vary. The shortest (and least informative) statement we have seen was that "the usual disclaimers apply" (Descamps 2008), while the detailed disclosures in biomedical literature can go on for pages (Baethge 2013; ICMJE 2016). Funding sources can be obscured by channeling funding through intermediaries, such as a critical review of cancer risks from talcum powder funded by a law firm involved in toxic tort litigation (Muscat and Huncharek 2008). Requirements for highly detailed disclosures risk diminishing their importance to that of the "fine print" cautions in commerce that are seldom read. Much like computer software user terms and conditions that have to be clicked past or the ubiquitous consumer product safety stickers that may be written more to avoid product liability claims than for practical safety, detailed conflict of interest disclosures may reach a point of diminishing returns. There is some evidence that over reliance on conflicts disclosure is ineffective or can give moral license to scientists to be biased (Cain et al. 2005). Our view is that true financial conflicts of interests should be avoided, not just disclosed. Yet for most scientists in ecotoxicology and the environmental sciences who sought and received funding in order to pursue studies, simple, unambiguous statements of the funding sources should generally be sufficient.

Non-financial competing factors may also compete with scientific objectivity. Factors or values such as these are usually termed "competing interests" reserving "conflicts of interest" for financial conflicts (Nature Editors 2018b; PLOS Medicine Editors 2008). In our observations, competing interests are rarely mentioned in environmental science publications. Rather, they are often discussed behind the scenes, such in correspondence between an editor and potential reviewers, along the lines of "*yes I would be happy to review this article and believe I can be objective, however, you should know that I used to be a labmate of the PI and we collaborated on an article 3 years ago.*" Marty et al (2010) give such an example of a disclosure of competing interests based on personal relationships. Whether or how competing interests or values affect the assumptions and perspectives of scientists' should be more formally stated is an area of rich debate in the philosophy of science literature (Douglas 2015; Elliott 2016; PLOS Medicine Editors 2008).

We reiterate our belief that the existence of a potential conflicts or competing interests is a ubiquitous part of the environmental science landscape and do not indicate poor science. Most scientists strive to present unbiased data and interpret their data evenhandedly. However, the varied experiences of scientists can influence their perspectives in ways that they may not recognize themselves. The transparency in disclosure reminds the reader to consider perspectives and alternate interpretations when judging the merits of a study, synthesis paper, or risk assessment.

Bias

Many of the published concerns in the environmental science literature come down to cognitive bias. Science is not value free, and personal bias in interpreting science is often related to differing worldviews (Douglas 2015; Elliott 2016; Lackey 2001; Nuzzo 2015). For instance, the collapse of major fisheries that ostensibly had been scientifically managed for sustainable yields helped inspire the Precautionary Principle. This philosophy sought more cautious management and the reversal of the burden of proof for sustainable exploitation of natural resources (Peterman and M'Gonigle 1992). Those with precautionary principle or risk assessment worldviews may interpret the same set of facts very differently. The precautionary principle adherent may emphasize absence of conclusive evidence of safety, and the risk assessment adherent may emphasize absence of conclusive evidence of harm (Fairbrother and Bennett 1999). In such settings, values and biases are interwoven. Even self-disciplined scientists who seek openness and objectivity carry some biases from experiences and acculturation (here meaning how working in different environmental organisations can lead scientists to modify their perception and thinking). Recognizing sources of bias does not imply ill intent, for just the process of acculturation to a particular place of employment can bias perceptions and inclinations (Figure 2; Brain et al. 2016; Suter and Cormier 2015a, b).

Professional societies such as SETAC can serve as a form of acculturation; some of the authors of this essay have been active members of SETAC for much longer than they have been employed by any single employer. Even self-disciplined scientists who seek openness and objectivity carry biases from their experiences. What becomes particularly difficult to self-regulate is the convergence of cognitive bias, a human nature to seek to please one's patron, and the interests of one's employer or client. For instance, studies funded by drug or medical device makers tend to find positive effects that favor the company funding the research (Lexchin et al. 2003; Smith 2006), and the funding effect for studies of chemical toxicity may lean toward finding negative effects (Bero et al. 2016; Krimsky 2003, 2013). However, concordance between a funder's self-interest and research findings does not alone indicate bias. Alternatively the industry-funded researchers who might have less extensive experience, the industry-funded

work was more thoroughly vetted based on prior internal research, or the industry-funded scientists might have better ability to obtain the resources and skill to carry out well focused and rigorous research (Krimsky 2013; Macleod 2014). It is doubtful that these influences can be completely separated. To us, disclosure, transparency and balanced external reviews are presently the best pragmatic approach to managing cognitive biases.

Tit for tat, adversarial claims of bias in the scientific literature doubtfully advance the science. Conflicting perspectives can become personalized and intractable. How to know which is more credible? Neither? Both? Food nutrition researchers pointed out examples of selective data interpretations and publication bias in obesity research in relation to sweetened beverage (soft drink) consumption and in the health benefits of breast feeding. They termed this distortion of information to further what may be perceived to be righteous ends as "white hat bias" (Cope and Allison 2009). However, their conflicted financial backing from the soft drink industry and from manufacturers of baby formula contributed to counter-criticisms of funding bias (Bes-Rastrollo et al. 2014; Harris and Patrick 2011; Mandrioli et al. 2016). Unresolved in the claims and counter-claims of bias and financial conflicts of interest was what advice was most credible.

In environmental toxicology as well, controversies over the best interpretation of sometimes ambiguous facts can become entrenched and focused on the people holding differing views as much as the evidence behind the different views. Examples include deeply held and personalized disagreements over risks of atrazine to amphibians (Benderly 2014; Hayes 2004; Kintisch 2010; Raloff 2010; Rohr and McCoy 2010; Solomon et al. 2008); sufficiently safe levels of selenium for fish and birds (Renner 2005; Skorupa et al. 2004); and a dispute that was maintained for over 20 years about whether an oil spill resulted in indirect harm to salmon (Burton and Ward 2012). These intractable, mutual bias criticisms make it very difficult for non-specialist readers to make informed judgements of which is the more credible science.

Some particularly challenging situations in ecotoxicology

Some situations that seem particularly challenging for researcher and institutions to maintain scientific credibility warrant mention. Elliott (2014) argued that scientific findings that are ambiguous or require a good deal of interpretation or are difficult to establish in an obvious and straightforward manner are prone to bias, particularly if strong incentives to influence research findings in ways that damage the credibility of research are present. In environmental toxicology, risk assessments or critical reviews fit that test and can be vulnerable to bias. Suter and Cormier (2015a) identified sources of bias in ecological risk assessment as including personal bias, regulatory capture, advocacy assessment, biased stakeholder and peer review processes, preference for standard studies, inappropriate standards of proof, misinterpretation, and ambiguity. These challenges may lead to differences of opinion on methods for drawing

conclusions to support decision-making that, while prone to bias, have, at their root, the need for drawing conclusions in the face of uncertainty.

Costs of large-scale projects to remediate contaminated environments such as sediments contaminated by urban and industrial sources, aged industrial facilities, or large mining operations can be enormous, running to the hundreds of millions of dollars or much more (Gustavson et al. 2007; McKinley 2016). In "polluter-pays" schemes, the potential financial liability associated with such projects could imperil the ongoing viability of companies, which in turn would harm the livelihoods of employees, among other social disruptions. In such a setting, the scientists working on behalf of the those who may have to incur the costs of cleanup might understandably be more cautious about the potential for misguided remediation following Type I error (e.g., falsely discovering environmental degradation) than Type II error (failing to discover degradation when in fact it is occurring), when the science is ambiguous. Conversely, the regulatory scientists entrusted to provide scientific advice to protect environmental quality might be obliged to err on the side of precaution and be more accepting of risk of Type I error, especially when it is "other peoples" money at stake.

