

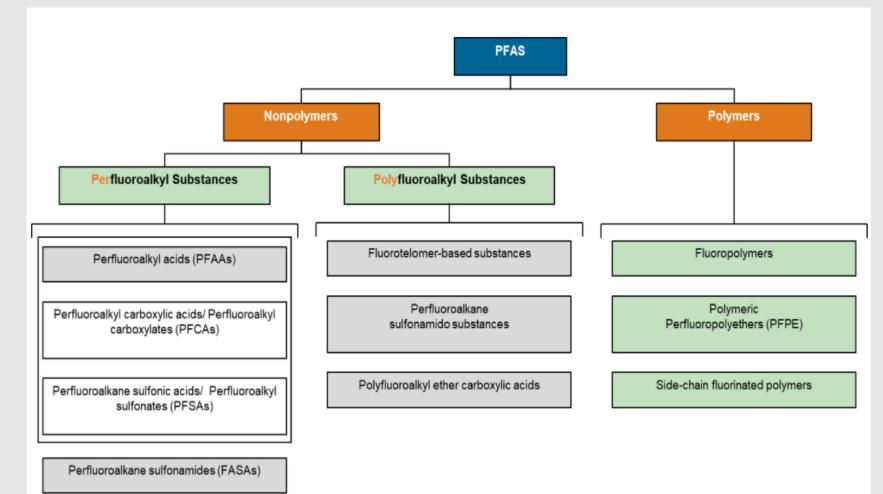
Overview of Per- and Polyfluoroalkyl Substances (PFAS) Activities and Considerations

Presented by

Miriam Calkins, PhD, MS Research Industrial Hygienist CDC/NIOSH/DFSE/FRB

April 21, 2023

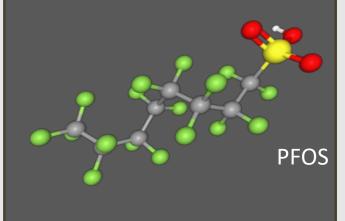
Classification



Synthetic, not naturally occurring

> 12,000 substances

Fluorinated aliphatic (carbon chain) structure



Interstate Technology & Regulatory Council (ITRC). 2020

Why are PFAS a concern?









Integrated extensively into products and processes since 1950s

Desirable properties

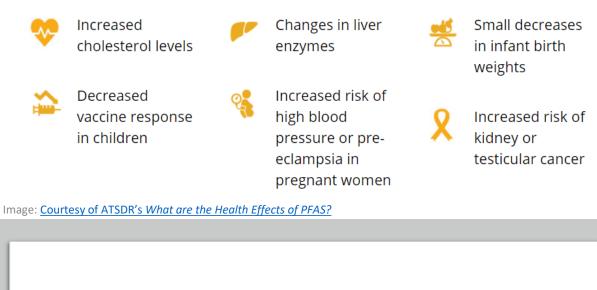
"Forever chemicals"

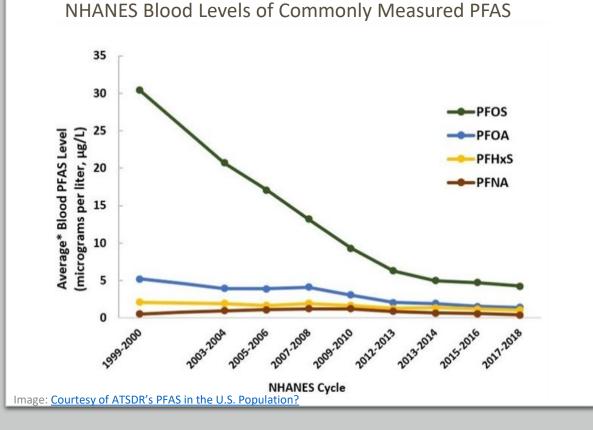
Toxicological and epidemiological evidence of health effects affecting multiple systems

