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1 **Addressing Urgent Questions for PFAS in the 21st Century**

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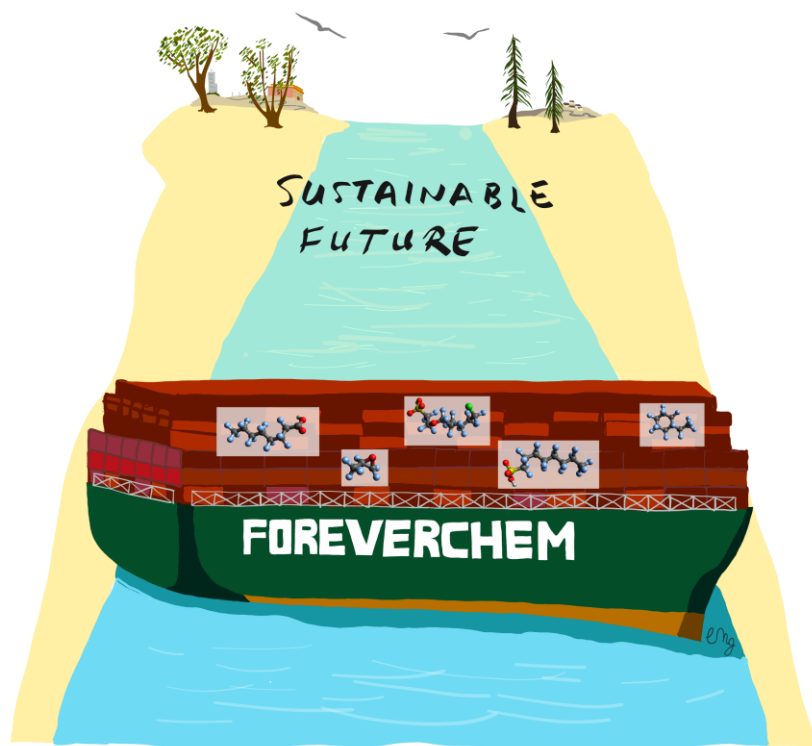
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27 **Table of Contents Art:**



28

29

30 **Abstract**

31 Despite decades of research on per- and polyfluoroalkyl substances (PFAS), fundamental obstacles
32 remain to addressing worldwide contamination by these chemicals and their associated impacts on
33 environmental quality and health. Here we propose six urgent questions relevant to science, technology,
34 and policy that must be tackled in order to address the “PFAS problem”: (1) What are the global
35 production volumes of PFAS, and where are PFAS used? (2) Where are the unknown PFAS hotspots in
36 the environment? (3) How can we make the measurement of PFAS globally accessible? (4) How can we
37 safely manage PFAS-containing waste? (5) How do we understand and describe the health effects of
38 PFAS exposure? And (6) Who pays the costs of PFAS contamination? The importance of each question
39 and barriers to progress are briefly described, and several potential paths forward are proposed. Given the
40 diversity of PFAS and their uses, the extreme persistence of most PFAS, the striking ongoing lack of
41 fundamental information, and the inequity of the health and environmental impacts from PFAS

42 contamination, there is a need for scientific and regulatory communities to work together, with
43 cooperation from PFAS-related industries, to fill in critical data gaps and protect human health and the
44 environment.

45

46 **Synopsis:** This article discusses key gaps in data, understanding, and technology to address the problem
47 of global PFAS contamination, identifies persistent barriers, and suggests useful paths forward.

48

49 **Introduction**

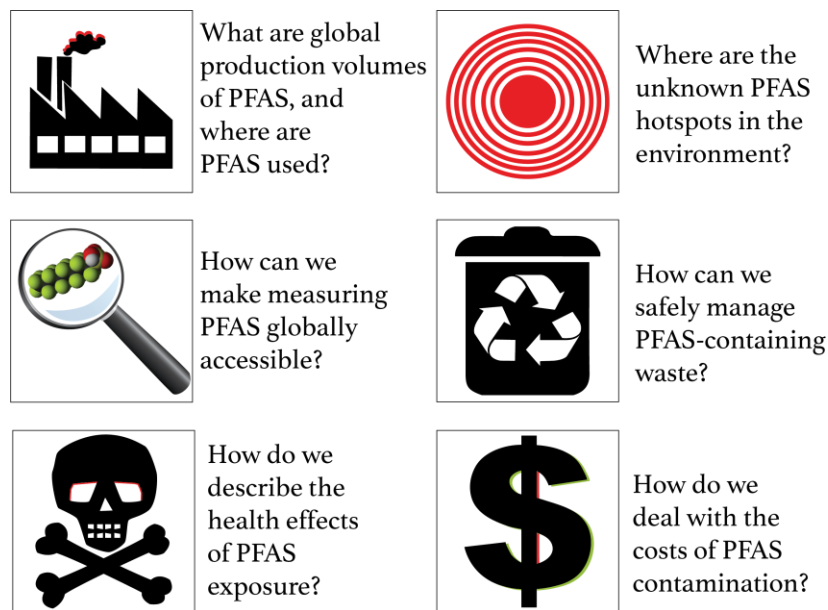
50 Per- and polyfluoroalkyl substances (PFAS) are a class of thousands of chemicals¹⁻³ with perfluorinated
51 carbon moieties that impart physical stability, chemical resistance, and, for most PFAS, extreme
52 environmental persistence⁴. For decades, PFAS have been incorporated into a vast array of products and
53 applications,⁵ and as a result, are pervasive environmental contaminants^{6,7}. The beginning of the 21st
54 Century saw increasing detection of long-chain perfluoroalkyl acids (PFAAs) in the environment and
55 organisms on a global scale. Recognition that some of these chemicals are globally transported,
56 bioaccumulate, and exert multiple adverse effects in biological systems led to regulation and phase-out of
57 several PFAS⁸⁻¹¹. In response, an array of other PFAS have been used as substitutes and are increasingly
58 detected in the environment, in wildlife, and in humans¹²⁻¹⁶.