While science ethicists and the NAS (Boden and Ozonoff 2008; Elliott 2014; Krimsky 2005; NAS 1992) have emphasized industry funding bias risks, these risks are not unique to industry funding of science. For examples, many countries have provisions for natural resource damage assessment and restoration (NRDAR) to compensate the public for lost opportunities following shipwrecks, oil spills, releases of industrial chemicals, and so on (Boehm and Ginn 2013; Descamps 2008; Flamini et al. 2004; Goldsmith et al. 2014). These assessments rely on science to some degree to establish linkages from the release to harm to the environment. In turn, trustees of natural resources rely on science advisors to assess the extent and scale of injuries (adverse effects) and the monies needed to restore the lost services. In large incidents, the responsible parties will inevitably retain their own science advisors. The resolution of complex situations is resolved either by negotiation or adversarial litigation (Flamini et al. 2004; Goldsmith et al. 2014). This environment produces an atmosphere with strong incentives for plaintiff/trustee science advisors to maximize the magnitude and spatial extent of effects to the environment and to downplay uncertainties or the influence of potential other, non-compensable stressors and vice versa for those scientists retained to help defend against claims. Maintaining objectivity and advancing science in such a work environment would require extraordinary self-discipline by the individual scientists, an institutional environment emphasizing science credibility, and an openness to external, disinterested review (Boden and Ozonoff 2008; Elliott 2014; Wagner 2005).

While at least in some jurisdictions, monies from NRDARs must go to restoring the damaged public natural resources (beyond paying for salaries, consulting fees, and expenses to support

claims) and cannot be used to enrich those pursuing the cases. Toxic torts by comparison, pursue damages on behalf of private individuals or groups who consider themselves to have been harmed by exposures to toxic chemicals. Toxic tort cases are adversarial proceedings with the lawyers expected to advocate only for their client, and expert witnesses are paid to present testimony to support just one side. These torts may be highly lucrative for the plaintiff attorneys who select the science testimony. For example, in the Vioxx litigation the share for plaintiff lawyers was about \$1.5 billion (32%) of the \$4.85 billion settlement (McClellan 2008), and in successful asbestos litigation the average share of payouts going to the victims was only 37% (Elliott 1988). The lures and risks of such immense payouts in toxic torts create strong incentives for biased science. At best, critical reviews or product defense studies conducted for toxic tort science should be regarded with skepticism.

Defense of science and engineering in favor of protecting enterprises reflecting years of devoted work is understandable but becomes dangerous when objectivity is compromised. Case studies such as the Vioxx case, in which the maker of the drug downplayed increased risks of mortality from a successful product in which they were deeply vested (Curfman et al. 2005; McClellan 2008) and the cross-claims of blame between the engineering consultants and the mine operator in the aftermath of the Mount Polley mine tailing dam failure (Amnesty 2017; Topf 2016), remind us that objective science (including recognizing and disclosing uncertainty, and encouraging additional science to narrow that uncertainty) is good business.

Academic – Industry Collaborations

The role of industry funding and concerns of perceived conflicts of interest in academicindustry collaborations have been addressed in literature and are a common element in institutional research integrity policies (Elliott 2014; Resnik and Shamoo 2011). Often through philanthropic foundations, industry may contribute to basic science education and research to strengthen regional universities and further the science literacy of potential workforce and society. Industry may also support applied ecotoxicology and other environmental science research to inform specific scientific questions that affect their business interests. When industry and academic research interests become at least partially congruent, academic scientists may actively seek out such interest and support for their projects and graduate students. Pragmatically, academic-industry collaborations are necessary since public funding alone may be insufficient to support graduate research or to address important questions relevant to industry and society. In the US, about 40% of national research and development is funded by the private sector (NAS 2017). In the US, public funding for university research on the effects of chemicals in the environment has consistently declined since 2000 (Bernhardt et al. 2017; Burton et al. 2017), which implies that without industry-academic collaborations, there would be much less substantive university research. The need for sufficient funding to support training and research

can trump concerns over the color of money, as captured in a university president's quip, "the only problem with tainted research funding is there t'aint enough of it" (Krimsky 2003).

Benefits of collaboration run both ways, with expertise from academic and public sectors helping industry find solutions to lessen or avoid contributing to environmental problems (Hopkin 2006). In a recent tripartisan commentary promoting collaborative research among academia, business, and government in environmental toxicology and chemistry, Chapman et al. (2018) argued that collaborative connections across sectors provide scientific structural integrity and generally foster scientific integrity, transparency, and environmental relevance through balancing perspectives. Collaborations may enhance the broad acceptability of research findings, over those put forth from noncollaborative research. However, collaborative research is not without challenges. Foremost of these is a perception of bias and known collaborators may be subject to ad hominem criticisms, ridicule, and marginalization. Strong attention to objectivity, transparency, and sometimes a thick skin are needed for successful collaborative research (Chapman et al. 2018). More generally, Edwards (2016) lays out several principles for successful, durable industry-academic collaborations, including: establish clear quality criteria and make them public; mandate data sharing; subject work to independent oversight before public release; and enshrine public ownership for all research outputs. Further, effective collaboration between industry and academic scientists requires industry to provide expertise as well as funds. Collaboration with industry scientists engenders a shared desire to succeed and creates a sense of ownership of a project (Edwards 2016). The interchange of science through academic, industry, and government scientists is deeply rooted in SETAC culture, and the favorable views of the authors toward working across sectors is undoubtedly influenced through our history with SETAC. However, industry support to academics or others in support of applied environmental questions may come with inherent conflicts of interest, and critics may consider scientists as collaborators in the pejorative sense of the word (Hopkin 2006). This setting requires vigilance from both industrial research sponsors and recipients to avoid unconscious bias.

While readers might presume situations in which individuals or institutions with strong incentives to influence research findings consistent with their financial interests will do so, it is important not to judge a study solely by its funder, nor to presume the sponsor's preferred outcome. For example, an energy company sponsored a study to see if they could develop a scientific case for relief from costly requirements for meeting dissolved oxygen criteria in a river downstream of its hydroelectric dam. Instead the testing showed that the existing criteria could impair hatching salmon (Geist et al. 2006). The company scientists easily could have buried the results, which could have been discounted as being from novel techniques. Their path of least resistance would have been to leave the study in the file drawer, rather than going to the trouble

of defending novel science and publishing it in the open literature. In the long-view, a reputation of science credibility may be more valuable for companies than short-term project benefits.

Other examples include scientists from mining and metals trade groups publishing studies showing that existing USEPA criteria for zinc and other metals could be under-protective of aquatic species or entire communities (Brix et al. 2011; DeForest and Van Genderen 2012). Conversely, a university quantitative ecologist accepted support from an environmental advocacy group (through university channels) to model the potential population-level effects of elevated selenium from mining on local native trout populations (Van Kirk and Hill 2007). As the advocacy group had been a persistent opponent of the mining operations, officials from the influential mining company apparently presumed that the academics' work would also be biased to favor the advocacy group's positions, and they questioned the researchers' probity (Blumenstyk 2007). In fact, the selenium concentrations projected by these academics to cause detrimental population-level effects were higher than concentrations previously derived by industry-funded consultants who themselves had been on the receiving end of bias implications because they were industry funded (Skorupa et al. 2004; Van Kirk and Hill 2007). Unfortunately, these favorable collaboration examples are countered by examples in which studies were funded as part of deliberate strategies to shape the science to fit business interests. This "tobacco strategy" has been asserted with various substances such as asbestos, benzene, chromium, lead, vinyl chloride, and more (Anderson 2017; Cranor 2008; Krimsky 2003; Michaels 2008; Oreskes and Conway 2011; Sass et al. 2005).

In keeping with the adage to be careful judging a book by its cover or wine by its label, judging science by its funder or by presumed interests or leanings of the scientists can lead to mistaken and unfair perceptions. Brain et al. (2016) pointed out that the career path of environmental scientists is often ambiguous and whether scientists ended up in careers with industry, academic, or government science has more to do with chance and timing of opportunities rather than a particular desire to work in one sector or another. Such is often the case with academic and government scientists who work with industry to jointly fund or investigate a science question of mutual interest (Hopkin 2006). The convergence of scientific interests with financial interests can lead to a good marriage, so long as the parties are principled and forthright with each other. While there may be a perception that research contracts are highly restrictive, in our experiences these agreements establish expectations of academic freedoms. "Interested science" should be viewed with open-minded skepticism, and studies with immense financial implications warrant a higher level of scrutiny than others (Krumholz et al. 2007; Suter and Cormier 2015b; van Kolfschooten 2002). It does not necessarily follow that interested science is wrong or tainted. Ensuring transparency and complete data reporting is one tangible

step researchers can take to improve credibility of and perceptions toward industry-academic collaborations.