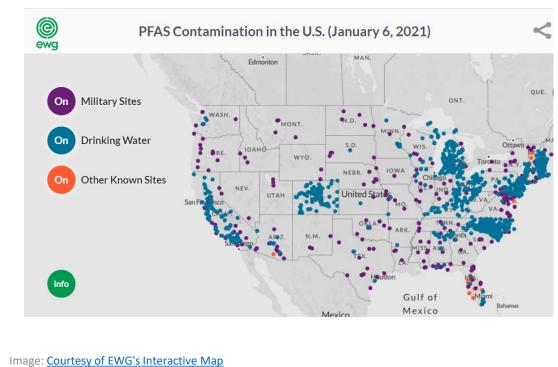


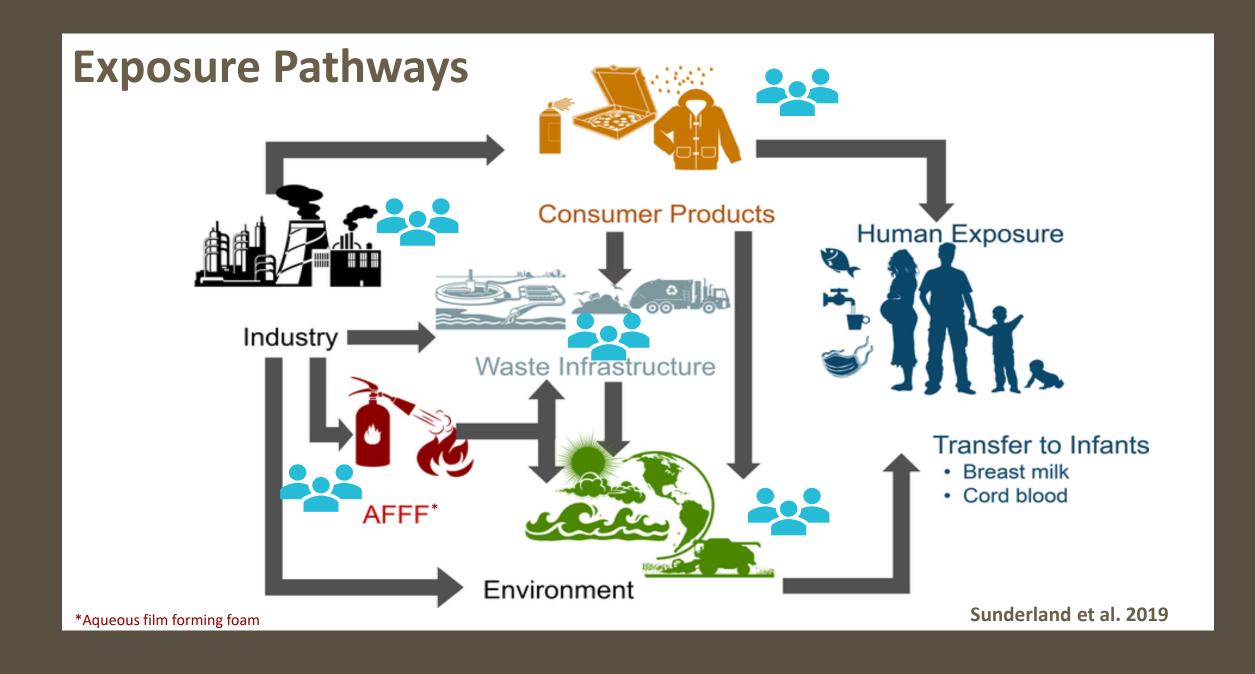


Community Exposure and Health









Big Picture

- High priority topic
- Challenges resulting from
 - PFAS as a class of chemicals
 - Different definitions
 - Availability of methods
- Voluntary phase out of select PFAS
- Highly litigious
- Legislation increasing

Federal, state, and local government





Unions and impacted parties

Standards organizations



Standards Worldwide



World's first ban on products with PFAS adopted in Maine

Law allows exemptions for health and safety when alternatives aren't available by Cheryl Hogue

July 19, 2021

Ski wax chemicals found in Park City's aquifer and groundwater wells

Leaders enact ban on use of fluoro ski wax due to 'forever chemicals'

By Amy Joi O'Donoghue | Mar 13, 2023, 6:55pm EDT

States Take on PFAS 'Forever Chemicals' With Bans, Lawsuits 3M to Exit PFAS Manufa

Read time: 7 mi

STATELINE ARTICLE

September 22, 2022 By: Alex Brown

3M to Exit PFAS Manufacturing by the End of 2025

ST. PAUL, Minn., Dec. 20, 2022 / PRNewswire/ -- 3M (NYSE: MMM) today announced it will exit

Biden-Harris Administration Proposes First-Ever National Standard to Protect Communities from PFAS in Drinking Water

substance (PFAS) manufacturing and work to discontinue the use of PFAS folio by the end of 2025. 3M's decision is based on careful consideration ion of the evolving external landscape, including multiple factors such as trends focused on reducing or eliminating the presence of PFAS in the

March 14, 2023

Contact Information EPA Press Office (press@epa.gov) ECHA publishes PFAS restriction proposal

ECHA/NR/23/04

The details of the proposed restriction of around 10 000 per- and polyfluoroalkyl substances (PFASs) are now available on ECHA's website. ECHA's scientific committees will now start evaluating the proposal in terms of the risks to people and the environment, and the impacts on society.

WHO International Agency for Research on Cancer (IARC)

- 2014 PFOA determined to be group 2b in Monograph 110.
- November 2023 PFOA and PFOS scheduled for Monograph 135.



National Academies of Sciences Engineering and Medicine (NASEM)

Guidance on PFAS Exposure, Testing, and Clinical Follow-Up

Released July 28, 2022

* Simple additive sum of MeFOSAA, PFHxS, PFOA (linear and branched isomers), PFDA, PFUnDA, PFOS (linear and branched isomers), and PFNA in serum or plasma

\geq 20 (ng/mL) PFAS*

Encourage PFAS exposure reduction if a source of exposure is identified, especially for pregnant persons.

In addition to the usual standard of care, clinicians should:

- Prioritize screening for dyslipidemia with a lipid panel (for patients over age 2) following American Academy of Pediatrics (AAP) recommendations for high-risk children and American Heart Association (AHA) guidance for high-risk adults.
- At all well visits:
 - Conduct thyroid function testing (for patients over age 18) with serum thyroid stimulating hormone (TSH),
 - Assess for signs and symptoms of kidney cancer (for patients over age 45), including with urinalysis, and
 - For patients over age 15, assess for signs and symptoms of testicular cancer and ulcerative colitis.

2-<20 (ng/mL) PFAS*

Encourage PFAS exposure reduction if a source has been identified, especially for pregnant persons.

Within the usual standard of care clinicians should:

- Prioritize screening for dyslipidemia with a lipid panel (once between 9 and 11 years of age, and once every 4 to 6 years over age 20) as recommended by the AAP and AHA.
- Screen for hypertensive disorders of pregnancy at all prenatal visits per the American College of Obstetricians and Gynecologists (ACOG).
- Screen for breast cancer based on clinical practice guidelines based on age and other risk factors such as those recommended by US Preventive Services Task Force (USPSTF).

