

Exploring Three-dimensional Bioprinting to Detect and Characterize Pathogens

Anne Arnold

Materials Scientist | National Security Directorate
December 14th, 2021, 5:00 PM PST



Community
SCIENCE & TECHNOLOGY
Seminar Series



Image courtesy of Andrea Starr

DOE's 17 **national laboratories** tackle critical scientific challenges



A regional, national, and international **scientific resource**



PNNL is DOE's **most diverse national laboratory**



5,300 Staff



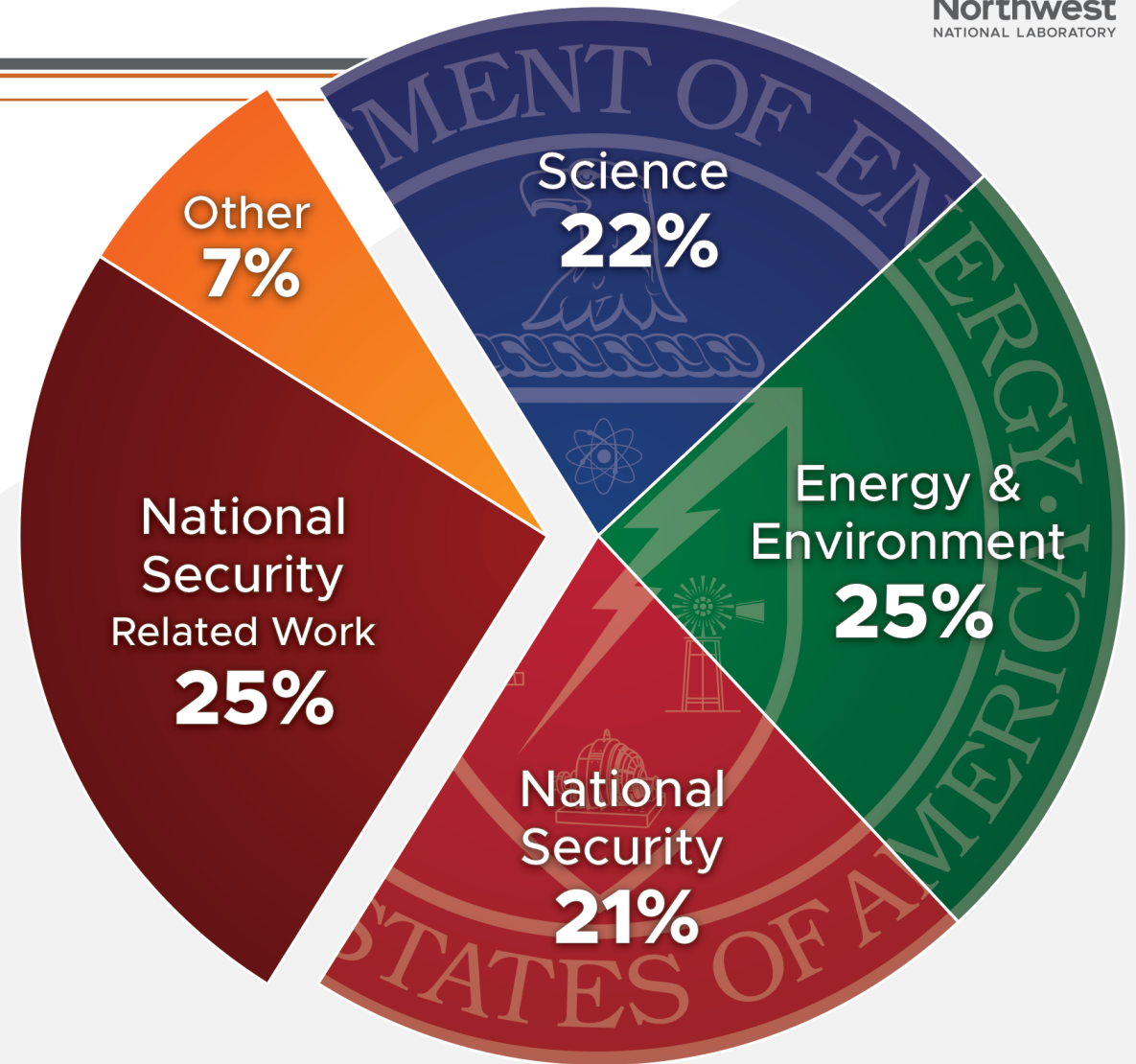
\$1.24B Spending



1,755 Peer-reviewed
Publications

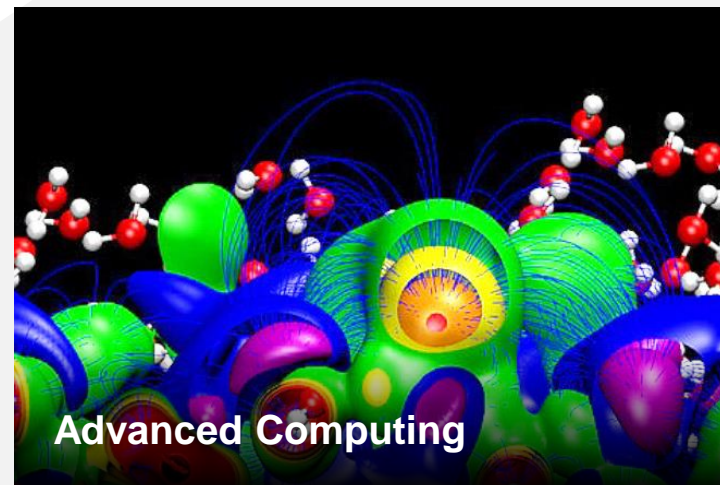
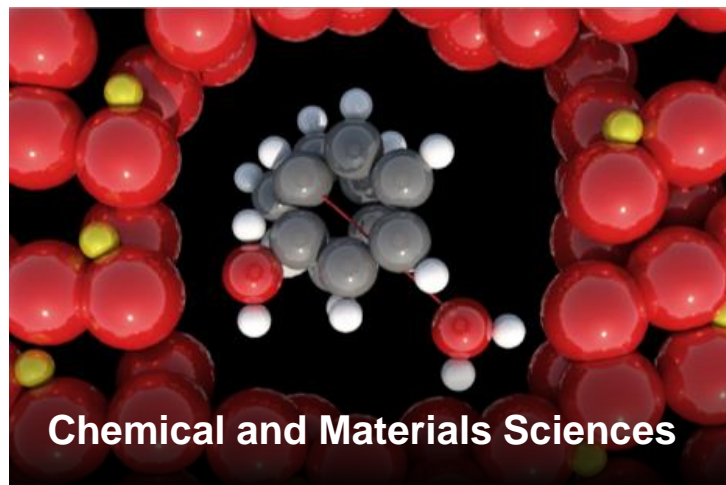
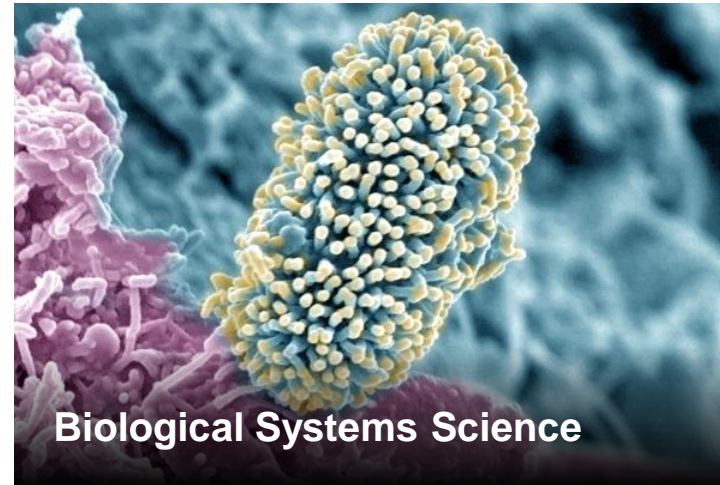
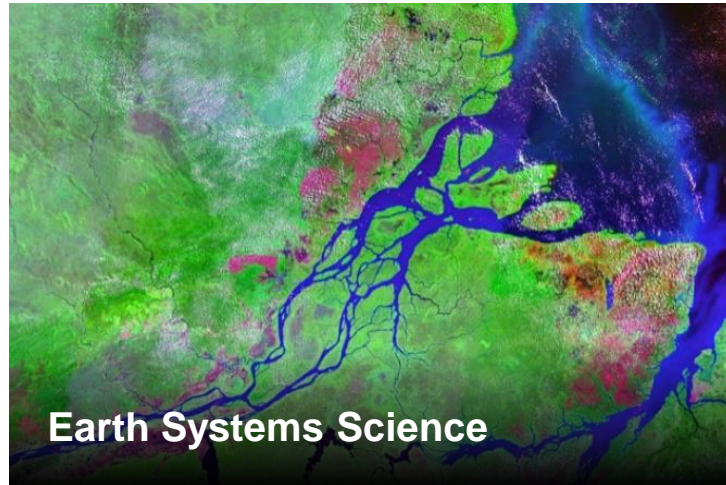