59

60 Despite two decades of research on fate and transport, biological effects, and environmental emissions,
61 critical gaps remain in our knowledge, preventing researchers and society from finding effective solutions
62 to the “PFAS problem”. This is due to the diversity of chemicals in the PFAS class, to ongoing analytical
63 challenges in detecting, characterizing, and quantifying different PFAS, and to a continued lack of
64 transparency on the part of industry concerning which PFAS are produced, where they are used, and in
65 what quantities. As society grapples with how PFAS may best be regulated and how to prioritize efforts
66 to minimize environmental and human exposure, major challenges remain. Here, we identify a set of six
67 urgent questions that must be addressed for the effective global management and eventual phase-out of

68 PFAS (Figure 1), building on the Zurich Statement on Future Actions on PFAS¹⁷. We also highlight
69 major barriers that impede progress in answering these questions, and provide potential paths forward
70 from the perspectives of science, technology, and policy.

71



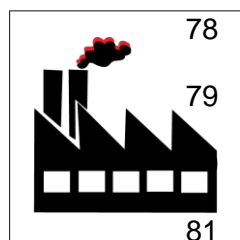
72

73 **Figure 1:** Six urgent questions relevant to science, technology, and policy that must be tackled in order to
74 address the “PFAS problem”.

75

76 **1. What are the global production volumes of PFAS, and where are PFAS used?**

77 **Importance:** This deceptively simple question highlights a fundamental gap in society’s knowledge about



nearly all PFAS. Despite painstaking emission estimates for the best-characterized
sub-classes of PFAS^{18,19}, there is a lack of information on historical and ongoing
production volumes of most PFAS, including even their identities^{1,20,21}. This
information is needed to build reliable emissions inventories, investigate

82 environmental fate and transport, and assess associated exposures and health risks. While this is a general

83 problem for most chemicals in commerce²², the multitude of uses for PFAS and the transformation of

84 various precursors into the same PFAS end-products make tracking the sources of PFAS exposure to

85 production and use particularly difficult. Without these data, society will fail to protect its members from
86 unknown exposures until or even after harmful and irreversible effects are discovered.

87

88 **Barriers:** Regulatory bodies in many countries have developed registries of chemicals produced or used
89 in their jurisdictions^{18,19,22–24}, but much of the collected information is confidential. In addition, many
90 newer uses of PFAS remain poorly documented in the technical literature. The Kirk-Othmer
91 Encyclopedia of Chemical Technology (2004)²⁵ and Kissa (2001)²⁶ are considered authoritative reference
92 sources for industrial applications of PFAS. However, most of the PFAS-relevant content in both were
93 written before the EPA’s Stewardship Program (2006)⁸, the addition of perfluorooctanesulfonic acid
94 (PFOS; 2009), perfluorooctanoic acid (PFOA; 2019) and their precursors to the Stockholm Convention,¹¹
95 and a number of PFAS restrictions under the European Union REACH legislation²⁷. The EPA’s Toxics
96 Release Inventory (TRI), designed to inform the public of releases of toxic chemicals in their
97 communities, can shed light on some larger sources of PFAS releases. However, it often falls short of the
98 level of detail needed to characterize environmental contamination because it requires only self-reporting
99 and contains extensive exemptions for many industry sectors (e.g., oil and gas), small businesses, facility
100 cleaning and maintenance applications, and trade secret claims, among others. A recently proposed new
101 rule under the Toxic Substances Control Act (TSCA) could overcome some of these key limitations for
102 PFAS, as discussed in the “Paths Forward” section that follows.