<2 (ng/mL) PFAS*

Provide usual standard of care



USDA USDA



National Institutes of Health



Industries

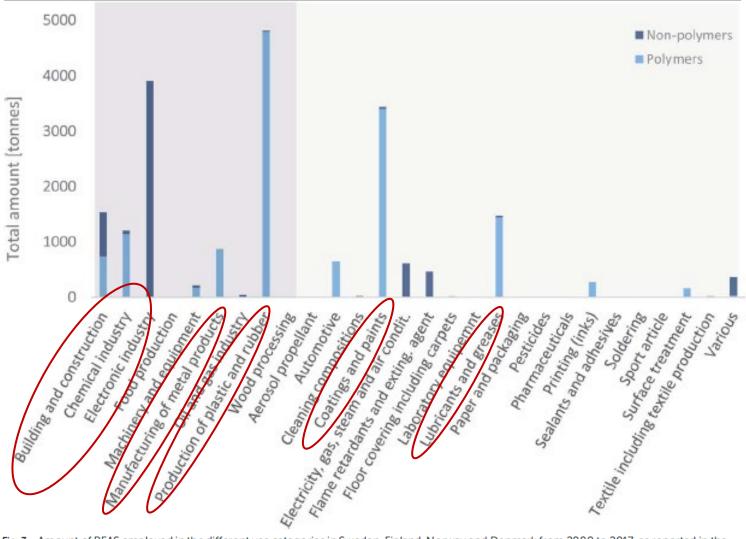


Fig. 3 Amount of PFAS employed in the different use categories in Sweden, Finland, Norway and Denmark from 2000 to 2017, as reported in the SPIN database.⁴⁴ Polymers include fluoropolymers and perfluoropolyethers. Side-chain fluorinated polymers have not been used above 0.2 t in any of the uses. Use categories with dark background are industrial branches, use categories with light grey background are other use categories.

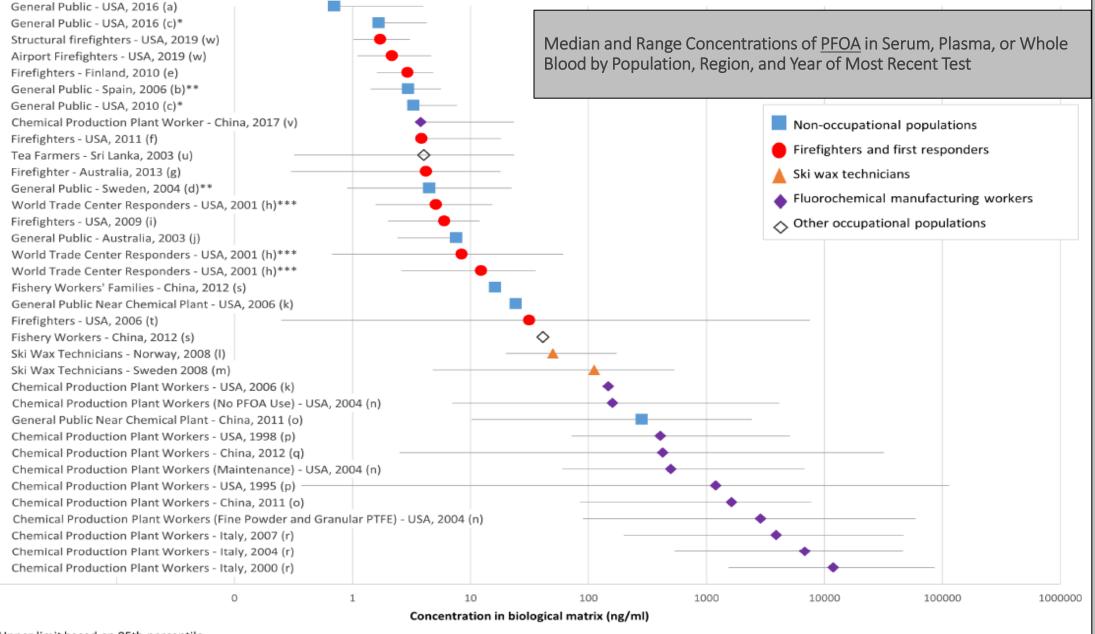
Gluge et al. 2019

Table 1 Industry branches and other use categories where PFAS were or are employed. The numbers in parentheses indicate the number of subcategories. No parentheses indicate no subcategories

Industry branches

	Aerospace (7)	Mining (3)
	Biotechnology (2)	Nuclear industry
	Building and construction (5)	Oil & gas industry (7)
	Chemical industry (8)	Pharmaceutical industry
	Electroless plating	Photographic industry (2)
	Electroplating (2)	Production of plastic and rubber
		(7)
	Electronic industry (5)	Semiconductor industry (12)
	Energy sector (10)	Textile production (2)
	Food production industry	Watchmaking industry
I	Machinery and equipment	Wood industry (3)
	Manufacture of metal products (6)	
ľ		
	Other use categories	
	Aerosol propellants	Metallic and ceramic surfaces
	Air conditioning	Music instruments (3)
	Antifoaming agent	Optical devices (3)
	Ammunition	Paper and packaging (2)
	Apparel	Particle physics
	Automotive (12)	Personal care products
	Cleaning compositions (6)	Pesticides (2)
	Coatings, paints and varnishes (3)	Pharmaceuticals (2)
	Conservation of books and manuscripts	Pipes, pumps, fittings and liners
	Cook- and bakingware	Plastic, rubber and resins (4)
	Dispersions	Printing (4)
	Electronic devices (7)	Refrigerant systems
	Fingerprint development	Sealants and adhesives (2)
	Fire-fighting foam (5)	Soldering (2)
	Flame retardants	Soil remediation

Floor covering including carpets and Sport article (7) floor polish (4)