247 Invention
Disclosures

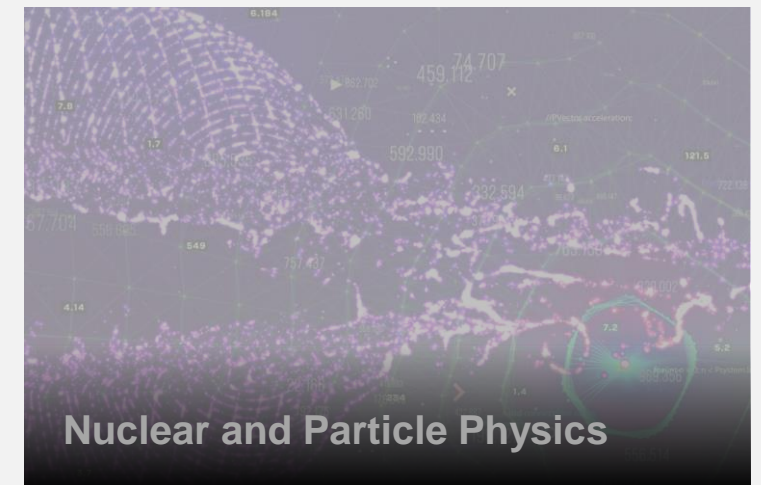
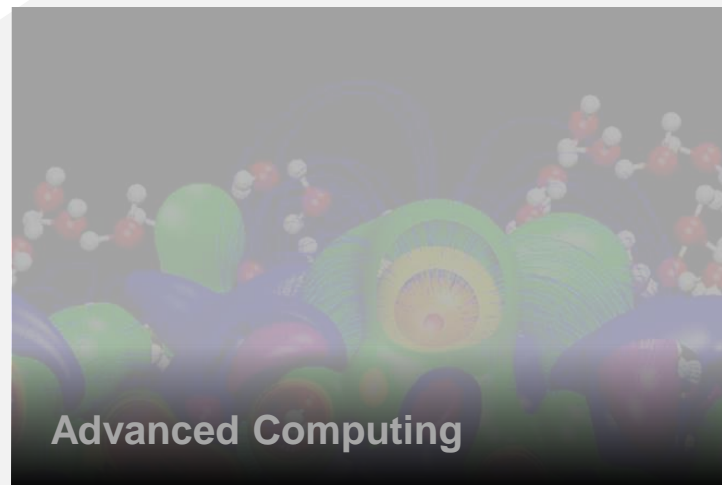
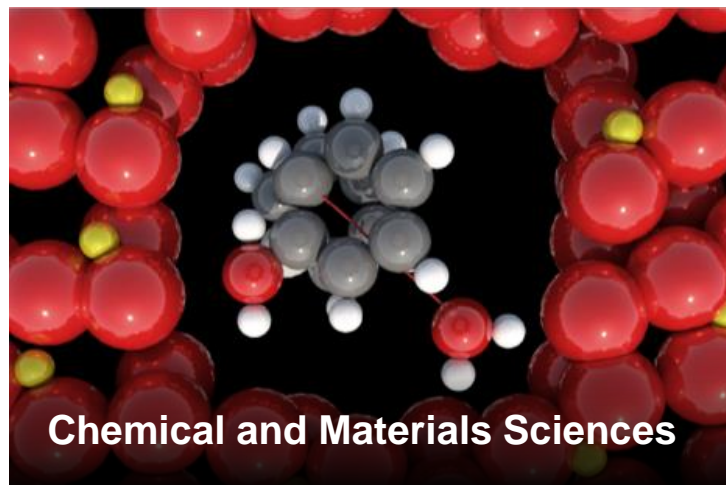
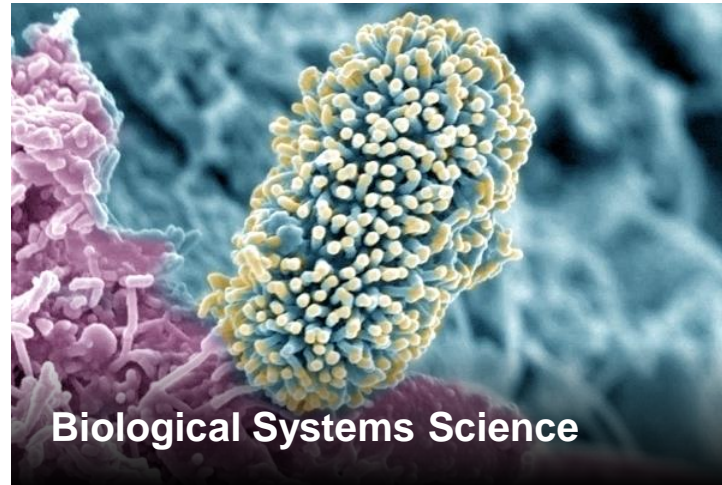