103

104 As a further complication, emissions and exposures vary depending on the properties, production, use
105 patterns and end-of-life treatments of the product and the PFAS applied. A recent broad overview of
106 PFAS uses⁵ in different consumer and industrial applications revealed a large number of little known uses
107 such as in ammunition, climbing ropes, guitar strings, artificial turf, and soil remediation. For other areas
108 (e.g., cosmetics, paints), PFAS use is known, but it is often less clear which specific PFAS have been
109 employed and at what quantities. Such lack of knowledge about PFAS in industrial processes and
110 products also impacts retailers and consumers. Public pressure to phase out hazardous chemicals has led

111 major retailers to remove certain PFAS from food packaging, clothing, and household furnishings^{28,29}.
112 However, retailers and product manufacturers often run into issues wherein PFAS are used somewhere
113 along the supply chain but the exact use, PFAS type, and concentration are unknown. Proprietary
114 information is used by industry as a justification for withholding the identity and concentration of
115 chemicals in commercial products, with Confidential Business Information (CBI) claims used to protect
116 details of formulas and manufacturing processes that confer an advantage over a company's competitors.
117 This means that often little is publicly known about the identity and quantity of specific chemical
118 structures present within a substance, formulation, or product.

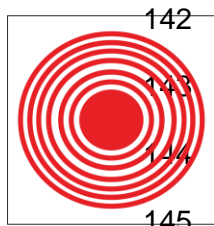
119
120 ***Potential Paths Forward:*** Chemical identities, production and consumption volumes, use locations and
121 emissions, including of byproducts and impurities, need to be reported by industry, and such information
122 needs to be made publicly accessible. Retailers and product manufacturers need to know and publish
123 where PFAS are present in their supply chains to foster greater transparency and confidence in the
124 composition and safety of end products. This will require public pressure, rules, and regulations. In June
125 2021, the US EPA published a proposed update to the reporting requirements for PFAS under TSCA³⁰
126 that could facilitate this type of reporting. The new rule potentially applies to a larger number of PFAS
127 and no longer exempts small-scale businesses that manufacture PFAS from reporting requirements, an
128 acknowledgment of the particular concern raised by these chemicals. However, this rule is still limited to
129 producers, and as such will not resolve the supply chain issues of identifying PFAS in and emissions from
130 downstream products. In addition, confidentiality of production and import volumes and chemical
131 identity are still supported under the proposed rule, thus continuing to limit public access to these critical
132 data under CBI claims. Another potentially useful mechanism is greater use of product registries, such as
133 are maintained by the Scandinavian countries³¹⁻³³. These require manufacturers and importers to declare
134 chemical substances and products (excluding food, cosmetics, and medicinal products) in excess of 100
135 kg per year per company. Finally, a researcher-led approach to identifying PFAS occurrence in products
136 and environmental emissions could entail greater use of coordination networks like NORMAN³⁴. Such a

137 network can serve as a central touchpoint to harmonize analytical methods and share information on
138 occurrence and effects of PFAS, but is limited to detecting pollution after it has occurred.

139

140 2. Where are the unknown PFAS hotspots in the environment?

141 **Importance:** The ability to identify geographic areas, environmental media, and populations with high



142 PFAS concentrations is crucial to manage exposures and for the development of
143 models to predict PFAS transfer across environmental media, geographic borders, and
144 food webs. The scientific community is well aware of certain contaminated sites such
145 as airports and military facilities,³⁵⁻³⁷ pulp and paper mills³⁸ and fluoropolymer

146 manufacturing facilities³⁹⁻⁴³, but others have only recently come to light⁴⁴. Certain activities can lead to
147 decade-long local releases that are poorly documented, because the respective PFAS amount is not
148 substantial on a regional or global scale, and therefore difficult to identify without local knowledge.

149

150 **Barriers:** Region- or country-specific uses exist that may constitute important but primarily local
151 contamination hot-spots. For example, high volumes of PFOS have been emitted in South America
152 through the use of Sulfloramid, an insecticide containing the PFOS-precursor N-ethyl perfluorooctane
153 sulfonamide used to control leaf-cutting ants^{45,46} Moreover, small-scale manufacturers in both developed
154 and developing countries have very different control practices in place, leading to PFAS emissions that
155 are poorly understood in light of the current knowledge of a few large industries, mostly in the developed
156 world. In developing countries, a general lack of access to the equipment, supplies, and infrastructure
157 needed to perform PFAS analyses can hinder identification of hotspots, a particularly critical barrier
158 discussed in detail under Question 3.