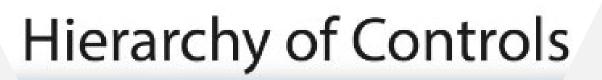
*Upper limit based on 95th percentile

**Whole blood sample. Serum concentration multipled by 1.8 for comparison to serum and plasma

***Serum sample

Occupational Exposure Limits and Guidance

- 3 ACGIH TLVs (all air)
 - Perfluoroisobutylene (PFIB)
 - Perfluorobutyl ethylene
 - Ammonium perfluorooctanoate (APFO)—a salt of PFOA
- Limited research noting protective practices
- Lacking recommendations for personal protective equipment (PPE)





Substitution

Engineering Controls

Administrative Controls

PPE

Availability of Methods

Analytical Chemistry

Targeted analyses

- Semi-volatile (LC-MS)
- Volatile (GC-MS)

•Semi-targeted analyses

•Non-targeted analyses

- Total organic fluorine (TOF)
- Particle-induced gamma-ray emission (PIGE)

Matrices

•Readily available

- Water
- Biological (blood/serum)
- •Less available
 - Air
 - Products/bulk materials
 - Dust
 - Other biological material (urine, milk, etc.)
 - Other

Routes of Exposure



Essential Use Principle

Category	Definition	PFAS examples	
(1) "Non-essential"	Uses that are not essential for health and safety, and the functioning of society. The use of substances is driven primarily by market opportunity	Dental floss, water-repellent surfer shorts, ski waxes	
(2) "Substitutable"	Uses that have come to be regarded as essential because they perform important functions, but where alternatives to the substances have now been developed that have equivalent functionality and adequate performance, which makes those uses of the substances no longer essential	Most uses of AFFFs, certain water-resistant textiles	
(3) "Essential"	Uses considered essential because they are necessary for health or safety or other highly important purposes and for which alternatives are not yet established ^{<i>a</i>}	Certain medical devices, occupational protective clothing	

Table 1 Three essentiality categories to aid the phase out of non-essential uses of chemicals of concern, exemplified with PFAS uses

^{*a*} This essentiality should not be considered permanent; rather, a constant pressure is needed to search for alternatives in order to move these uses into category 2 above.

Clarifying Questions and Discussion



Limited occupational guidance, including

OELs (3 ACGIH TLVs) Guidance control mechanisms Monitoring methods

Å

Challenges in identifying specific PFAS

Broad class of chemicals Analytical methods Changes in production over time

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Limited research on occupationally relevant exposure routes



Considered essential in some settings



NIOSH Research, Technical Support, and Strategies

Presented by

Susan Moore, PhD

Associate Director for Science

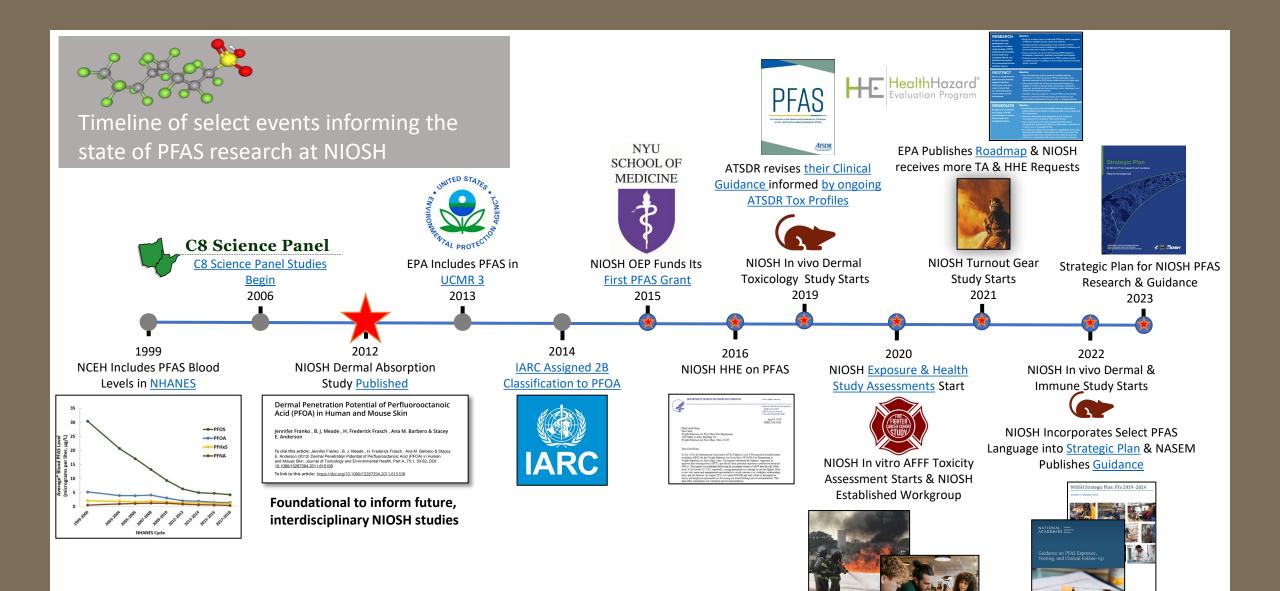
Co-Coordinator, Public Safety Sector

Co-Coordinator, Personal Protective Technology

CDC/NIOSH/NPPTL

Projects Leads

Miriam Calkins, Crystal Forester, Stacey Anderson, Lisa Weatherly, Todd Stueckle, Jen Roberts, Cindy Striley, Jess Rinsky, and Jessica Li



Exposure Assessments and Epidemiology

Project Lead: Miriam Calkins

Occupational Exposure and Health Indicator Assessment of PFAS: A Feasibility Study