FY 2021 Staff

PNNL's **Science** mission advances understanding of the world around us



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Exploring **Three-dimensional Bioprinting** to detect and characterize **Pathogens**

1

Motivation

» Critical need for more sophisticated methods to detect and study pathogens that cause disease



2

Approach

» Creating synthetic tissue to study disease transmission using 3D bioprinting

3

Outcomes

» Developed a 3D bioprinting platform for disease detection
» Applied technology to further STEAM education

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Image courtesy of Andrea Starr

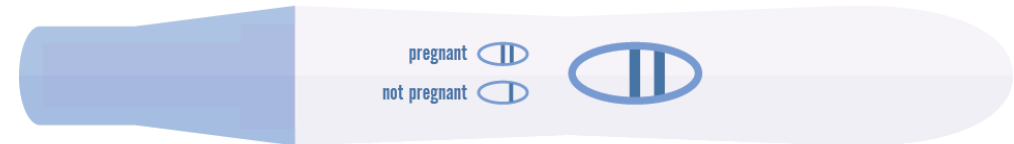
Biosensors: Nature-inspired detection

Biosensor: A device that uses a living organism (or a piece) to detect if there is a compound of interest present

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Over-the-counter pregnancy test



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Biosensors: Nature-inspired detection

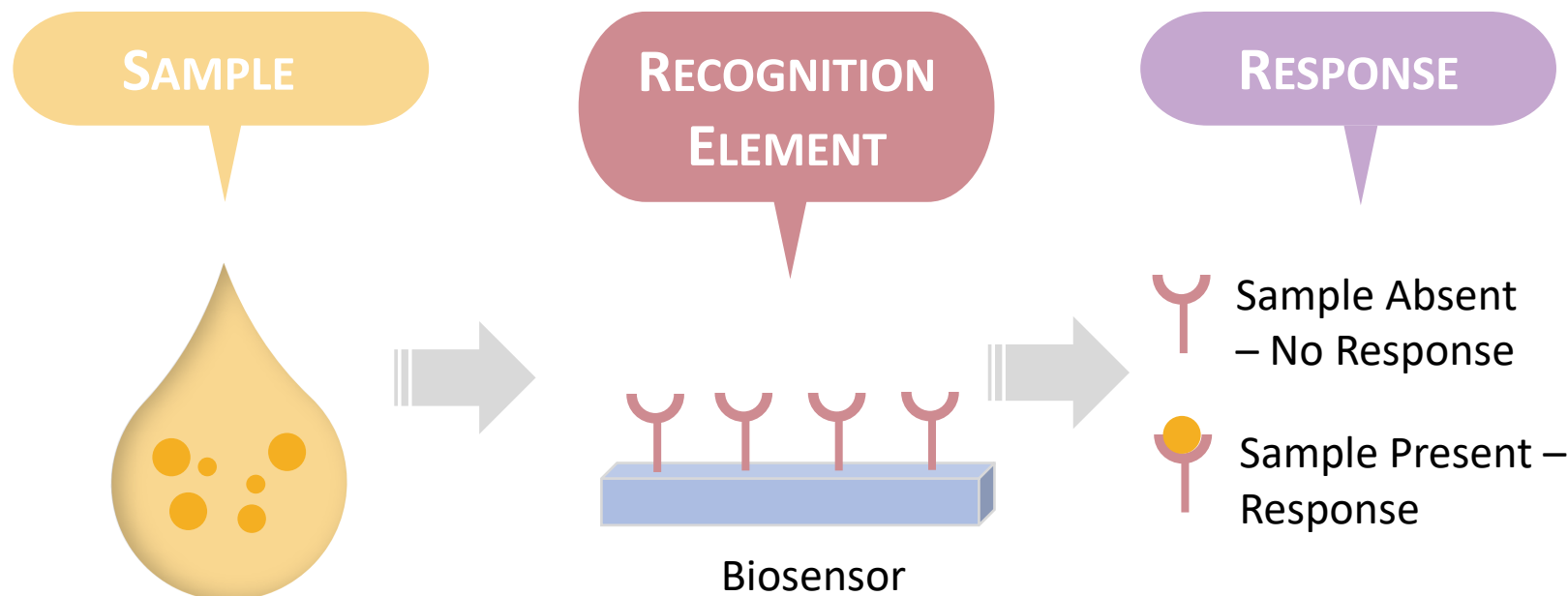
Biosensor: A device that uses a living organism (or a piece) to detect if there is a compound of interest present

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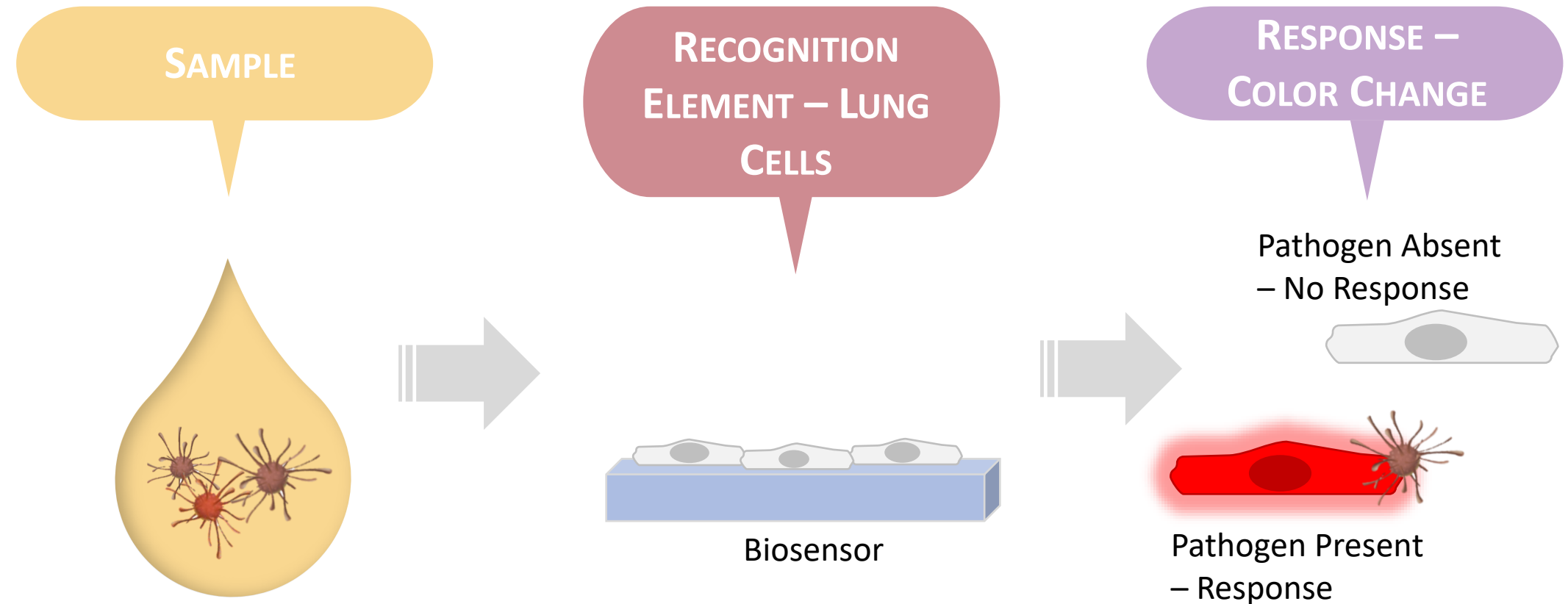
Basic biosensor components