A scientific society founded on the principles of balancing competing interests

Scientific societies have important roles in promoting scientific integrity and ethical conduct, such as establishing codes of ethics which include disclosure of conflicts of interest, being a focal point for developing and communicating discipline-specific standards to foster research integrity, and providing educational material (AAAS 2000; NAS 2017).

We think the Society of Environmental Toxicology and Chemistry (SETAC) is notable for its directed and sustained efforts to balance competing perspectives in its deliberative processes and other activities. The founding principles and structure of SETAC sets out a tripartisan structure with regulatory, industrial, and academic scientists (Bui et al. 2004; Menzie and Smith 2018). As a result, SETAC now has well developed norms for balancing interests, inclusiveness of differing viewpoints, and neutrality in the reporting. These norms have enabled SETAC to be regarded as a source of consensus-based science with successful partnership or advisory roles in United Nations programs and conventions such as the United Nations Environment Programme's (UNEP) Global Mercury Partnership, Stockholm Convention on persistent organic pollutants, UNEP-SETAC Life Cycle Initiative for reducing hazardous waste as well as informing national-level legislation (Augspurger 2014; Mozur 2012). In contrast to multi-sector, non-profit organizations such as the Health and Environmental Science Institute (hesiglobal.org) which brings scientists together from academia, government, industry, and non-governmental advocacy organizations (NGOs) to conduct original research in the public domain, SETAC is more a forum for dialog and promotion of best practices within the discipline.

The intended balanced representation of industry, government, and academia isn't always achievable, for there are also guidelines for gender equity, geographic representation, and of course people have to be willing to volunteer. Further, the tripartisan emphasis underrepresents scientists from environmental advocacy groups or other NGOs. These groups are influential for shaping public debate, policy and law on environmental issues, but their low participation in the Society suggests that they may not be attracted to or feel welcomed by a "hard" scientific society such as SETAC, and meeting costs may be a barrier. Despite these imperfections, the norms of seeking to balance potentially conflicting interests and to provide a safe forum to express differing scientific viewpoints are deeply ingrained in the Society's culture and activities.

Promoting scientific integrity in ecotoxicology

While "scientific integrity" is ultimately a subjective judgment that cannot easily be reduced to review checklists, there are some general points to maintain in ecotoxicology and related science. These include relevance, rigor, reproducibility, objectivity, and transparency.

Environmental Relevance

By definition, environmental chemistry and ecotoxicology are concerned with how chemicals, both natural and synthetic, pose a threat or influence the natural world (Johnson et al. 2017). Because of pragmatic and ethical constraints, research in this domain is often done in laboratory environments, testing cultured laboratory organisms or cell lines or other in vitro surrogates for organisms. However, the intent of such research invariably still has some intended relevance to conditions that occur in the environment. We have seen articles in ecotoxicology literature discussing some novel research based on under-tested taxa, underappreciated endpoints, unexpected multiple stressor effects, or unanticipated indirect effects via untested commensal microbes. An article may start out with an introduction on the ecological importance of the novel work, the work is reported, and then the discussion closes arguing that ecological importance of their work, how it should change the thinking in the field, and management implications. Yet to obtain their desired experimental effects, exposure concentrations may have been orders of magnitude higher than those typical in the real world, or exposure routes, chemical forms, or dilution media may be unlike those that the organisms could encounter in nature (Johnson and Sumpter 2016; Mebane and Meyer 2016; Weltje and Sumpter 2017). When authors present such studies with a narrative on the ecological importance of their topic, this may be a form of misrepresentation.

Environmental relevance and regulatory relevance may not always be one and the same. Still with studies of potential ecological effects of chemicals, investigators often hope that their research will inform future regulatory assessments of risk and safety. In practice, environmental regulators may pass over results of academic studies in favor of research sponsored by industry. Steps academic ecotoxicologists could take to improve the utility of their research for informing policy include developing an understanding of environmental regulatory frameworks, using existing chemical assessments to inform new studies, conducting and reporting studies to include sufficient rigor, quality assurance, and detail to enable regulatory use, and placing academic studies in a regulatory context (Ågerstrand et al. 2017).

Rigor

Funders, journals, and institutions reward novelty, such as the short-lived discovery of a bacterium that grows with arsenic instead of phosphorus (Alberts 2012). Highly selective journals with article acceptance rates of 10% or less preferentially publish findings that are sensational or at least surprising. These incentives are influential because universities and research institutes often hire and promote scientists based on their record of acquiring grant money and the number of publications times the journal impact factors of the journals published therein (Parker et al. 2016). With finite career opportunities and high network connectivity, the marginal return for being in the top tier of publications may be orders of magnitude higher than

an otherwise respectable publication record (Smaldino and McElreath 2016). The editorial quest for novelty has led to publication of questionable articles in elite journals, such as one positing that caterpillars were the results of accidental sex between insects and worms (Borrell 2009). Top tier journals also tend to have higher retraction rates than mid-tier journals, suggesting that rigor has sometimes been compromised in the competition for paradigm shifting results (Nature Editors 2014).

In ecotoxicology, Harris et al (2014) describe 12 basic principles of sound ecotoxicology that should apply to most environmental toxicity studies. These principles range from carefully considering essential aspects of experimental design through to accurately defining the exposure, adequate replication, unbiased analysis and reporting of the results, and repeating experiments that yielded surprising or ambiguous responses. There are ample opportunities for improvement. For example, Harris and Sumpter (2015) asked a very basic question of a sample of studies published in 2013 in three leading ecotoxicological publications: was the concentration of the test chemical actually measured? Of the studies reviewed from *Environmental Toxicology and Chemistry*, 20% failed this basic aspect of experimental credibility, as did 33% and 41% of ecotoxicology studies published in *Aquatic Toxicology*, and *Environmental Science and Technology*, respectively (Harris and Sumpter 2015).

While Harris et al. (2014) emphasized laboratory-based studies, field-based environmental effects studies replace the challenges of the artificiality and questionable relevance of some laboratory-based toxicity testing, with different, messy, real world challenges. Closely related to the 12 principles described by Harris et al, we suggest 8 basic principles relevant to most field-based ecotoxicological studies or environmental effects monitoring.

- 1. Development of a thorough understanding of the issues to ask informed questions. The available literature should be thoroughly vetted to inform the need for experimentation or field studies;
- 2. A thorough understanding of the questions being posed is an essential prerequisite for designing a robust, reliable, reproducible study. Incomplete understanding of the questions leads to vague and often misguided study that undermines the scientific process (Lindenmayer and Likens 2010; Suter et al. 2002);
- 3. The ability to identify and reliably measure sensitive indicators (Melvin et al. 2009),
- 4. Careful attention to appropriate reference conditions to avoid potential, actual effects being masked by variability or confounding factors introduced by differences between the reference and test site environments (Arciszewski and Munkittrick 2015; Mebane et al. 2015). For example, beaches on rocky headlands and protected bays will have very different benthic invertebrate communities, as do flowing rivers and impounded reservoirs. Study designs that attempt to detect pollution effects on communities across such disparate habitats may have very low discriminatory power and by failing to account for natural variability, adverse pollutant effects could be obscured (Buys et al. 2015; Parker and Wiens 2005; Wiens and Parker 1995);

- 5. Try to study a number of locations that vary in the degree of the factor under investigation, such as chemical pollution, in order to accurately estimate a relationship between exposure to the environmental factor of interest and the effect of that factor, if such a relationship exists.
 6. Time and patience: Just as experimental exposures need to be of appropriate duration for effects of interest to be manifested, environmental monitoring needs to be maintained long enough to pick up true trends if present, or to convincingly argue that trends are not present (Lindenmayer and Likens 2010).
 - 7. Specific definitions of what effects are considered negligible or of concern (Melvin et al. 2009; Munkittrick et al. 2009; Power et al. 1995).
 - 8. Avoid power failures: use a statistical approach appropriate to the question, considering statistical burden of proof issues. For instance, P>0.05 in testing for trends or differences between locations does not by itself show the lack of trend or effects (Dixon and Pechmann 2005; Mudge et al. 2012).
 - 9. Transparent reporting with detailed methods and raw data sufficient for others to reproduce the analyses or to further examine the data using alternative analyses (Duke and Porter 2013; McNutt et al. 2016; Schäfer et al. 2013).