159

160 **Potential Paths Forward:** A systematic inventory of all PFAS industries is needed to identify current and
161 former sites of emissions on a global scale. This requires international collaboration to integrate
162 knowledge about locally important industries and practices. These inventories of industrial activities can

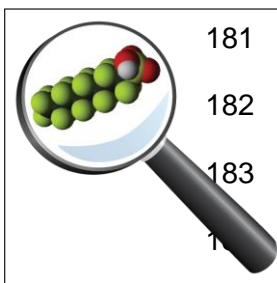
163 then be connected to known PFAS uses, enabling a systematic population of maps of potential PFAS
164 contamination on a global scale, and bringing into focus areas that have been historically neglected in
165 monitoring campaigns and/or research. This type of approach, for example using geographic information
166 systems (GIS) to share and distribute data, is a means to organize knowledge and plan sampling
167 campaigns on a global scale.

168
169 At the same time, funding from multiple sources (industries, governments, foundations) for monitoring
170 campaigns that screen diverse media (e.g. air, water, soils, sediments) for PFAS can identify geographical
171 hot spots not connected to a known or suspected PFAS-associated activity. Data on emissions and
172 environmental occurrence could be integrated and evaluated through the use of environmental fate and
173 transport models^{47,48}. Mismatches between model predictions and measurements can provide clues to
174 missing emissions sources or hot spots. The data generated through these concerted efforts will be key to
175 raising awareness at the governmental level on the urgency and scale of PFAS pollution, with the intent to
176 motivate sufficient funding for monitoring and remediation activities on a large scale, as well as stopping
177 ongoing emissions of identified local sources.

178

179 3. How can we make measuring PFAS globally accessible?

180 **Importance:** Overcoming uncertainties in global and local PFAS distribution and exposure, and closing



181 critical geographical and biological data gaps as discussed above also requires,
182 fundamentally, the ability to actually detect and measure a wide range of PFAS
183 compounds in myriad locations and in diverse environmental and biological
184 media. Analytical methods are needed for environmental media, drinking water,

185 sewage sludge, foods, blood, fat, and various kinds of products and technical mixtures for monitoring and
186 enforcement of current and upcoming regulations. Giving more researchers, communities, health-care
187 providers, utilities, and businesses the ability to accurately detect PFAS will facilitate efforts to minimize
188 exposure, protect vulnerable populations of humans and wildlife, and evaluate the effectiveness of

189 interventions. Making resources available to scientists in developing countries and developing rapid and
190 cost-effective analytical approaches that are reliable and accessible will greatly improve the
191 understanding of PFAS sources, fate and transport in areas where relatively little is currently known, such
192 as Africa, Central America and parts of Asia.

193

194 **Barriers:** Until now the ability to measure and monitor PFAS has largely been restricted to well-
195 resourced groups and countries with access to equipment, standards, infrastructure, and expertise. Well-
196 established methods that can achieve high sensitivity with robust quality control require sophisticated
197 analytical equipment (e.g. liquid chromatography tandem mass spectrometry, LC-MS/MS) that is
198 expensive to acquire and requires specialized expertise to operate and maintain. In the past, the analysis of
199 PFAS has been particularly challenging due to the presence of PFAS in certain laboratory and sampling
200 materials and equipment, requiring control and monitoring of contamination, though measures have been
201 developed to overcome this challenge^{49,50}.

202

203 Reliable and well-documented protocols are still limited to a narrow range of PFAS, and high-quality
204 analytical reference standards that enable targeted analysis with reliable quantification are expensive, and
205 still unavailable for many PFAS. Commercial standard providers^{51,52} cover only about 80 different PFAS,
206 plus variations (i.e., branched isomers or mass-labeled compounds). Without the availability of analytical
207 standards, non-targeted analysis methods with expensive equipment and expertise are needed to identify
208 unknown PFAS^{53,54}. Recent actions by a PFAS producer may set a worrisome precedent. According to a
209 letter sent by Wellington Laboratories to its customers in January 2021, the PFAS manufacturer Solvay
210 has threatened to sue Wellington for patent infringement for their sale of a standard for a novel PFOA-
211 replacement in Solvay's fluoropolymer production (CAS 1190931-41-9)⁵⁵. This raises the potential for
212 industry to monopolize access, maintain secrecy, and delay progress in establishing occurrence and
213 toxicity data for these substances.

214

215 **Potential Paths Forward:** While the low (part per trillion) limit levels instituted for PFAS in drinking
216 water in many jurisdictions^{9,56} require high sensitivity methods and rely on the availability of standards,
217 for purposes such as screening of sites or products, simpler lower-cost methods may suffice. There are
218 several Total Fluorine (TF) methods to detect the presence of fluorine or fluorinated compounds (e.g.
219 CIC⁵⁷, PIGE⁵⁸, and XPS⁸³), which can be combined with sample preparation methods such as extractable
220 organic fluorine (EOF⁵⁹) to provide rapid screening of both abiotic and biotic matrices. Much research is
221 ongoing to develop additional methods, such as versatile and low-cost PFAS sensors⁶⁰⁻⁶². Whatever their
222 technical approach, methods should be validated across laboratories and ideally standardized. Positive
223 steps in this direction were recently illustrated for EOF measurements in water compared to total targeted
224 PFAS in a Swedish interlaboratory comparison study⁶³.