Exploiting **human lung cells** to design **biosensors** for pathogen detection

Biosensor: A device that uses a living organism (or a piece) to detect if there is a compound of interest present

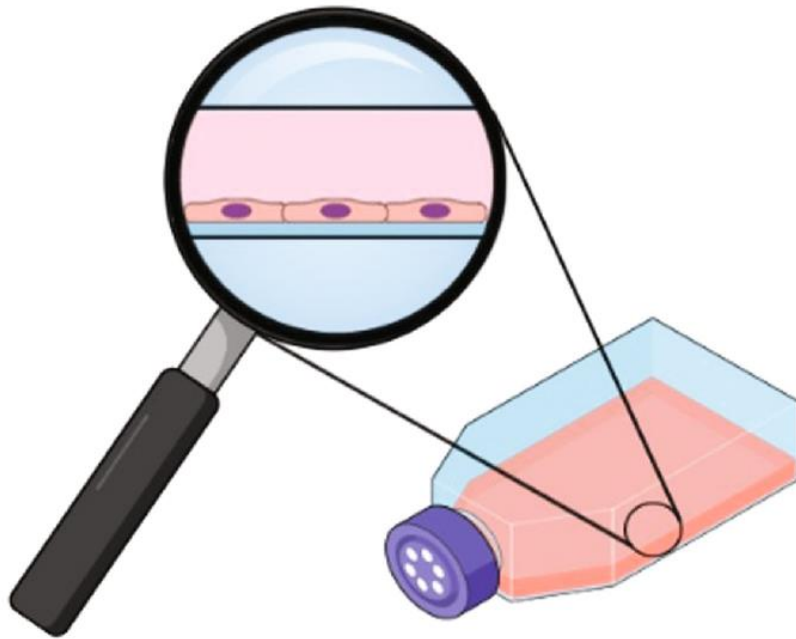
Lung cell biosensor for pathogens



Cell-based biosensors are traditionally 2D

Biosensor: A device that uses a living organism (or a piece) to detect if there is a compound of interest present

Two-dimensional cell growth in a laboratory



Cell Growth Lab

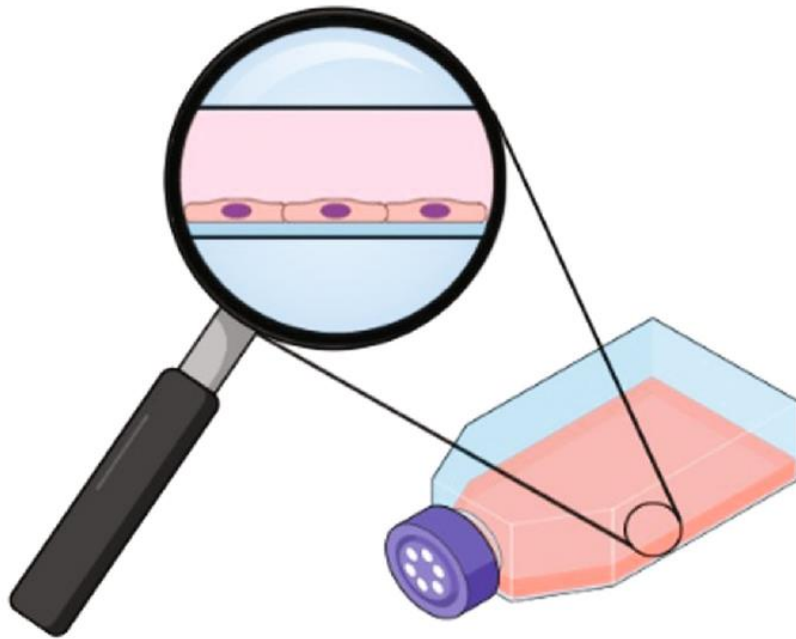


V. Charwat, D Egger. Cell Culture Technology, 2018. "The Third Dimension in Cell Culture: From 2D to 3D Culture Formats."

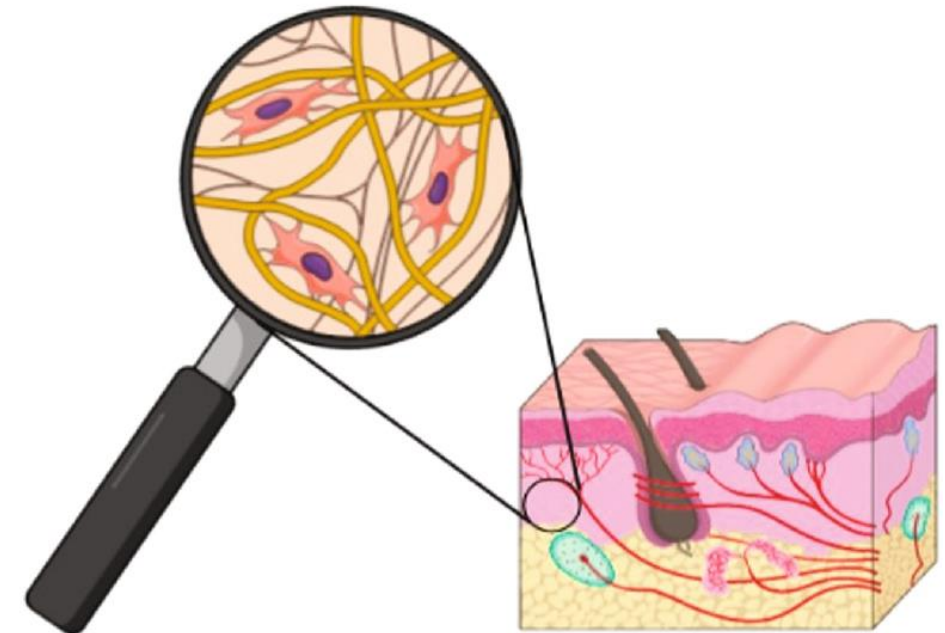
3D biosensors create new opportunities to study disease transmission

Biosensor: A device that uses a living organism (or a piece) to detect if there is a compound of interest present

Two-dimensional cell growth in a laboratory



Three-dimensional cell growth in a tissue



V. Charwat, D Egger. Cell Culture Technology, 2018. "The Third Dimension in Cell Culture: From 2D to 3D Culture Formats."