Reproducibility

Reproducibility is one indicator of reliable research. However, the inability of researchers to reproduce influential studies of others or their own has garnered enough attention to be called a "reproducibility crisis" (Baker 2016a; Henderson and Thomson 2017; Munafò et al. 2017). However, not all studies are easily reproduced. Environmental data are often messy and field studies are more often observational than experimental. Large scale, ecologically realistic studies such as long-term, experimental lake studies difficult to do even once; and (hopefully) no one wishes for mishaps such as tailings dam failures or oil spills to study (Parker and Wiens 2005; Schindler 1998; Wiens and Parker 1995). Such studies require a logical system for causal inference to separate cause-and-effect from serendipitous correlations (Norton et al. 2002; Suter et al. 2002).

Even rigorous laboratory studies may be difficult to replicate due to the highly variable nature of biological systems and unanticipated responses to unknown factors. Demands for reproducibility may favor industrial science over academic science. Industry often works within strict Good Laboratory Practice (GLP) rules and with well-studied species tested through standardized protocols (Elliott 2016). Academic science is often framed around education, and grants and graduate student researchers are usually encouraged to go after something new and novel; protocols may be developed as they go, and quality control may be uneven (Baker 2016b). Obstacles to adopting formalized quality management systems such as GLP in small research settings may include costs, lack of resources, lack of mandate, independent cultures, and high turnover (Figure 3). Adherence to GLP does not therefore mean that a study is well-designed. It only means that a study is well-documented, quality assured, and that the study design protocol

was expressly followed. But if the study design wasn't appropriately relevant and rigorous for the study questions, the well documented and reproducible results will doubtfully be meaningful. Nevertheless, even if regulatory GLP compliance is not required, small academic research facilities can benefit from embracing core components of GLPs, such as defining responsibilities, maintenance and sanitation of common lab spaces, equipment and materials, well defined experimental protocols, quality control testing including positive and negative controls, data reviews, audits, and archiving (Bornstein-Forst 2017; Moermond et al. 2016).

Better experimental protocols that are easier to follow are one tangible way to strive for better reproducibility and transferability of both novel and standard experimental methods (Figure 4). Multimedia experimental protocols could be much easier to explain and teach techniques than the conventional, densely worded, printed protocols. The Journal of Visualized Experiments (JoVE) is an innovative peer reviewed, science methods journal in which its articles are a unique blend of the conventional printed article with professionally produced videography. Ecotoxicology methods articles have begun to be published in this format (Calfee et al. 2016; van Iersel et al. 2014). The field would benefit from broader use of new visualization techniques to document new methods and to improve education and training on techniques that need to be highly standardized to be repeatable. At the minimum, with the availability of electronic data repositories and supplemental information in journals, there is no reason why detailed methods including video demonstrations cannot be published.

Reproducing a statistical summary or model run reported in a scientific publication when the underlying data and code are provided and explained is one thing. Reproducing an actual complex experiment is hard and is rarely attempted, unless perhaps the results are novel and have a high regulatory or societal impact. Even under the best of circumstances, such as when the original investigators have the resources to and are motivated to repeat an experiment in the same lab, with organisms from the same culture, using as close to identical methods as they could manage, and including positive controls, the investigators may be unable to produce the same result twice (Mebane et al. 2008; Owen et al. 2010). Positive controls (testing a substance with well characterized effects) are not always used in toxicity testing programs, but their routine use can help investigators understand variability in test results (Glass 2018). Nosek and Errington (2017) caution that if investigator #2 reports that the results of study #1 could not be reproduced, that by itself does not indicate which is more credible: result #1, #2, neither, or both. Further, much of the "reproducibility" debate in the natural sciences is focused on cell biology or human behavior (psychology) experiments, which may be more tractable to reproducibility studies than messy environmental observational or experimental studies. Especially with complex biological testing such as multi-generation tests, a green thumb husbandry factor may bring together art and science to environmental chemistry and toxicology (Figure 4). Subtle

methods differences, strain differences or stochastic events can be so puzzling that investigators are left thinking demons must have snuck into their study and interfered with one treatment but not others (Hurlbert 1984). (We presume that Hurlbert's (1984) suggestions for exorcisms or human sacrifice for troubleshooting suspected demonic intrusions, might run afoul of some contemporary institutional review board policies.)

Still, reproducibility is a core tenet of science and successful reproduction adds confidence in the credibility of novel findings. Divergent but individually credible results may further advance the science by illuminating important aspects missed in the initial study (Owen et al. 2010). If for instance, an investigator were to find a novel, major adverse effect of a class of chemicals to a previously untested taxonomic group, then other equally diligent and skilled investigators should be able produce similar effects in other research settings, even if the test conditions were only similar. For instance, a standalone paper from the 1970s that a snail was anomalously sensitive to Pb was regarded skeptically. Over 30 years later, this open-minded skepticism led to follow-on studies from a new generation of scientists that not only affirmed the anomalous early report of sensitivity but also led to important advances in comparative physiology and underlying mechanisms of toxicity (Brix et al. 2012). Similarly, early reports that freshwater mussels and other mollusks were unusually sensitive to ammonia were not widely persuasive. After repeated studies across multiple laboratories and species showed similar findings, the issue gained traction with standardized method development, inter-laboratory round robin testing, and attention by environmental managers (Farris and Hassel 2006; USEPA 2013).

Individual investigators may not always have the opportunities for self-replication, but best practices call for repeating what one can (Harris and Sumpter 2015). In field studies, multiple measures of exposure, multiple years of field data, and so on give credence to findings. We recognize that all science has practical resource limits and we are not going as far as arguing that novel findings from small-sample studies should never be published. Rather, the appropriate conclusion from such studies is along the lines of "if these findings turn out to be repeatable, they could be an important development." In our view, novel, major findings that are supported only by a one-off study are best regarded as tentative.

Transparency

Transparency in reporting research, including all the relevant underlying data that were relied upon in the paper, has become a critical element of integrity in science. Science's claim to self-correction and overall reliability is based on the ability of researchers to replicate the results of published studies (Nosek and 39 co-authors 2015). Studies cannot be replicated or even reconstructed if scientists will not share additional data, information, or materials from published studies, and we believe that upholding such ethical norms is every scientist's responsibility. The

embrace of the principle of transparent reporting has been uneven across disciplines, and the field of ecotoxicology has certainly not distinguished itself as a leader in this regard (McNutt et al. 2016; Meyer and Francisco 2013; Parker et al. 2016; Schäfer et al. 2013; Womack 2015).

Researchers in ecotoxicology and environmental chemistry have long only presented highly reduced data summaries. The only "data" included in some publications are crowded figures and tables with results of statistical outputs, such as F- values, effects concentration point estimates (EC50, EC10, etc.), or no-and lowest-observed effects concentrations (NOECs, LOECs). These derived values are not data. Such data-poor publications were understandable before the early 2000s, in which strict page limits and word limits precluded authors "wasting" space publishing data tables. With the provisions for electronic supplemental material beginning in the 2000s, and dedicated data repositories becoming widely available at low or no costs to authors in the 2010s, these reasons for opaque publication are no longer justified. Researchers who choose not to transparently report the actual data underlying their scientific findings may have other reasons for doing so. They may be concerned about others scooping them on their own data (McNutt 2016), although counterintuitively, publishing data may actually help establish priority and reduce scooping concerns (Laine 2017). Other less charitable reasons why researchers might resist publishing data include that they haven't devoted the needed time to organize their data in a coherent fashion that is interpretable by others, because reported results might not be able to be reconstructed from the underlying data, they are not keen to facilitate alternate statistical analyses or interpretations of their data, that they wish to publish unfalsifiable findings, or because there's simply less there than they led readers to believe (Smith and Roberts 2016).