225
226 Capacity building efforts can support a pipeline for training and technology transfer from better resourced
227 countries and institutions. Some programs already exist for instrument donation, such as the Seeding Labs
228 program on Instrumental Access that donates equipment to promote research and education in developing
229 countries⁶⁴. Such programs are important, but represent only a small part of the solution to this enormous
230 challenge. In addition to equipment, access to supplies (e.g. standards, solvents) and reliable infrastructure
231 (electricity, water, gases) is crucial and often unavailable. To make these efforts accessible and
232 sustainable, traineeships could be established for scientists in under-resourced regions to learn PFAS
233 analysis at host laboratories. This would provide the opportunity for scientists in regions without
234 adequate infrastructure to collect local samples to be analyzed at the host institution, while retaining
235 ownership of the data and authorship in resulting publications.

236

237 **4. How can we safely manage PFAS-containing wastes?**

238 **Importance:** As PFAS are phased out of specific products and uses, safe disposal of existing stockpiles



239 becomes an urgent need. There are many diffuse sources of PFAS, such as textiles,
240 food contact materials, personal care products, and household furnishings, that
241 eventually enter landfills and wastewater, and are later re-emitted to the
242 environment through the air, landfill leachate, or into soil from biosolids

243 application⁶⁵⁻⁶⁷. Within recycling streams, separation and safe disposal of PFAS contained within
244 complex matrices become extremely challenging, given knowledge gaps on which types of PFAS are
245 present, and at which levels, in various types of waste. Knowledge on how to deal with PFAS-containing
246 waste is also critical for legislation related to regulations such as EU REACH and the Stockholm and
247 Basel Conventions and ongoing PFAS restrictions.

248

249 **Barriers:** Multiple technologies are being developed to remove PFAS from contaminated soil and water,
250 some of which have proven effective, but high long-term cost and energy use remain major challenges.
251 For example, sorptive or membrane-based processes result in contaminated wastewater streams (spent
252 sorbent, membrane rejectate) that must be disposed of. Most desirable are in-situ clean-up methods (not
253 “pump and treat”) but, so far, such a remediation solution has not been found. Large-scale water treatment
254 facilities can be equipped with advanced treatment technologies (e.g. reverse osmosis) to remove
255 persistent and mobile (water-soluble) chemicals like PFAS, but these are prohibitively costly to install
256 and maintain for small systems⁶⁸⁻⁷² and also generate PFAS-containing waste.

257

258 High-temperature incineration has been proposed for some concentrated stocks (e.g. aqueous film-
259 forming foams), but given the high stability of the carbon-fluorine bond, there are concerns whether
260 incineration is consistently operated under conditions that ensure the full mineralization of PFAS. In
261 Europe, flue gases from municipal waste incinerators are meant to run at a temperature of 850 °C for at
262 least two seconds⁷³, but studies show that complete combustion of PFAS such as PFOA and PFOS
263 requires temperatures of at least 1000 °C⁷⁴. Limited work is underway to monitor incineration plants for

264 emissions of PFAS, but few data from full-scale studies are yet available⁷⁵. While intensive research is
265 ongoing to identify and optimize routes of PFAS biodegradation⁷⁶⁻⁷⁸ as a potentially less energy- and
266 cost-intensive solution, none are currently effective at complete mineralization under reasonable time-
267 scales.

268

269 **Potential Paths Forward:** Given the difficulties and costs associated with the disposal of PFAS, an
270 upstream solution (i.e. avoiding PFAS except for cases of essential uses) is the most effective means of
271 dealing with future PFAS-waste. The production of PFAS for essential uses should also be carefully
272 controlled to result in close-to-zero emissions, because the few options available for safe disposal will
273 always be costly based on currently available and foreseeable technologies. Recovery of PFAS from such
274 uses is another important measure to ensure the need for energy-intensive destruction is avoided. Product
275 labeling can be effective in reducing use and emissions of hazardous chemicals including PFAS, but trace
276 PFAS contamination within recycling streams may prevent recycled materials from being incorporated
277 into goods labeled PFAS-free. Given existing background levels, it may be necessary for PFAS-free
278 labeling to include an allowance for trace, non-functional levels of PFAS for industry partners trying to
279 move away from fluorinated chemicals.