Industries with moderate to high potential for worker exposure

Biomonitoring, industrial hygiene measures, and survey materials



Fire Fighter Cancer Cohort Study (FFCCS) – Exposure and Epigenetic Markers

30-year, multi-center prospective study – carcinogenic effects and cancer prevention Firefighter sub-groups and conducting assessments of fire dept practices Exposure routes and epigenetic changes



UNIVERSITY OF MIAMI





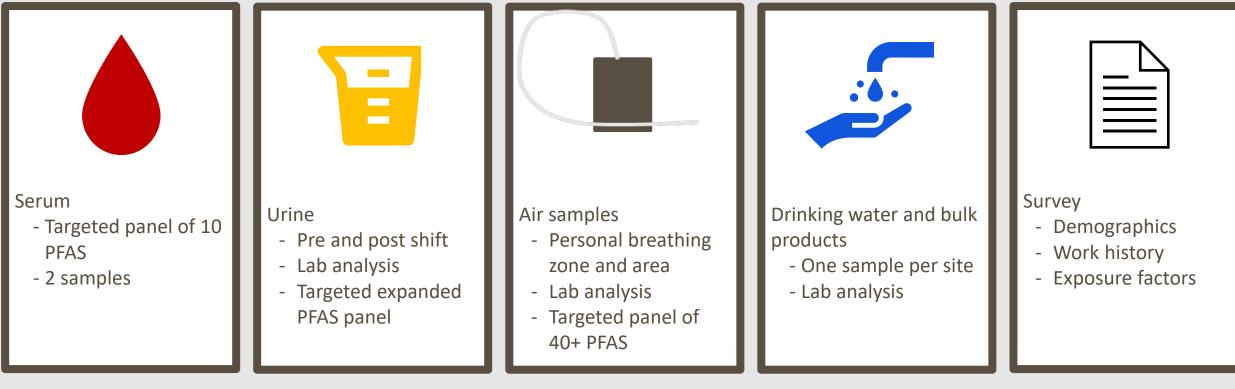




U.S. Department of Health and Human Services Centers for Disease Control and Prevention

Occupational Exposure and Health Indicator Assessment of PFAS: A Feasibility Study Ongoing

Personal measurements: two biological matrices, breathing zone samples, employment history **Worksite measurements:** drinking water and area air



- Cross-sectional study; manufacturing and services sectors
- Targeting 150 participants and 10 worksites currently in recruitment phase → reach out to Miriam Calkins for more information on how to be involved

Fire Fighter Cancer Cohort Study (FFCCS) Ongoing



Volume 13, Issue 20, October 2021, Pages 1619-1636 https://doi.org/10.2217/epi-2021-0225 Research Article Per- and polyfluoroalkyl substances, epigenetic age and DNA methylation: a cross-sectional study of firefighters Jaclyn M Goodrich 10¹, Miriam M Calkins², Alberto J Caban-Martinez 10³, Todd Stueckle⁴, Casey Grant⁵, Antonia M Calafat⁶, Amy Nematollahi⁷, Alesia M Jung⁸, Judith M Graber⁹, Timothy Jenkins¹⁰, Angela L Slitt¹¹, Alisa Dewald¹, Julianne Cook Botelho⁶, Shawn Beitel⁷, Sally Littau⁷, John Gulotta¹², Darin Wallentine¹², Jeff Hughes¹³, Charles Popp¹⁴ & lefferev L Burgess^{7,*}

WILEY
ni ¹ nD ⁹ ∋

Paola Louzado-Feliciano¹⁰ | Simi O. Oduwole¹⁰ Alberto J. Caban-Martinez DO, PhD, MPH¹⁰ ©

Future Medicine Ltd Epigenomics

<u>Example of information provided in publications</u> Serum levels from different fire departments similar to NHANES with some departments having higher levels than others

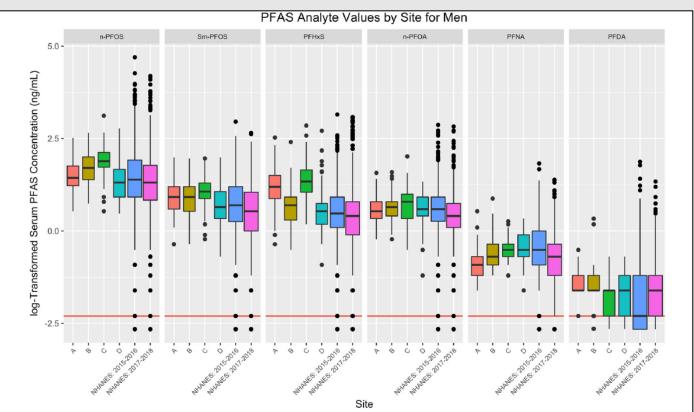


FIGURE 1 Log-transformed serum per- and polyfluoroalkyl substances concentrations (ng/ml) for men, split by site. National Health and Nutrition Examination Survey (NHANES) data are weighted. The limit of detection is plotted as a red line. Numbers of participants for each site are 77 (Department A), 59 (B), 59 (C), 61 (D), 964 (NHANES 2015–2016), and 952 (NHANES 2017–2018). [Color figure can be viewed at wileyonlinelibrary.com]

Personal Protective Equipment: Dermal exposures and materials testing Ongoing

Project Lead: Miriam Calkins

Part of FFCCS – Dermal exposure to PFAS

New garments worn by firefighters during physical activities (excluding live burn exercise)

Serum (pre and post), urine (daily), silicone wristbands (during activity)