Data sharing may still be regarded more as an imposition from science funders to be complied with rather than as a universal principle embraced by those conducting and publishing scientific research (Burwell et al. 2013; Collins and Verdier 2017; European Commission 2016; Holdren 2013; Nelson 2009; Nosek and 39 co-authors 2015). There are many pragmatic obstacles to effective data sharing, such as the expertise, extra work, and costs to researchers to organize, serve, and preserve their data in a comprehensible manner, privacy and anonymity concerns for environmental data collected from private property, about human subjects, and balancing intellectual property concerns. Some environmental science research is intended to be confidential, such as private sector economic geology, agricultural chemical product develop products and recoup research and development investments. However, in our view, researchers on such ventures cannot have it both ways, by publishing some outcomes in the peer reviewed literature, but withholding the supporting data as private. A recent corporate initiative to make available traditionally protected crop safety information is noteworthy in this regard (https://cropscience-transparency.bayer.com/).

Most environmental science journals have policies encouraging and facilitating data sharing. SETAC journals are probably typical in requiring a statement by the authors' whether and how the data underlying their analyses are available, with an admonition that authors should share upon request. A passable statement may be something as feeble as "*Contact the Corresponding Author for data availability*."

The strongest data disclosure policy for journals publishing in the environmental sciences is probably that developed for the Public Library of Science (PLoS) family of journals. "*PLoS journals require authors to make all data underlying the findings described in their manuscript fully available without restriction, with rare exception*" (PLOS 2014). Exceptions are limited to privacy or vulnerability concerns such as data on human research subjects that could not be fully anonymized, locations of archeological, fossil, or endangered species, that could be exploited or damaged, or safety and security considerations. Penalties for authors who fail to comply include rejection, or if they decline to provide data for an already published article, the editors could flag their article with a cautionary correction or even retract it (PLOS 2014). Whether PLoS's stand requiring authors to make available all data underlying their findings will lead other journals to stiffen their resolve, or whether the comparatively lax policies of competing journals will undermine PLoS and other open-science advocates remains to be seen (Davis 2016; Nosek and 39 co-authors 2015).

Implicit to such requirements is the assumption that common understandings of what constitutes "raw data" will be contextual. For instance in toxicity tests, usually the counts or measurements at each time interval, and all associated chemical and physical measurements are considered "raw data." Metrics of species counts, average treatment concentrations, average of replicates and such are not raw data, they are derived values. Some "data points" such as a streamflow measurement or a chemical concentration in a medium are actually derived values, and the true raw data behind a data point includes survey data, unprocessed sensor readings, spectral outputs, and such. Unless the study involves methods comparisons or forensic data audits, usually the researcher just wants the resultant derived values at a level of detail sufficient to reconstruct and further analyze the original detail.

While the notion that investigators should preserve and share underlying data is simple, the reality of doing so is much more complicated and challenging. To us, it is a priority to strongly encourage, for without data, the credibility of science cannot be evaluated. Some research has shown the willingness and ability for authors to share data declines significantly with time, and having a weak data availability policy is only marginally better than having no policy at all (Vines et al. 2014). Computer servers get replaced, directories flushed, offices moved, files dumped, and investigators move on, retire, and eventually die.

Rather than mandates, one simple incentive to improving openness in reporting has been for journals to award prominent open data "badges" for articles verified as being supported by available, correct, usable, and complete data. By showing an open data badge on the issue table of contents, article web page, and including a "verified open data" statement in the bibliographic indexing metadata, articles without such badge endorsement may be seen as incomplete. Over time, this might shift the norm toward open preservation and sharing. In at least one journal, this approach appeared to markedly improve the sharing and preservation of data through linked, independent repositories (Kidwell et al. 2016; Munafò et al. 2017).

Critical Reviews and Literature Syntheses

In ecotoxicology, published literature can roughly be broken down into two categories: original research and the review article. The original research article usually is based upon field observations, laboratory experiments, modelling, or blended approaches. Generalizing original articles through reviews and syntheses are critical parts of the ecotoxicology and most environmental science literature. Critical reviews, risk assessments, environmental quality standards, are based on syntheses of the literature, and not on individual studies. Synthesis articles have rather distinct scientific integrity problems from the original research article. Decisions must be made on how studies were located, results categorized, and a host of data reduction, data standardization, and analyses decisions need to be made. These decisions and associated biases may be deliberate and clearly explained or the analyst may not even recognize that they have made a decision (Roberts et al. 2006; Suter and Cormier 2015a). In some cases we suspect analysts obscured their decisions. Systematic review methodology is now being used also for some chemical assessments in which case data synthesis may be highly structured, with criteria clearly defined for data inclusion and search strategies (Hobbs et al. 2005; Van Der Kraak et al. 2014; Whaley et al. 2016). Other situations may follow the wending path of the present article: discussions among the authors "have you seen so-and-so?", and readings that led to other relevant material through forward and backward citing, along with by some specific subject searches. This path led to much relevant and thoughtful material across many disciplines. But it was hardly systematic or reproducible.

Literature searches from different sources can yield very different results. For example, using a 2007 original research article on population modeling of selenium toxicity to trout (Van Kirk and Hill 2007), four leading bibliographic indexing services were searched for articles citing that study. Web of Science (WoS), Elsevier's Scopus, Digital Science's Dimensions, and Google Scholar found 7, 10, 15, and 22 citing publications respectively. Scopus found all articles found by WoS, plus articles WoS missed in *Human and Ecological Risk Assessment* and *IEAM*. Google Scholar found all articles found by Scopus and WoS, plus articles in *Ecotoxicology Modeling*, *Water Resources Research*, 3 government reports, 2 books, a thesis, a conference proceeding, a

duplicate, and 2 ambiguous citations. It follows from this 3-fold difference in valid citations that a critical review of published literature on a topic or a regulatory assessment could miss relevant science if the assessors relied too heavily on a single search provider.

This simple example was from the "current era" of science, which began by 1996 or so, depending on which bibliographic indexing service scholars are using. Web sites for WoS and Scopus respectively report their indexing databases are reliable from 1971 and 1996 forward. Relying exclusively on bibliographic index searching may omit important, relevant older research.

Thus, we have the indexing bias problem in meta-analyses and assessment (that not indexed won't be retrieved), and the related problem of reviewing the secondary source but citing the original. We have seen assessments that omitted seminal research published before the current digital era, which may reflect indexing bias. Ecotoxicology syntheses often rely on variations of species-sensitivity distributions, which may provide more explanations of statistical characteristics of the datasets, data extrapolations, transformations, normalizations, than on where the data came from in the first place. We have seen micrograms and milligrams mixed up, and rankings that mistakenly commingled endpoints such as time to death in hours with effects concentrations. Some of these issues are undoubtedly related to the online availability of well curated databases such as ECETOC Aquatic Toxicity (EAT) Database from the European Center for Ecotoxicology and Toxicology of Chemicals or the U.S. Environmental Protection Agency's EcoTox databases. These compiled databases are valuable resources but reliance on secondary, compilations deprive the original authors of credit via citations. At least for publicly funded science, citations may be a way that authors demonstrate the value of their work to the scientific community, and thus build the case for further funding. Further, reliance on secondary sources is a good way to introduce or repeat inaccuracies (Rekdal 2014). We echo previous calls for better training and rigor when conducting and reporting secondary analyses of ecotoxicology and related literature. Practices from other fields, such as the Cochrane systematic review approach and guidelines for the ethical reuse of data could be adapted to the ecotoxicology practice (Duke and Porter 2013; Roberts et al. 2006; Suter and Cormier 2015a).

Objectivity

Beyond the topics discussed, some explicit steps to ensure objectivity bear consideration from initial planning through the interpretation of results. Even very careful scientists are not immune from cognitive biases that affect objectivity. These include hypothesis dependency (alternatives never considered will not be evaluated), hypothesis tenacity (sticking with an initial hypothesis despite evidence to the contrary), and anchoring, where beliefs are excessively influenced by initial perceptions (Norton et al. 2003).

There some steps scientists could consciously take to avoid these pitfalls. Strive to be selfskeptical of possible personal predilections toward expected causative factors or outcomes. Consider the opposite – maybe it is correct. For topics with competing schools of thought, consider how a scientist from a different school might interpret the data; or how might one from a different sector analyze the problem (Suter and Cormier 2015a)? If considerations such as these are sincere and not just done to anticipate and refute criticisms, they can help to ensure the objectivity and balance of interpretations.