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3D Printing: New technology to create detailed, 3D objects of **any shape** or **size**



<https://www.youtube.com/watch?v=WRGJUXVXoj8>



<https://gfycat.com/gifs/search/3doodler+printing+pen>

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3Doodler Pen



<https://giphy.com/gifs/search/3doodler+printing+pen>

Traditional, plastic-based 3D printer



<https://interestingengineering.com/7-3d-printers-to-start-your-3d-printing-journey-in-2019>

3D printers that use **plastic** are **not compatible** with **cells**

Plastic filament for a 3D printer



<https://3dprinting.com/tips-tricks/choosing-a-good-filament/>

Not compatible with printing cells

1) Requires high heat to print

2) Lacks necessary permeability

3D Bioprinting: Bioinks are used as a replacement for plastics to 3D print synthetic tissue

Bioink: A thick liquid that can be combined with cells and printed into 3D shapes

Example of viscous liquids

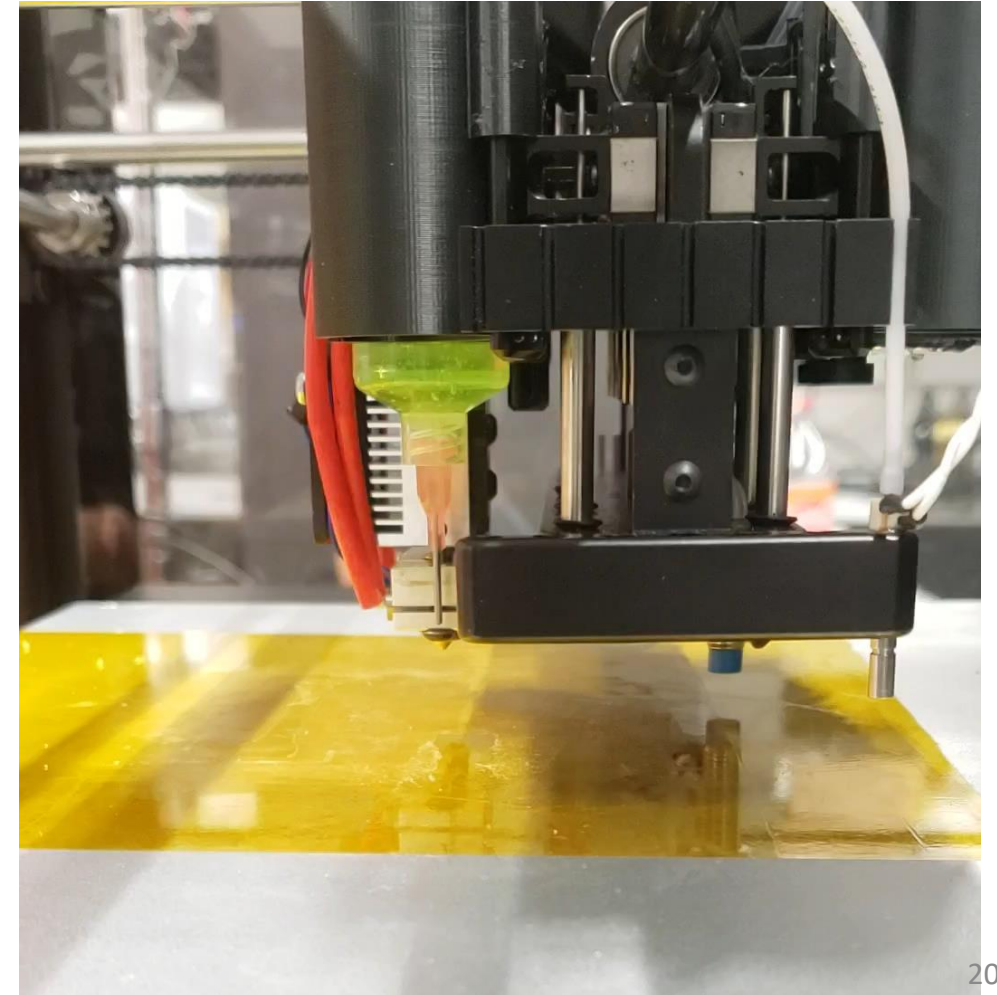


<https://energyfactor.exxonmobil.asia/science-technology/how-viscosity-helps-the-engine-go-round/>



<https://www.primogif.com/p/SZMzyHawXVmMw>

3D bioprinting a bioink



Bioink **composition** is dictated by the **application**

Bioink: A thick **liquid** that can be combined with **cells** and printed into 3D shapes

“**Liquid**” component of a bioink

Materials harvested from nature

Gelatin



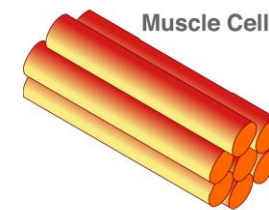
Royalty-free stock photo ID: 1840600699 by Ahanov Michael