Project Lead: Crystal Forester

Evaluation of Firefighter Textiles for PFAS

New textiles laundered

PFAS concentration on each layer determined





Acute PFAS toxicity of firefighter turnout gear

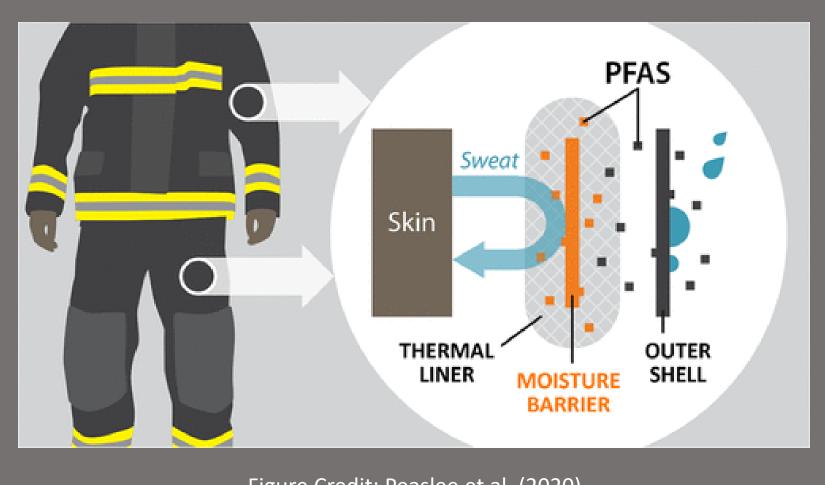
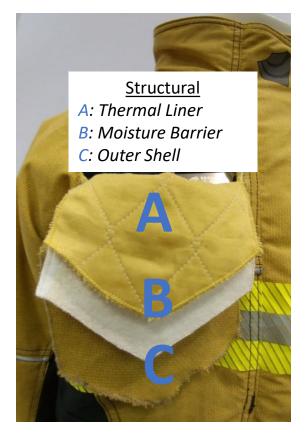


Figure Credit: Peaslee et al. (2020)

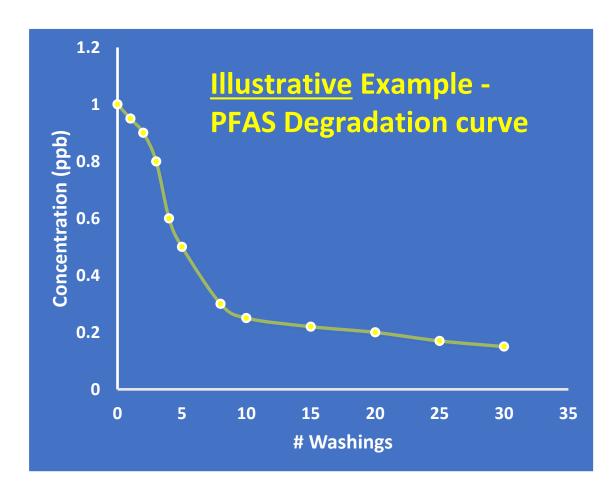
Evaluation of firefighter textiles for PFAS



Outer Shell (n=7) Scoured Outer Shell (n=2) Moisture barriers (n=5) Thermal liners (n=7



Wildland (n=5)



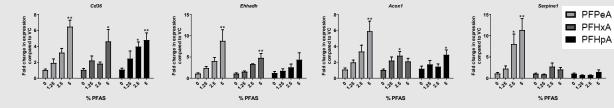
Toxicology Ongoing

Project Leads: Lisa Weatherly and Stacey Anderson

Toxicology Following Dermal Exposure to PFAS – animal study

4 Carboxylate and 4 sulfonate PFAS

Serum, urine, organ weight, histology, gene expression, skin integrity, immune phenotyping



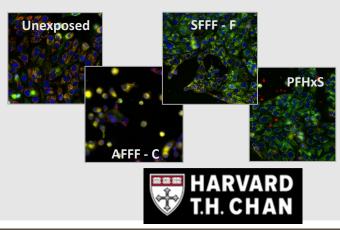


Aqueous Film Forming Foam, AFFF (n=5)

Synthetic Fluorine-Free Foam, SFFF (n=6)

Cell viability, live cell counts, mitochondrial membrane potential, intracellular reactive oxygen species, gene expression via RNAseq

Project Lead: Todd Stueckle



Toxicology following dermal exposure shows that some PFAS:

Are dermally absorbed (long- and short-chain)

Alter organs (weight, histology, phenotyping, and gene expression)

Disrupt skin (fibrosis, mild/moderate inflammation, and cell death)

Produce liver toxicities that are <u>not</u> associated with carbon chain length (sulfonic acid PFAS shows opposite association)



Food and Chemical Toxicology 156 (2021) 112528

Contents lists available at ScienceDirect

Food and Chemical Toxicology

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

Systemic toxicity induced by topical application of heptafluorobutyric acid (PFBA) in a murine model

Lisa M. Weatherly $\overset{*}{,}$ Hillary L. Shane , Ewa Lukomska , Rachel Baur , Stacey E. Anderson

Allergy and Clinical Immunology Branch, Health Effects Laboratory Division, National Institute for Occupational Safety and Health, Morgantown, WV, USA



Immunotoxicity and allergenic potential induced by topical application of perfluorooctanoic acid (PFOA) in a murine model