Environmental Chemistry

We discuss environmental chemistry separately because it has different scientific integrity challenges than the biological side of ecotoxicology. Unlike the situation in the biological side of ecotoxicology where serious questions have been aired about the reproducibility of some of the published research (Scott 2012, 2018; Sumpter et al. 2014; Sumpter et al. 2016), analytical environmental chemistry does not appear to suffer from such problems to the same extent. The likely reason for this is that quality assurance mechanisms are routinely incorporated into analytical projects involving the measurement of environmental concentrations of chemicals, thus ensuring that the results are accurate. These include the use of high quality standards, which are widely available, the use of high specification instruments, and general guidelines proposed by national and international institutions. That combination enables recovery rates to be determined, preferably at different concentration ranges, for intra- and inter-day precision to be assessed, detection limits to be quantified, and matrix effects (interference from other substances) to be investigated. These quality assurance procedures are adopted routinely, are always checked by reviewers of analytical papers, and ensure quality is maintained.

Unlike method development however, the reporting and interpretation of environmental chemistry has common pitfalls, particularly in analyses from large datasets or compiled databases, and in citation practices. For example, metadata specifying fundamental details may be missing or misunderstood, such as whether concentrations of metals or other elements in water are from filtered or unfiltered samples or if they reflect the total mass of the element or only one speciation state (Sprague et al. 2017). Aquatic metals concentrations declined from mg/L levels in reports from the 1980s to μ g/L or sub μ g/L levels by the late 1990s. This remarkable, widespread decline was not due to better pollution controls or global geochemical change, but to improved recognition and control of ubiquitous contamination in field and laboratory sampling and analysis methods. There are ongoing debates over the most appropriate sampling and analysis methods for inorganic water quality constituents particularly for environments that are expensive and difficult to sample representatively, such as large rivers. (Horowitz 2013). Such sampling biases and analytical method differences may be substantial enough to confound analyses.

Organic environmental chemistry datasets have similar pitfalls that can confound subsequent reviews and secondary analyses. For example, Kolpin et al (2002a) published a summary of a survey of different pharmaceuticals, hormones and other organic contaminants from 139 streams. This highly influential paper showed that some organic contaminants were widespread in streams and contributed to heightened concern and research interest in their potential health and environmental risks. The paper is presently the most highly cited paper ever published in the journal *Environmental Science & Technology* (1st of 46,011 papers published, with 5,104 citations in the Scopus database as of 16 August 2018). However, reported concentrations of at least 1 of the 95 chemicals reported, 17α -ethinyl estradiol (EE2), were questioned because the median and maximum concentrations of 73 and 831 ng/L, respectively were about 10 to 1000 times higher than those from other surveys or analyses (Ericson et al. 2002; Hannah et al. 2009). Kolpin et al. (2002b) responded that upon further inspection, they had discovered that their maximum reported EE2 value of 831 ng/L was indeed incorrect owing to analytical interferences. They further explained that they had defined "median" in a peculiar way, as the median of detected values in streams, not in its usual meaning as a central tendency of all values. Because EE2 was undetectable in 94% of streams sampled, the median of detected values was skewed far above the median of all streams (<5 ng/L). However, despite the discovery of the mistakes, no correction was issued for the original publication. The Kolpin et al. (2002b) acknowledgment of the mistaken values was buried among the other 5,103 citing papers and the subtle, peculiar definition of a "median" was likely overlooked by most readers. As of August 2018, at least 50 citing papers were identified that re-reported and perpetuated the exorbitant and mistaken 831 ng/L maximum EE2 value for U.S. streams.

Thus, it is easy for authors to easily misinterpret or to perpetuate erroneous relevant values from the literature. The problem of citing unreliable maximum values would be avoided if authors simply cited extreme statistics, such as percentile concentrations (e.g., the 10th to 90th, 5th to 95th, 1st to 99th) instead of ranges (Weltje and Sumpter 2017). Whereas a single extreme value defines the range, extreme percentiles are more representative of severe conditions that organisms may actually encounter and will be more stable and are far less vulnerable to be mistaken. For instance, Santore et al. (2018, their figure 9) elegantly summarized about 29,000 paired reports from aggregated data sources of dissolved and total aluminum (Al) in freshwater. Logically, the dissolved fraction of trace metals in water can be no greater than the total. In practice, results don't always come out that way especially when the two values are close. Factors such as differences in sample digestion, differences between instruments, or slight differences in technique may introduce subtle analytical biases that produce the dissolved fraction greater than the whole (Paul et al. 2016). In the Santore et al. (2018) comparison, at least 150 of the 29,000 Al pairs show the dissolved fraction is greater than the whole. While such logically impossible values should usually simply indicate that close to 100% of the total Al was

present in dissolved form, some are obviously impossible values with the dissolved fraction 2-3 orders of magnitude higher than the total concentration. Should an imprudent analyst uncritically report on ranges of dissolved and total Al, they would report nonsense results. Simply backing off to the 99th or 95th percentile for a large dataset such as this one would still reflect the extremes of environmental conditions but be somewhat insulated from dubious, single values.

The counterpart to avoiding pitfalls working with big data is avoiding pitfalls when working with small datasets. Small datasets may be the best that can be acquired from studies in extreme environments where sampling is difficult, dangerous, or very expensive or where conditions are ephemeral or rare. Dismissing high-quality data that may have been collected under heroic or unrepeatable circumstances on pedantic statistical grounds would be foolhardy. Small datasets can be important, so long as one is cautious about inferences. Efforts to "improve" small datasets through sophisticated statistical techniques should be resisted (Murtaugh 2007).

Advocacy

Science is the enterprise for answering questions and making predictions about the how the universe works. Science can inform issues, but science can never answer "should" questions. For example, science cannot tell societies whether they should restrict chemical uses and releases, whether natural preserves should be set aside from human exploitation, or whether biodiversity should be protected. These are among the myriad value judgements that societies must make, and while science can support societies in making these choices through predictions founded upon a body of knowledge, there are never "scientifically correct" answers to questions of human values, morals, and ethics (Snyder and Hooper-Bui 2018). Scientists are humans, and like all people, hold ethical and moral values which drive assumptions which may not be explicitly stated if even recognized. For example, the notion of "environmental protection" in the environmental toxicology field is rooted in societal norms, statues, and international agreements with goals of minimizing harm (a human concept) from activities such as extraction, manufacture, use, and disposal of chemical products. Scientists in the field develop informed opinions toward the "should" questions relating to their experiences, which leads to questions of whether and how scientists advocate for "should" questions.

The underpinnings of science are that researchers have no vested interest in the results of their observations, that they objectively record and analyze these results, and that they fairly report the outcomes in the peer-reviewed literature. Advocacy can compromise these underpinnings, at the cost of scientists' credibility (Fenn and Milton 1997). Scientists tend to be passionate about their science, which has led to controversy over the role that scientists should play in related public policy debates. While we think most scientists would agree that advocacy for science having a role in environmental policy debates is appropriate, there is likely much less agreement whether

it is appropriate for scientists to advocate for particular outcomes in policy debates. If the policy debate turns on questions of science central to a scientist's particular area of study, probably no one is better positioned than that scientist to lay out the evidence for or against a particular course of action. If the scientist is regarded as a neutral and informed voice, their advice may be valued by all sides in a policy dispute (Sedlak 2016). However, if the scientist's experience or analyses leads them to the strong conviction that one policy direction is more correct and should be adopted, then they are no longer a neutral broker and have become an advocate.

When questions of science are central to adversarial adjudicated proceedings, the protagonists controlling the proceedings are often lawyers. The lawyers are expected to advocate for their client's interest; not for objective science. The lawyers retain consulting scientists as expert witnesses to support their side of the case. The lawyers' will presumably seek out scientists whose research findings and views will increase their chance of winning. In the close, intense working environment of a team preparing for a complex, science-based legal strategy, it is easy for scientists to get caught up in the enthusiasm of a "team spirit" with a loss of detached impartiality and objectivity. Scientists who begin to function as "hired guns" focused on team wins are no longer scientists but advocates (Christensen and Klauda 1988).