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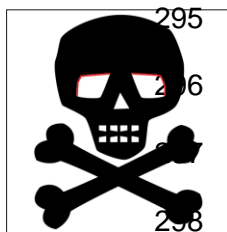
281 Even when an “ideal” future can be achieved where only essential uses of PFAS occur and PFAS from
282 these uses are recovered and not released, there are still the problems of legacy PFAS contamination and
283 ongoing PFAS emissions. To address existing and ongoing waste issues, funding and research should be
284 targeted towards technologies that can destroy PFAS with reasonable cost and environmental
285 performance. Hybrid technologies that combine sorption and mineralization (“concentrate and destroy”)
286 approaches may be particularly helpful in dealing with initially complex and dilute waste streams.
287 Whatever the approach, the re-emission and shifting of contamination across environmental media (e.g.
288 from soil to air) must be prevented. This also argues against testing of destruction technologies at scale
289 until proven strategies are in place to prevent re-emission. Until these technologies can be better

290 developed, confined disposal facilities that store PFAS wastes while preventing emissions via air and
291 leachate may be a best imperfect choice.

292
293

5. How can we understand and describe the health effects of PFAS exposure?

294 **Importance:** Toxicological assessment of each of the thousands of PFAS is not required to decide that



295 further environmental contamination by PFAS and subsequent exposure should be
296 avoided. However, pressing questions remain about how to deal with historical and
ongoing PFAS pollution and associated health effects. To address the potential
effects of existing exposures, and to prevent the extensive use of similarly

299 bioavailable and toxic substances in the future, it is important to understand how to link measured
300 exposures (e.g., levels of specific PFAS in blood) to current or anticipated health effects. It is also critical
301 to link those health effects to specific physical-chemical properties and modes or mechanisms of
302 toxicological action of PFAS, for example through adverse outcome pathways, AOPs. Concerns about
303 their bioaccumulation and toxicity led to the global phase-out of a number of PFAAs. Yet advances in
304 non-targeted analysis have facilitated discovery of many other structurally similar compounds in the
305 environment, wildlife, and humans^{16,42,80,81}. Some of the newly detected compounds are attracting
306 increasing attention as they replace phased-out PFAAs in processes and products^{41,82,83}, although
307 they have in fact been released for decades in certain industries⁸⁴⁻⁸⁶ but were under the radar of the
308 scientific and regulatory communities. The tissue distributions and bioaccumulation potentials are still not
309 well understood^{79,80}, but laboratory studies suggest that several replacement PFAS bioaccumulate and/or
310 exert toxic effects similar to the compounds they have replaced, as well as other distinct toxic effects<sup>44,81-
311 84</sup>.

312

313 **Barriers:** One of the most difficult questions scientists working on PFAS face is that of causality: is a
314 health condition suffered by a community member the result of their exposure to PFAS, or does a blood
315 test indicating the presence of PFAS mean that they will become sick in the future? Communities with

316 contaminated water supplies face challenges in court to having their health and remediation costs covered
317 by the parties thought responsible for the contamination. To make the link between exposure and effect,
318 clear lines of evidence are needed to both document the exposure and explain how it leads to an observed
319 adverse health impact⁸⁷. A striking feature of PFAS toxicity is the diversity of biological pathways that
320 are affected⁸⁸, especially given that most of the toxicological data currently available for PFAS are for a
321 few single PFAAs. Understudied groups of PFAS (e.g. neutral, cationic, zwitterionic, cyclic) may have
322 substantially different biological behavior that could be missed by established sampling approaches. For
323 example, if their tissue distributions vary from those of anionic PFAAs, focusing on only serum or liver
324 concentrations could miss critical accumulation sites for these PFAS (e.g. in lipids⁸⁹). The structural
325 diversity of PFAS and the fact that exposures are nearly always to mixtures rather than single substances
326 complicates the search for mechanisms and structure-activity relationships.

327

328 ***Potential Paths Forward:*** The use of class-based methods to evaluate PFAS can work as a
329 precautionary approach in the face of continuing uncertainty, particularly with respect to curtailing new or
330 continuing uses of PFAS⁹⁰. For existing exposures, additional, appropriately funded epidemiological
331 studies that target large populations with a diversity of primary exposure routes can help to develop better
332 links between exposure and effect, especially for less-studied PFAS and exposure routes. Analyses in
333 these studies should include not only blood but also other matrices (urine, breast milk, hair, lipid tissues)
334 to capture a wider diversity of PFAS physicochemical properties, half-lives of elimination, and potential
335 internal storage sites. When occurrence data in populations are combined with PFAS identities and
336 concentrations in products and environmental matrices, as discussed under questions 1 and 2, scientists
337 can begin to develop “signatures” for exposures to PFAS from specific sources. Such information would
338 be highly useful in the design of effective interventions to minimize exposures. Strategic and periodically
339 implemented human biomonitoring studies combined with environmental exposure assessments can also
340 evaluate effectiveness of regulatory initiatives^{91,92}.