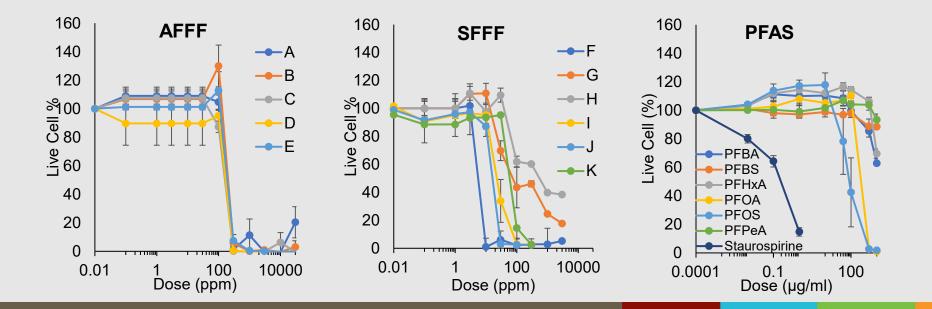
Hillary L. Shane*, Rachel Baur, Ewa Lukomska, Lisa Weatherly, Stacey E. Anderson Altergy and Clinical Immunology Branch, Health Effects Laboratory Division, National Institute for Occupational Safety and Health, Morganown, WV, 26505, USA

Acute toxicity of firefighting foams – in vitro study

Exposures to AFFF (a PFAS mixture) had substantially greater acute toxicity than single PFAS exposures but lower acute toxicity than SFFF (PFAS-free alternative)

CAVEAT 1: PFAS is only one type of surfactant in AFFF – the other surfactants and solvents may be responsible for the higher acute toxicity (being explored by ongoing work)

CAVEAT 2: Dosages tested were above recorded serum levels in firefighters (ongoing work is reducing dosage to sub-toxic levels)



Methods Development Ongoing

Air Sampling and Biological Monitoring to support other/future studies

Air sampling: develop standardized method; differentiate PFAS forms; 12 targeted PFAS analytes

Biological monitoring (animal study): support toxicology studies; high and low PFAS concentrations

Project Leads: Jen Roberts and Cynthia Striley



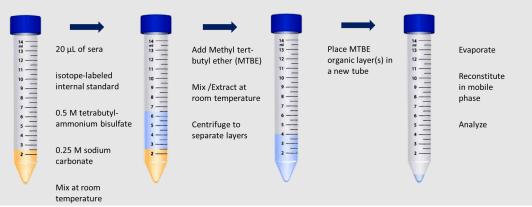


Air sampling (custom OVS-2 sampler) & analysis (LC/MS/MS)





Biological monitoring: liquid extraction from mouse serum



Technical Assistance and Health Hazard Evaluations Ongoing

Project Leads: Jess Rinsky and Jessica Li

Requests for Technical Assistance

ATSDR –recommendations to improve training materials for public safety staff (mainly firefighters) in Pennsylvania

EPA – possible PFAS exposure during field inspections



U.S. Department of Health and Human Services Agency for Toxic Substances and Disease Registry



Project Leads: Jess Rinsky and Jessica Li

Requests for Health Hazard Evaluations

2016 – acute firefighter exposure to AFFF (no PFAS-specific monitoring)

New (n=2) – airport and aerospace occupational settings



NIOSH-funded extramural outputs since 2015

Monitoring Technology



Chemical Engineering Journal Volume 417, 1 August 2021, 129133



Review

Sensors for detecting per- and polyfluoroalkyl substances (PFAS): A critical review of development challenges, current sensors, and commercialization obstacles

Colorado Ag Center at CSU (U54OH008085-17)

Exposure



Environmental Research Volume 187, August 2020, 109686



Perfluoroalkyl substances exposure and hearing impairment in US adults



Environmental Research Volume 175, August 2019, Pages 186-199



Determinants of per- and polyfluoroalkyl substances (PFAS) in midlife women: Evidence of racial/ethnic and geographic differences in PFAS exposure

Currently funded extramural projects

Exposomic Approach to Identifying WTC Exposures and Effects in Survivor Youth

exposomic Approach to id	entitying wild exposures and effects in a	Survivor Youth.
roject Number U010H012472-01	Contact PI/Project Leader HERBSTMAN, JULIE BETH	Awardee Organization COLUMBIA UNIVERSITY HEALTH SCIENCES
PROJECT SUMMARY Pregnant wo	men and children who lived, worked or attended school n	ear the World Trade Center (WTC) on September 11,
contrast to psychological effects, p understood. This is particularly imp exposures occurring during sensiti that neonates and children expose adverse health outcomes including exposure—including both psycholo effects, which are likely to emerge high-resolution methods to measur powerful exposomic and metabolo	hological and chemical exposures with potential adverse hysical health effects of early life exposure to the World ortant as the well-known developmental origins of health ve developmental windows may lead to adverse health or d to chemicalis (e.g., dioxins, perfluoroalkylsubstances (P adverse birth outcomes, altered lipid levels, and asthma gicial and chemical exposures—has been challenging, lin as exposed children age. Here, we employ advances in m ie thousands of exogenous chemicals and their endogen mic approaches have been used to enhance disease pro children and adults . Using two cohorts of WTC- exposed	Trade Center (WTC) disaster remain poorly h and disease (DoHAD) concept indicates that utcomes in later life. We have recently demonstrated PFAS)) known to be associated with WTC experience . However, characterizing the totality of WTC niting our ability to identify WTC-related health olecular laboratory technology, which have enabled ous biological responses to these exposures. These gnosis and diagnosis as well as provide insight into
targeted biomarkers associated W conventional WTC exposure measu related exogenous chemical signat children. If confirmed, the exogeno	t is associated with conventional WTC exposure measure TC exposure indices (Aim 1); 2) identify a metabolomic s ures and psychological stressors associated with WTC e ures (Aim 3a) and metabolomic signatures (Aim 3b) are us chemical and metabolomic signatures we identify in b nique fingerprints that can be used to predict adverse WT	ignature that reflects the biological response to both xposure (Aim 2); and 3) determine if WTC-exposure associated with adverse health outcomes in plood stored from these two vulnerable WTC-
1	LU010H012472-01 (fund	ed FY22)
	WTC Populations	
Prenatal WTC Chemical E	xposures, Birth Outcomes and Cardiome	etabolic Risks-Resubmission-1
Project Number 5U010H011299-04	Contact PI/Project Leader TRASANDE, LEONARDO	Awardee Organization NEW YORK UNIVERSITY SCHOOL OF