Policy advocacy is potentially problematic because it may compromise use of research findings in policy and management deliberations if the information is not viewed as credible by all sides (Scott et al. 2007). In some situations, advocacy is beyond reproach, such as a university scientist who uncovered a lead poisoned community water system. Simply reporting the findings to the responsible officials likely would have been ineffective, if the ineptitude or indifference of those same responsible officials contributed to the situation in the first place (Sedlak 2016). However, not all situations are so clear cut, and reasonable people who share similar motivations, skills, and agree that researchers should do the right thing may not agree on what that is. Deliberations on major environmental issues are complex and science may only be one element of the deliberations. Developing and providing technical and scientific information to inform policy deliberations in an objective and relevant way is a formidable challenge (Meyer et al. 2010; Nelson and Vucetich 2009).

Institutional constraints aside, how scientists balance these competing issues and choose when or whether to engage in advocacy is a deeply personal choice and is situational (Meyer et al. 2010; Sedlak 2016). However, just as science journals discourage comingling original research results and commentary, scientists should keep science and advocacy distinct in their publications and speaking. In particular, we argue that scientists should be watchful for stealth policy advocacy. Stealth advocacy is the use of value-laden language in scientific writing that assumes a policy preference (Lackey 2007; Pielke 2007). Rather than openly disclosing assumed values or policy preferences, biases may be unconsciously (or deliberately) cloaked through

normative science. Normative science is science developed, presented, or interpreted all based on an assumed, usually unstated, preference for a particular policy or class of policy choices. This covert advocacy may be reflected in word choices, and such advocacy is not always apparent even to the advocate. For instance, value-laden words such as *stressors, impacted, degraded, improved, good,* and *poor* may be used to describe habitats or other environmental features. Less value-laden words would be *factors, exposed, altered, changed, increased,* or *decreased.* The use of normative science is potentially insidious because the tacit, usually unstated, preference for a particular policy or class of policy choices is not perceptibly normative to policy makers or even to many scientists (Lackey 2007).

Criticisms of normative science can be excessive, as taken literally, the entire discipline of conservation biology could be considered too normative. Similarly, the mission statement of SETAC "to support the development of principles and practices for protection, enhancement and management of sustainable environmental quality and ecosystem integrity" could be too much for some. Science is normative, with topics and study questions influenced by normative treaties and regulations. Areas of study or techniques once considered appropriate areas of science inquiry such as craniometry, eugenics, or experimentation on human subjects without informed consent are no longer considered to be within the norms of ethical science. Within environmental toxicology, pressure to reduce the use of animal testing might be an example of normative science.

Our point is not to argue for or against scientists engaging in overt policy advocacy, which is a personal decision, but for clarity and transparency. Just as original results, opinion, judgements and speculation should not be blended in a scientific paper, science and advocacy need some separation (Scott and Rachlow 2011). Covert advocacy is a form of bias. Environmental scientists should clearly differentiate between research findings and policy advocacy based upon those findings.

Weaponizing scientific integrity and transparency

We recognize that "scientific integrity" discussions can easily be diminished to going down the path carved by "sound science" strategic initiatives, which often boiled down to campaigns to call "*my science good science and your science junk*" (Doremus 2007; Kapustka 2016; McGarity 2003; Oreskes and Conway 2011). The goal may be to recast policy, ideological, or economic disputes as doubt or created conflicts in science. In countries with a tort-based, adversarial legal system for resolving injuries or damages, science-based information becomes just another tool for dueling experts, who often have primary responsibility for advocating for the interests of their client (Wagner 2005). Research integrity policies or requirements for data transparency can be used as weapons to bury public university or government scientists with vexatious, intrusive,

and costly demands for records such as raw laboratory notebooks, instrument calibration records, emails between coauthors, working drafts, and peer comments and responses. Such demands can be effective tools for interfering with the work of public-sector scientists, including academics in public institutions (Folta 2015; Halpern and Mann 2015; Kloor 2015; Kollipara 2015; Lewandowsky and Bishop 2016), or academics in private institutions but who receive research support from public sources (Hey and Chalmers 2010; Shrader-Frechette 2012). For example, Deborah Swackhamer, an environmental chemist at the University of Minnesota, was targeted under state open records laws with legal demands for raw unpublished data, class notes, purchase records, telephone records, and more from a 15-year period. Ironically, the identities of those seeking the information were themselves shielded from disclosure (Halpern 2015). Some scientists have learned to use transparency laws against their peers in the highly competitive arena of grant funding. Through freedom of information demands for competing grant proposals, scientists have been able to obtain details on competitors' new research direction, preliminary results, and cost structure. For those targeted scientists, such information gathering may be seen as research espionage under the rubric of transparency (Carey and Woodward 2017).

The sunshine laws enacted in many jurisdictions were intended to illuminate the business of government officials and were doubtfully intended by their crafters to sweep up university professors. Nevertheless, some see scientists are fair targets of such tactics, as inspections of their erstwhile private communications have uncovered peer review misconduct, undisclosed conflicts of interest, or bias (e.g., Fellner 2018; Krimsky and Gillam 2018; Russell et al. 2010). Privately funded research is generally shielded from such practices (Brain et al. 2016; Wagner and Michaels 2004). However, researchers at private institutions may also be subject to baseless litigation to intimidate scientists and deter others by inflicting long and costly legal processes, disruption, and threats of personal financial liability. Such harassing lawsuits have been employed often enough to get a name, SLAPP suits for Strategic Litigation Against Public Participation (Johnson 2007; Nature Medicine Editors 2017; Robbins 2017). While legal, such strategies represent detrimental practices cloaked in the vernacular of transparent science (Johnson 2007; Levy and Johns 2016; McGarity and Wagner 2008; Wagner and Michaels 2004).

Education for a culture of science integrity

It is one thing to realize that there is a problem, but quite another to find effective solutions to that problem. For high scientific integrity to be the norm, the culture of science has to systematically embrace exemplary practices and discourage bad behavior (Benderly 2010). However sometimes the system and its incentives are part of the problem. Many of the reliability concerns and detrimental behaviors we have discussed are related to the perverse incentives under which scientists may operate. These incentives are grants and publications. In the case of grants, the more, and bigger, they are, the better, as far as institutions are concerned. In the case

of publications, the number of these seems to be much more important than their quality. This is probably because assessing the quality of a scientific article is not easy; there is no established, widely accepted way of doing this. The 'status' of the journal in which an article is published, which is most often taken to be the impact factor of that journal, also is considered an important factor. Hence scientists strive to get their papers published in journals with high impact factors; and may act unethically to do so (Nature Editors 2014). These incentives, particularly those concerning publications ('the more the merrier'), probably contribute to many of the lapses in integrity in ecotoxicology. This problem became particularly severe in China with its high scientific output and system of tying substantial cash bonuses or other overt rewards to researchers to the impact factors of journals in which their articles were published. This contributed to ingenious and widespread misconduct (Hvistendahl 2013). In response, China recently initiated sweeping reforms with strong disincentives for academic misconduct – institutions will no longer investigate themselves, funding would be withheld from institutions at which misconduct occurred, researchers will be deterred from publishing findings in journals that are deemed to be of poor academic quality and set up merely for profit, and scientists found to have conducted willful misconduct would face severe penalties (Cyranoski 2018; Nature Editors 2018a). How the education for and enforcement of these initiatives develops is likely to influence research communities and funders elsewhere. Likewise in ecotoxicology, moving to more ethical practices in which integrity is central to any endeavor will not be easy to accomplish, nor will it be achieved quickly.

Education of ecotoxicologists, both young and old, is a key way forward towards better culture of integrity in our discipline. That education can be delivered in a variety of ways, the two most obvious and practical being (1) the publication of articles in journals in which integrity and ethics are discussed, and (2) courses run by scientific societies such as SETAC. This article is an example of a very direct attempt at highlighting integrity issues in our field, with the hope that by making ecotoxicologists aware of these unethical practices they will change their behavior and act more ethically. Other, less direct, attempts have involved the publication of papers covering suggestions for how to improve the quality of ecotoxicology research, from the planning stage (Harris et al. 2014) to the publication stage (e.g., Hanson et al. 2017; Moermond et al. 2016). However, it seems unlikely that published papers alone will have a significant influence on the quality of ecotoxicology research because few scientists will be aware of them, and even fewer will read them carefully and subsequently act on the advice in them.