Synthetic materials

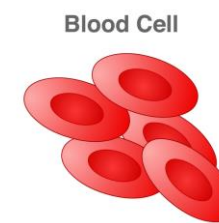


“**Cell**” component of a bioink

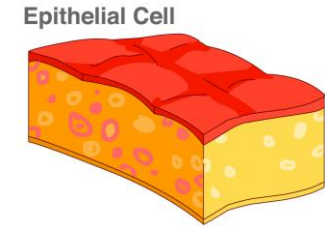
Human cells



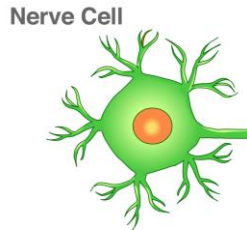
Muscle Cell



Blood Cell



Epithelial Cell



Nerve Cell

Royalty-free stock vector ID: 1544771027 by grayjay

Bacterial cells



Clostridium
botulinum

Royalty-free stock vector ID: 1496096153 by Elena Istomina



Escherichia coli



Salmonella typhi

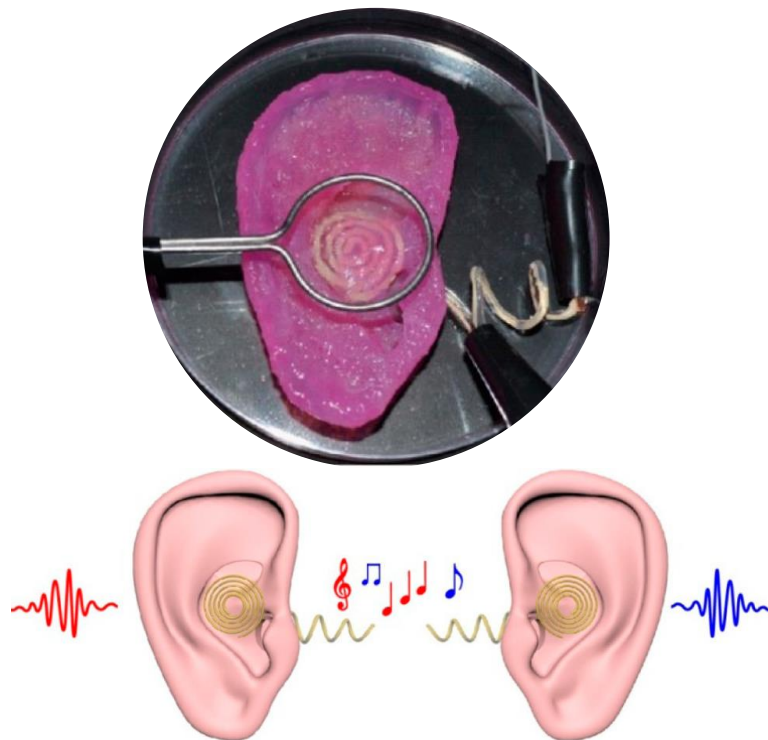


Clostridium
tetani

Bioink **composition** is dictated by application

Bioink: A thick **liquid** that can be combined with **cells** and printed into 3D shapes

3D printed bionic ears



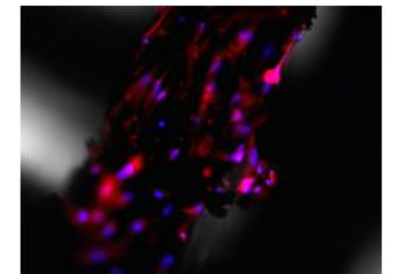
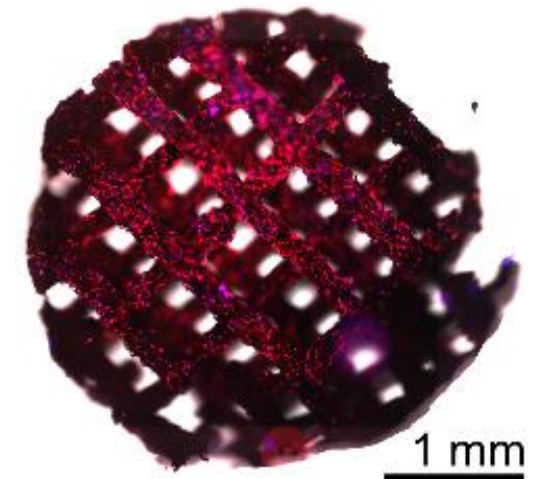
Nano Lett. 2013, 13, 6, 2634-2639

3D printed human heart



ACS Biomater. Sci. Eng. 2020, 6, 11, 6453-6459

3D printed bone



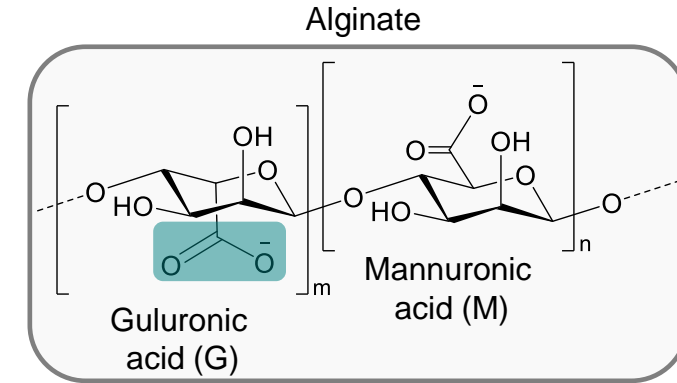
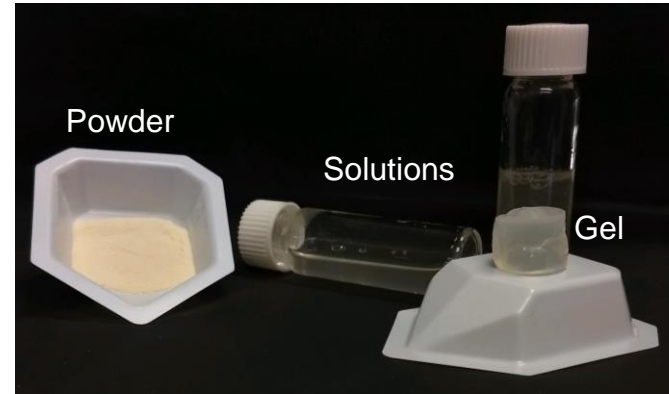
200 μm
L. Daneshmandi, et al. Unpublished

Our **bioink** is composed of **alginate**, a naturally-derived polymer

Brown seaweed



Public Domain, <https://commons.wikimedia.org/w/index.php?curid=408029>

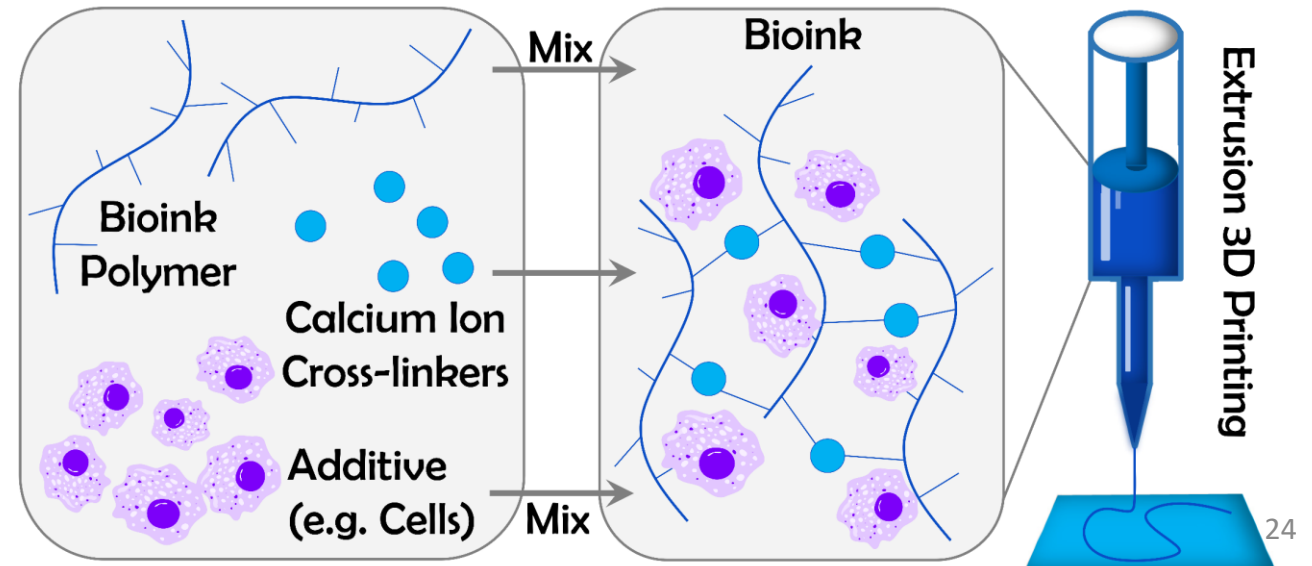
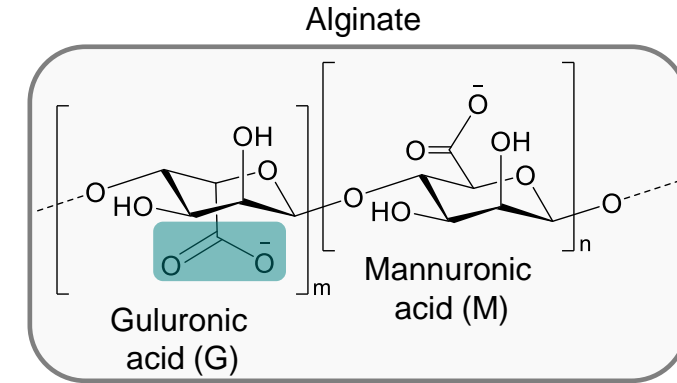
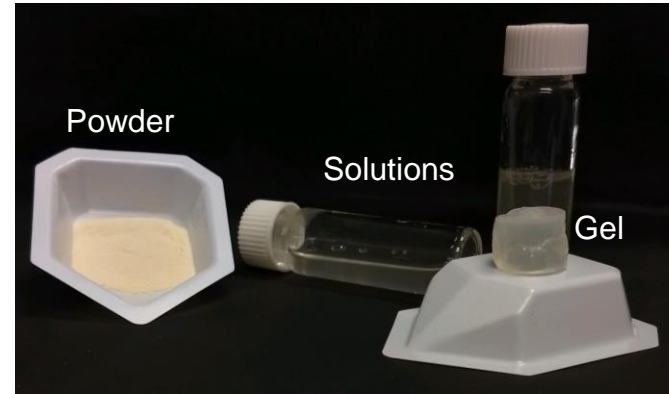