TRASANDE, LEONARDO	NEW YORK UNIVERSITY SCHOOL OF
ne, the Mailman School of Public Health, the Wads	worth Laboratories of the NYS Department of Health,
al Center respond to PAR-16-098, proposing to leve	erage two unique and contemporaneous cohorts to
essors in relationship to proximity to the WTC site a	and self-reported exposures, and evaluate birth,
her is the northern Manhattan-based Columbia Chi	delivered in one of three lower Manhattan hospitals in Idren's Environmental Health Center (NM) cohort. The ted evaluations of stress-related exposures. Except
are already available including freshly obtained me	asurements of POPs, which we will extend to include
in these cohorts, we will compare both psychosoc I were not prenatally exposed to the WTC disaster.	ial and chemical exposures and their association with This study leverages previously measured
cal exposures in relation to these endpoints in add cardiovascular endpoints reflecting environmenta	lescence. Preclinical measures included in the l influences in homogeneous populations such as
	he, the Mailman School of Public Health, the Wads al Center respond to PAR-16-098, proposing to leve ssors in relationship to proximity to the WTC site if outcomes; The first is comprised of mothers who her is the northern Manhattan-based Columbia Chi fore and after September 11, 2001 permitting ness are already available including freshly obtained me ations, neurodevelopmental outcomes have been a

5U01OH011299-04 (wrapping up, COVID-19 delay)

Michigan ERC (T42OH008455-18)

NIOSH Priorities

NIOSH Strategic Plan: FYs 2019–2024

Version 5: October 2021





















Centers for Disease Control and Prevention National Institute for Occupational Safety and Health

Management groups, labor organizations, and consensus standard bodies use NIOSH information to prevent exposures to known or suspected carcinogens among public safety workers

Updated "Need" to include, "Research assessing newer, emerging, and unstudied sources of PFAS exposure are needed to address gaps in exposure assessment, toxicology, and worker protection for the fire service."

Changes adopted

Changes on hold pending NIOSH PFAS Research Agenda

Summit on Fire Prevention and Control

National Roundtable Testimony: Firefighter Cancer

Edward Kelly, General President, International Association of Fire Fighters

"We ask the federal government to establish a comprehensive firefighter strategy that invests in research, provides access to cancer screening for all firefighters, and reduces and ultimately eliminates PFAS exposure." Consensus standard bodies, labor organizations, and management groups use NIOSH information to reduce risk factors to cardiovascular disease among public safety workers.

Requested to add content about PFAS link to CVD

Employers, workers, professional associations, policy-makers, researchers, and consensus standard organizations use NIOSH information to prevent transmission of infectious disease among public safety workers

Requested to add content about PFAS link to immune suppression

Employers, workers and their representatives, researchers, safety and health professionals, and authoritative bodies use NIOSH information to prevent

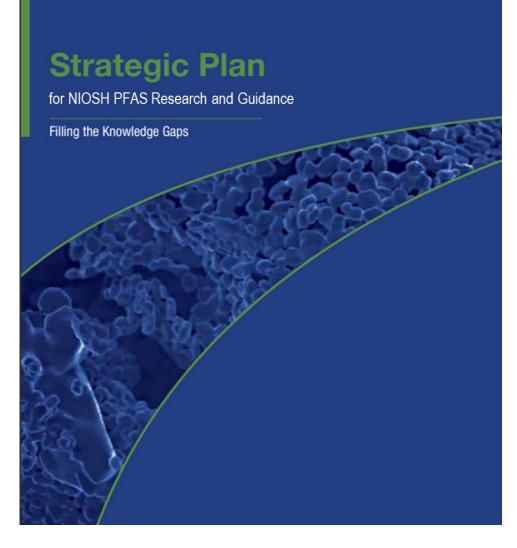
- exposures to known or suspected carcinogens among manufacturing workers
- adverse reproductive outcomes among manufacturing workers



Challenge: insufficient burden data to explicitly name PFAS *lack of data ≠ lack of an issue* Employers, policy-makers, trade associations, and manufacturers use NIOSH information to prevent immune and dermal diseases among manufacturing

Program leaders planning to request PFAS be added as example exposure leading to immune suppressive disorders (extramural research)

> Changes on hold pending NIOSH PFAS Research Agenda



In January 2023, NIOSH initiated an effort to develop PFAS Research Agenda to focus future NIOSH investments

For more information, contact

Miriam Calkins, PhD, MS NIOSH PFAS Workgroup Chair <u>MCalkins@cdc.gov</u>

Susan M. Moore, PhD NIOSH PFAS Workgroup Co-Chair <u>SMMoore@cdc.gov</u>

For more information, contact CDC 1-800-CDC-INFO (232-4636) TTY: 1-888-232-6348 www.cdc.gov

The findings and conclusions in this report are those of the authors and do not necessarily represent the official position of the National Institute for Occupational Safety and Health Centers for Disease Control and Prevention.



Questions to the BSC



With PFAS being incrementally phased out, what are the key areas of impact that can be addressed with high-quality research?



What unintended worker health or safety consequences may be created as PFAS are phased out?



What else should we consider to assist employers and protect workers against exposure to PFAS?