Although there has been some public discussion about what training and skills the ecotoxicologists of the future will require (Harris et al. 2017), this crucial aspect of producing better ecotoxicologists, capable of doing better, and hence more useful, research has rarely been addressed. Yet there are undoubtedly things that could be done to better educate the

ecotoxicologists of the future. A radical proposal would be to require aspiring ecotoxicologists to pass examinations before they are allowed to practice ecotoxicology, either as researchers or regulators. Many professions do insist that its practitioners pass examinations before they are allowed to practice: doctors, dentists, accountants and lawyers are obvious examples. This strategy ensures that practitioners are adequately trained. As a first step towards the goal of ensuring all ecotoxicologists are appropriately trained, specific courses could be introduced, and attendance become mandatory. Courses on topics such as experimental design, statistical analysis, data presentation and how to write a scientific paper could be designed easily. In fact, many research organizations and some industries already run 'in house' courses on these topics, which may include short, memorable messages such as David Glass's series of animations on common pitfalls in experimental design (Glass 2018). Munafò et al.'s (2017) "manifesto for reproducible science" is a framework that could readily be adapted to a course on integrity in ecotoxicology research. In fact, as the issue of integrity (or, more accurately, the lack of it) has gained in prominence in the last few years, some organizations have responded by running training courses for their young scientists on integrity and ethical behavior in research. SETAC could offer such training courses and does so to a limited extent already. Another possibility would be for consultancy companies to develop and run these training courses for clients, who could be universities, research organizations or industrial companies. Consultancy companies that specialized exclusively in providing training could be established; this has happened already in many other professions.

In summary, identifying that there are problems with the way ecotoxicologists are trained currently about integrity issues in their discipline is only the first step. Better education of ecotoxicologists (both those starting their careers and those well-established already) is needed. Such education will need to be provided in a range of formats, to maximize its chances of succeeding. The environment cannot be protected by poor quality ecotoxicology.

Promoting scientific integrity in environmental toxicology

Scientific integrity is harnessed by high quality environmental research characterized by rigor, relevance, reproducibility, and objectivity. Our review suggested several conclusions, tangible actions and less tangible directions that professional societies such as SETAC could do to encourage scientists, their supporting institutions, and science journals to maintain and improve a culture of science integrity. Scientific integrity is reinforced through full transparency exemplified by full disclosures of potential conflicting and competing interests that could contribute to bias, and by making all data and observations readily accessible. Specifically:

1. Scientific integrity in ecotoxicology and the environmental sciences cannot be ensured by impeccable policies or checklists. It is an attitude to be embraced, maintained, and

enforced through the support, guidance and approval of one's peers through a community of practices.

- 2. Reliability, rigor, relevance and reproducibility of science are more important than novel advances, if those advances neglect these "four Rs."
- 3. Increased attention to a culture of quality management training and transparency could improve the confidence in published findings.
- 4. Studies that are not supported by primary data released through data repositories or detailed supporting information are not fully credible.
- 5. Journal publishers and editors could strongly encourage the complete presentation of supporting data, with prominent labeling on the journal and article front matter indicating whether data are available. They should caution authors at the outset that the inability to produce data upon request could be cause for retraction.
- 6. One practical step investigators can take toward improving reproducibility of experiments would be to produce detailed video illustration of their methods.
- 7. As a community, be aware of and disclose potential conflicting or competing interests that could contribute to, or be perceived as, bias and not tolerate extreme conflicts or bias.
- 8. Discourage judging science by its funder; rather, open-minded skepticism is applicable when the funder has a stake in the outcome of a study.
- 9. Scientists, like all people, have moral and ethical assumptions, based upon their values. These should not be intermixed with their interpretations and reporting of science. If scientists' values lead them to cross the lines from analysis to advocacy, they need to be particularly careful about distinguishing between science, values, assumptions, and opinion.
- 10. Professional societies such as SETAC have an important role in fostering respectful evidence-based dialog, in meetings and correspondence on published works.
- 11. Professional societies such as SETAC could support a standing training seminar on principles of scientific integrity, the transparent conduct of science and best practices for peer review in conjunction with its annual meetings.
- 12. Professional societies such as SETAC have a valuable role in facilitating balanced, expert reviews of controversial science topics, such as has been done with their Pellston Workshops series and resulting publications.

Supplemental Information

SI 1. Peer review process

Acknowledgements

We thank the four anonymous peer reviewers and the two institutional reviewers for their constructive criticisms and insights. This commentary and perspectives followed discussions by the Society of Environmental Toxicology and Chemistry (SETAC) ad hoc subcommittee on Scientific Integrity (Fairbrother 2016). Author roles: CAM wrote most of the paper, with contributions by JPS and AF. All other authors participated in discussions, reviewed and edited the drafts, supported the recommendations, and approved of its publication. Specific required disclaimers: The views expressed in this article are those of the authors and do not necessarily represent the views or policies of SETAC, the U.S. Environmental Protection Agency, or the U.S. Fish and Wildlife Service. Affiliations are given for identification, but do not necessarily imply endorsement. Author affiliations reflect a variety of academic, business, and governmental employment, and all authors have competing interests from our previous experiences and work environments, but hopefully these were somewhat balanced out. No specific funding was provided for the writing of this manuscript and no financial conflicts of interest were declared by any author.

Data Accessibility

No datasets are associated with this policy analysis.

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List of Figures

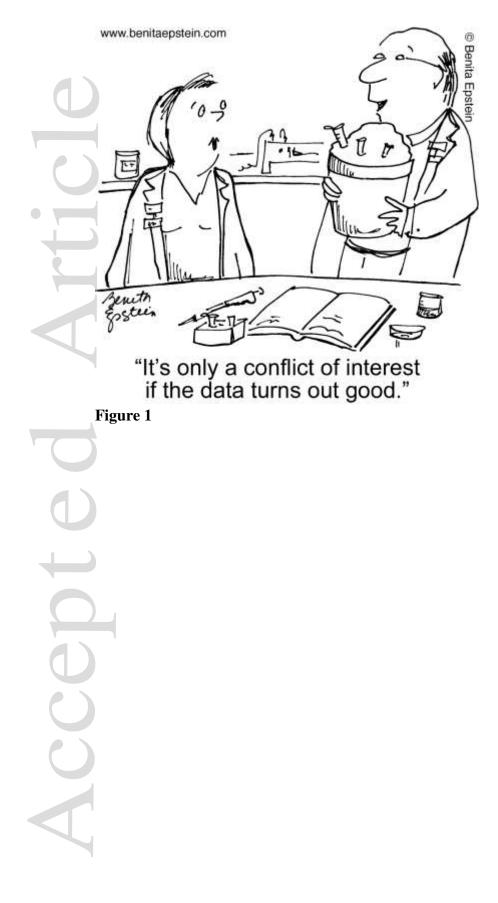
Figure 1. Conflicts of interest in science arise when secondary interests such as financial gain or maintaining professional relationships compromise the primary interest of upholding scientific norms such as the objective design, conduct, and interpretation of studies and the open sharing of scientific discoveries to advance our collective learning (© Benita Epstein, used with permission.)

Figure 2. Confirmation bias is the tendency to seek and interpret evidence in a way that confirms preexisting beliefs and gives less consideration to alternative hypotheses (© Benita Epstein, used with permission).

Figure 3. Large environmental chemistry and toxicology laboratories that use standard methods to produce results that may be submitted to regulatory agencies usually have a well-established quality management structure. Quality management in academic research laboratories focused on novel methods may be more ad hoc, especially if the research work force is dominated by transient scientists, such as students or those on short-term postgraduate appointments (Credit: S. Harris, sciencecartoonsplus.com).

Figure 4. The brief methods descriptions in journal articles are seldom sufficient to be reproducible by others. Step-by-step video documentation of experimental protocols can be published as video articles, uploaded to online repositories, or published as supplemental information. Video protocols are underutilized in environmental toxicology (Credit: S. Harris, Sciencecartoonsplus.com).

Figure published info









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"WE JUST DON'T GET INVOLVED WITH THINGS LIKE DOUBLE-BLIND TESTS AND PEER REVIEW. WE'RE JUST A LITTLE MOM-AND-POP LABORATORY."

Figure 3



"OF COURSE YOU CAN'T REPLICATE MY EXPERIMENT. THERE'S A SECRET INCANTATION THAT YOU HAVE TO CHANT, AND I'M NOT TELLING IT TO ANYONE."

Figure 4