341

342 Better integration of mechanistic and observational studies can reveal how PFAS induce adverse health
343 outcomes in humans and wildlife. Computational and in-vitro approaches (e.g. toxicokinetic models^{93,94},
344 food-web bioaccumulation models⁹⁵⁻⁹⁷, protein and phospholipid interaction models and in-vitro
345 studies⁹⁸⁻¹⁰³) can provide insight into expected exposures and effects in diverse species. However, these
346 newer approaches still face substantial barriers to inform policy, as regulatory approaches still often
347 require that risk assessment used to support regulatory standards be based on human epidemiology data or
348 in vivo animal toxicology data. These data are largely lacking for many of the PFAS now widely detected
349 in the environment. Strategies to incorporate in vitro and computational data into regulatory framework
350 would allow for more rapid expansion of risk assessment to emerging PFAS. Such studies could be
351 further strengthened by systematic reviews of existing data to confirm or refute linkages between
352 exposures and outcomes. To avoid regrettable substitution with existing PFAS and non-PFAS
353 alternatives, information revealed about modes or mechanism of toxic could also be used to inform future
354 chemical design. Chemists should incorporate principles of hazard assessment, including structure-
355 activity relationships, early in the molecular design phase to aid in the development of chemicals that are
356 less persistent, bioavailable and toxic.

357

358 **6. Who pays for the impacts of PFAS contamination?**

359 **Importance:** A 2019 study for the Nordic Council of Ministers estimated the costs for Europe of water



360 treatment and soil remediation due to contamination of a sub-set of PFAS at between
361 EUR 10-20 billion over a 20-year period¹⁰⁴. Testing of publicly supplied drinking
362 water sources indicates that as many as 80 million US residents may be receiving
363 water with PFAS levels exceeding limits recommended by regulatory agencies and
364 toxicologists.^{9,105-107} These communities may face costs ranging from purchase of replacement (bottled)
365 water to major capital expenditures and long-term maintenance of water treatment technologies by their
366 water utilities, which are transferred to consumers through their water bills¹⁰⁸⁻¹¹¹. Removal and disposal
367 of contaminated soil or treatment of groundwater (e.g., pump and treat) is particularly expensive¹¹², and is

368 therefore rarely undertaken. Indirect costs can include loss of property value or closure of a business if
369 contamination is found. Examples include an organic farm in Colorado that had to stop growing crops
370 because its water supply had been contaminated by PFAS from fire-fighting foam¹¹³, and a dairy farm in
371 Maine that had to cull its herd because the milk had levels of PFAS 60 to 150 times higher than health
372 advisory levels, due to applications of contaminated paper mill sludge to pastures as fertilizer¹¹⁴.

373
374 Moreover, projected health-related costs due to effects of PFAS exposure are many times higher than the
375 costs of environmental remediation. The Nordic study estimated the costs of human-health impacts from
376 exposure to PFAS to be a minimum of EUR 54-82 billion *each year* in Europe. Direct costs will include
377 medical treatment for PFAS-related health impacts such as cancer, high blood pressure, obesity and low
378 birth weight. Indirect costs range from lost years of life and/or lost quality of life, impacts on family or on
379 mental health because of anxiety about PFAS exposure, and ongoing health monitoring.

380
381 **Barriers:** Costs of environmental and health impacts from PFAS exposure, like most environmental
382 damages, continue to be treated as negative externalities – costs not borne by the polluter carrying out the
383 activity causing the exposure, but by society at large. The major barrier to covering these enormous costs
384 is lack of political agreement concerning who is responsible for this contamination and exposure, and who
385 should pay. While the “Polluter Pays Principle” was first defined and championed by the OECD in 1972,
386 it has rarely been implemented¹¹⁵. When local, regional, or national governments step in to finance clean-
387 up of drinking water and other remediation processes, the costs are ultimately passed on to the taxpayer.

388
389 The costs of health impacts from PFAS exposure are often borne directly by the individuals who have
390 developed the disease and by healthcare systems, because of complexities associated with establishing
391 direct causal links between pollution and the health impact. The relationship between exposure and
392 disease can be particularly difficult to verify when impacts of exposure do not arise until many years later
393 (e.g., cancer). In the US, a few legal actions for compensation have been successful, e.g., a class action

394 suit against Dupont/Chemours on behalf of 70,000 persons exposed to industrial discharges in West
395 Virginia settled for \$670 million and a State of Minnesota lawsuit against 3M for water contamination
396 settled for \$800 million. However, the PFAS released by these companies remain in the environment and
397 will likely remain a source of exposure for generations, not covered by these lawsuits.