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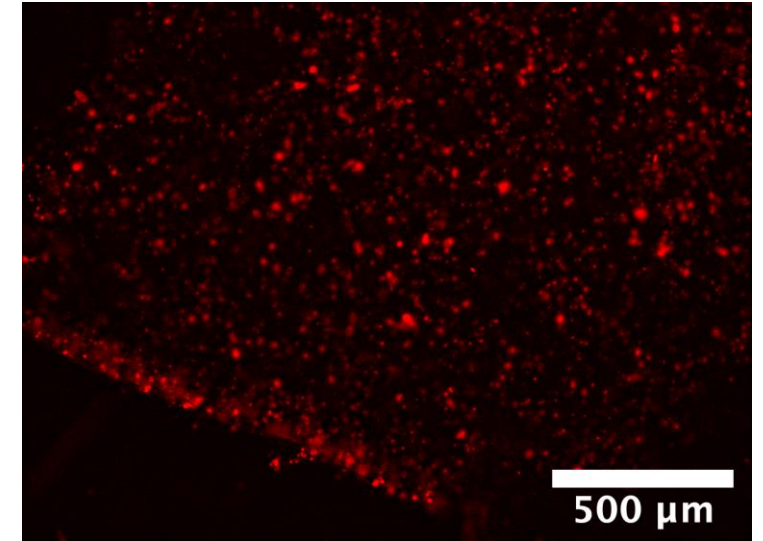
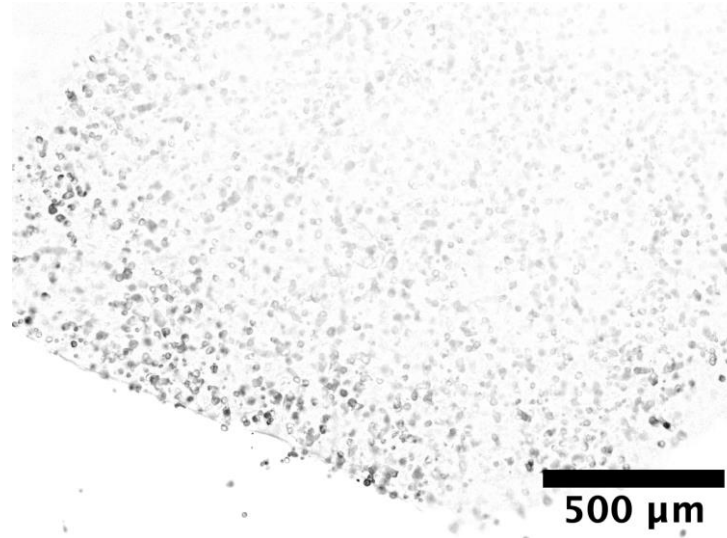