398

399 **Potential Paths Forward:** The extreme persistence of nearly all PFAS highlights the absurdity of
400 continuing to treat environmental damage—including damage to public health—as a negative externality
401 that can be ignored or even denied by the emitter. Such long-lived environmental contamination does not
402 simply shift a burden but rather extends it, indefinitely, to future generations and all species. This is not a
403 transaction that can be supported in a sustainable society for the sake of preserving a specific market or
404 manufacturer. Mechanisms already exist that could be activated to shift cost burdens away from
405 communities and taxpayers, such as the aforementioned Polluter Pays Principle. The Superfund program
406 under the Comprehensive Environmental Response, Compensation, and Liability Act of 1980 (CERCLA)
407 in the United States¹¹⁶ can hold polluters retroactively liable, but requires that the chemical to be
408 remediated is first designated as a hazardous substance. The designation of PFAS as hazardous substances
409 in the US is still under debate¹¹⁷, but would mark an important step forward in assigning liability.

410

411 However, liability might justifiably lie with different parties under different circumstances. Should the
412 polluter be defined as the company that released the PFAS-containing material into the environment or
413 the company that manufactured the material in the first place? Was the product that contained PFAS
414 properly used? Was it properly disposed of? Was the user sufficiently informed about the risks of release?
415 How should that liability be treated when companies merge, split, and otherwise change their structure
416 and identities, such as when Dupont spun off Chemours in 2015 and offloaded much of their PFAS-
417 related liability¹¹⁸? A number of cost recovery mechanisms have been suggested under the Strategic
418 Approach to International Chemicals Management¹¹⁹ that could help countries to address these issues, by
419 funding assessment, remediation, and health care costs. These include collecting fees from companies

420 who wish to register chemicals for use, charging environmental protection taxes, and charging for
421 permits.

422

423 **Conclusions: Answering Urgent Questions to Address the PFAS Problem**

424 While these urgent questions highlight critical gaps in current understanding of the PFAS problem,
425 enough is already known to take action. Costs associated with environmental cleanup and ongoing health
426 effects of chemicals are magnified for extremely persistent environmental contaminants^{4,120}, adding
427 urgency to efforts to phase out current non-essential uses of PFAS¹²¹. Beyond these well-founded
428 precautionary actions, the most important step is to improve the transparency about where and in what
429 quantity PFAS are used. This will aid in identifying and phasing out all non-essential uses of PFAS and
430 provide opportunities to identify less hazardous substitutes for PFAS. Production of safer chemicals and
431 products must be seen as a competitive advantage and as a driver for innovation and the opening of new
432 markets.

433

434 Consumers are increasingly demanding that the products they use minimize their own health risks as well
435 as risks to environmental health. These consumer-driven initiatives place pressure on major retailers to
436 remove known problematic chemicals—e.g., bisphenol A (BPA)¹²², polybrominated diphenyl ethers
437 (PBDEs)¹²³, and, now, PFAS—from their products, and have proven enormously effective. However, this
438 is not a perfect system, as illustrated by the case of BPA, where consumer pressure led to its replacement
439 by bisphenol S (BPS), which has turned out to be just as harmful as BPA¹²⁴. Thus, while consumers can
440 demand that known harmful chemicals be removed from their products, it is up to industry under the
441 purview of scientific and regulatory communities to ensure that regrettable substitutions do not occur. A
442 first step would be to move towards household goods, cosmetics, food-packaging materials, and personal
443 care products with a smaller total number of ingredients, simplifying the assessment of a particular
444 formulation.

445

446 While consumers have direct purchasing power, their ability to use this to avoid hazardous substances is
447 impeded by the lack of transparency in product ingredients and increasing cases of ‘greenwashing’. Major
448 retailers and institutions in charge of public procurement, on the other hand, can wield much more
449 concentrated power as well as knowledge about product supply chains. When large multinational
450 corporations demand that their product lines remove certain hazardous chemicals, it helps in the voluntary
451 restriction of those chemicals and also serves as a driver for innovation in the search for less hazardous
452 alternatives. One particularly effective means for public agencies and retailers is through the use of lists
453 of prohibited chemicals, such as the “Substitute it Now” (SIN) list,³¹ which can serve as a scientifically-
454 vetted ‘manual’ of chemicals to avoid. Compilation and curation of such lists, as well as their
455 counterparts—lists of preferred less hazardous chemicals and products such as US EPA’s Safer
456 Choice¹²⁵—can help to prevent the chemical whack-a-mole game of regrettable substitutions.

457
458 The environmental health impacts of a chemical used in a product are often not borne by the same
459 population who benefits from the sale and use of these products. Production of PFAS has shifted to
460 China, India, Brazil, and other countries where there is little awareness of the public health risks from
461 PFAS and almost no environmental or human health monitoring. Extremely high exposures are already
462 occurring, as was recently documented near a production facility in China⁴⁴. A key component of the
463 solutions we propose here is to ensure that PFAS research and monitoring is supported in more countries,
464 with the goal to alleviate the impacts of “off-shoring” the negative repercussions of emissions associated
465 with the production and end-of-life of PFAS and PFAS-containing products. In answering urgent
466 questions for the sustainable management of PFAS, technological and policy interventions cannot be
467 effective without also addressing environmental equity.

468

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479

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