Significant work to **refine bioink** formulation for our application

Early bioink recipe



Lung cells inside bioink



Refined bioink recipe

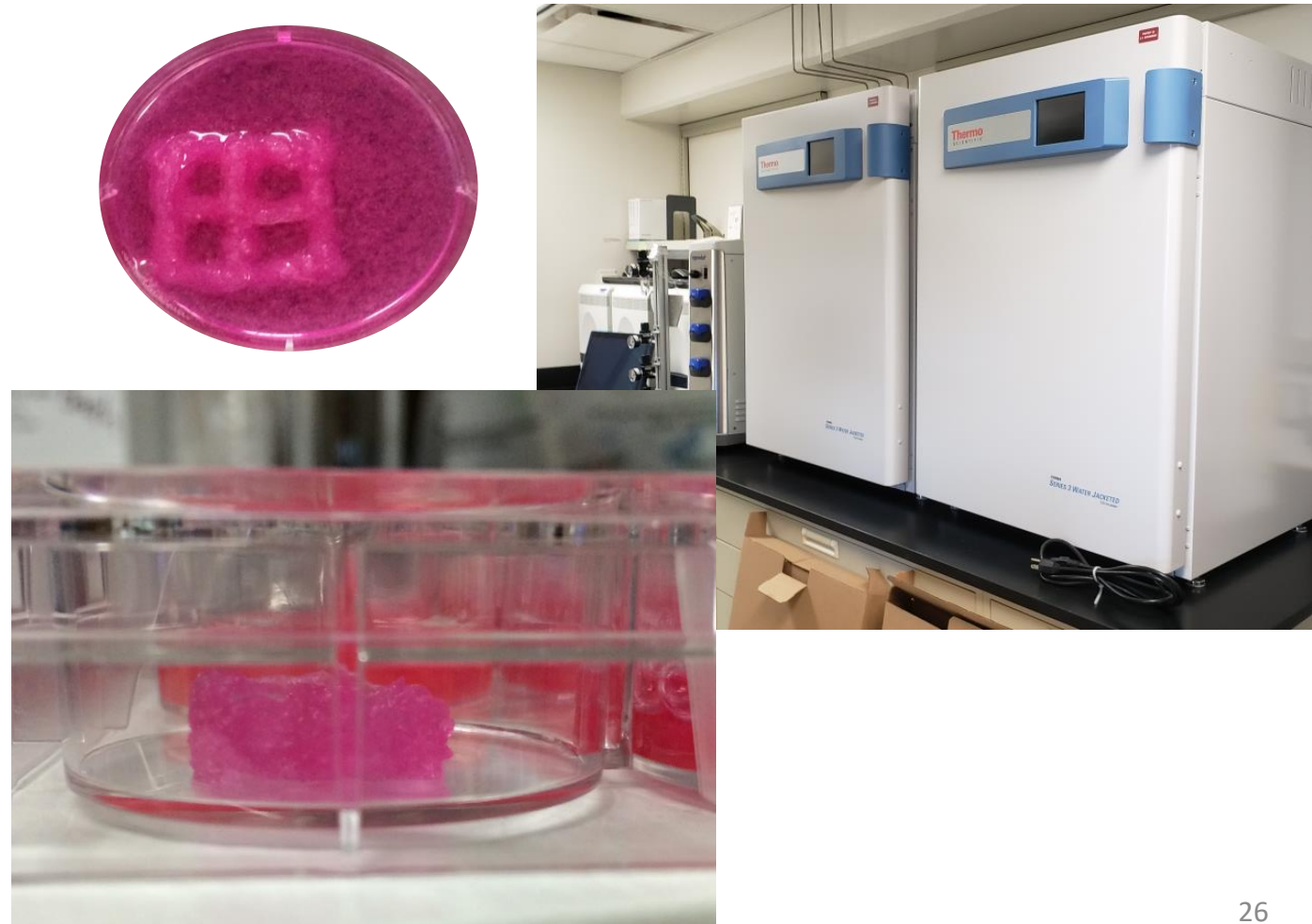


Significant work to **refine bioink** formulation for our application

Refined bioink recipe



Synthetic tissue prints



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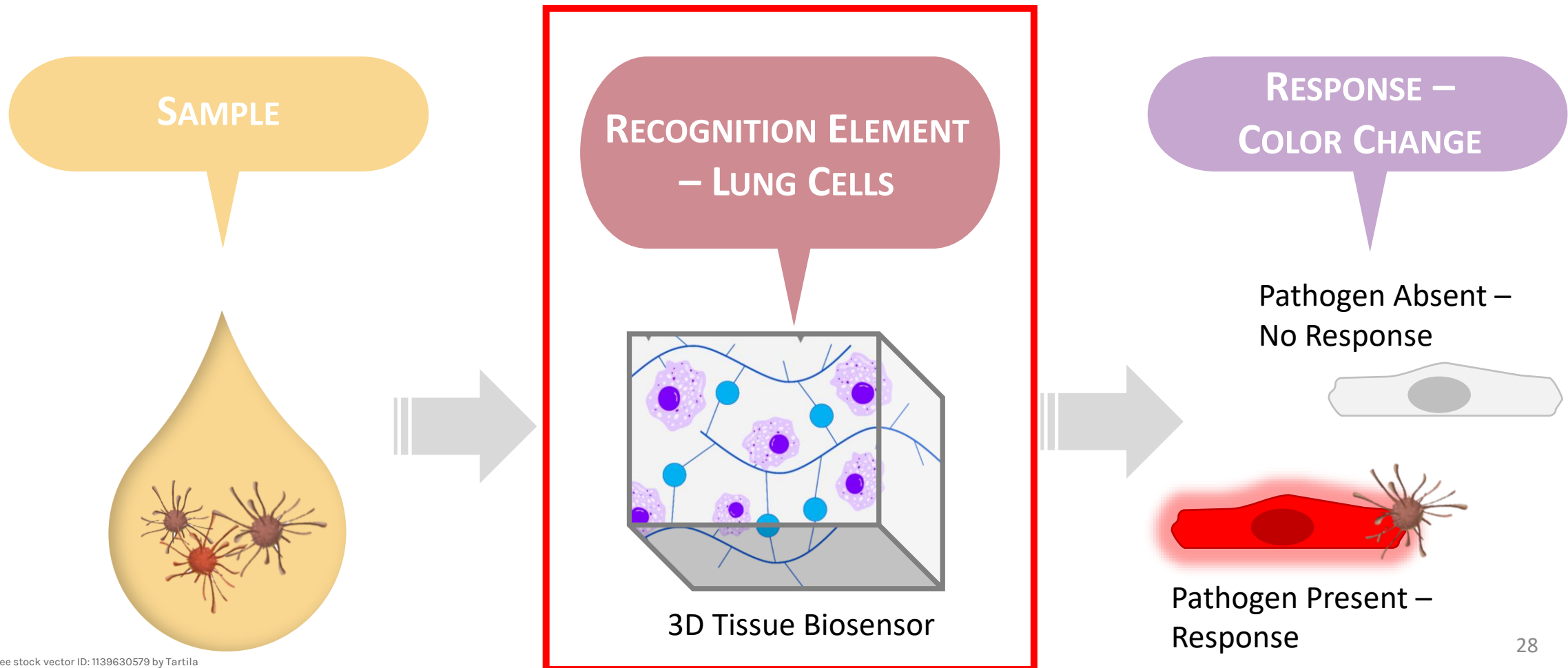
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Outcomes

» Developed a 3D bioprinting platform for disease detection
» Applied technology to further STEAM education

Developed a **3D bioprinting platform** to print synthetic lung tissue for disease detection

Biosensor: A device that uses a living organism (or a piece) to detect if there is a compound of interest present



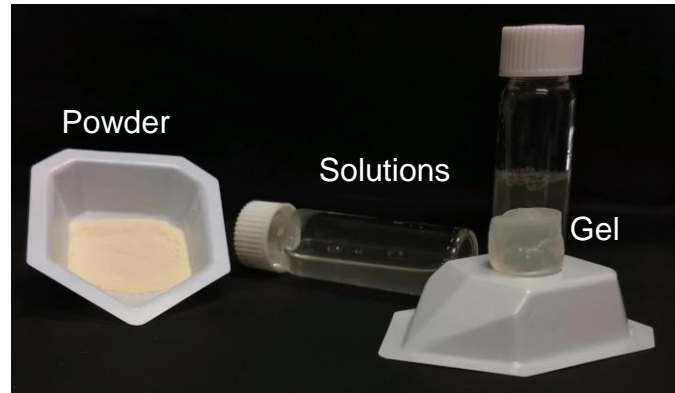
Increasing
knowledge and
interest in **STEM**
through hands-on
learning with PNNL's
STEM
Ambassadors



Photo taken before the COVID-19 pandemic

Transforming our **bioink** recipe into **STEAM** educational tools

Alginate bioink



Mica powders



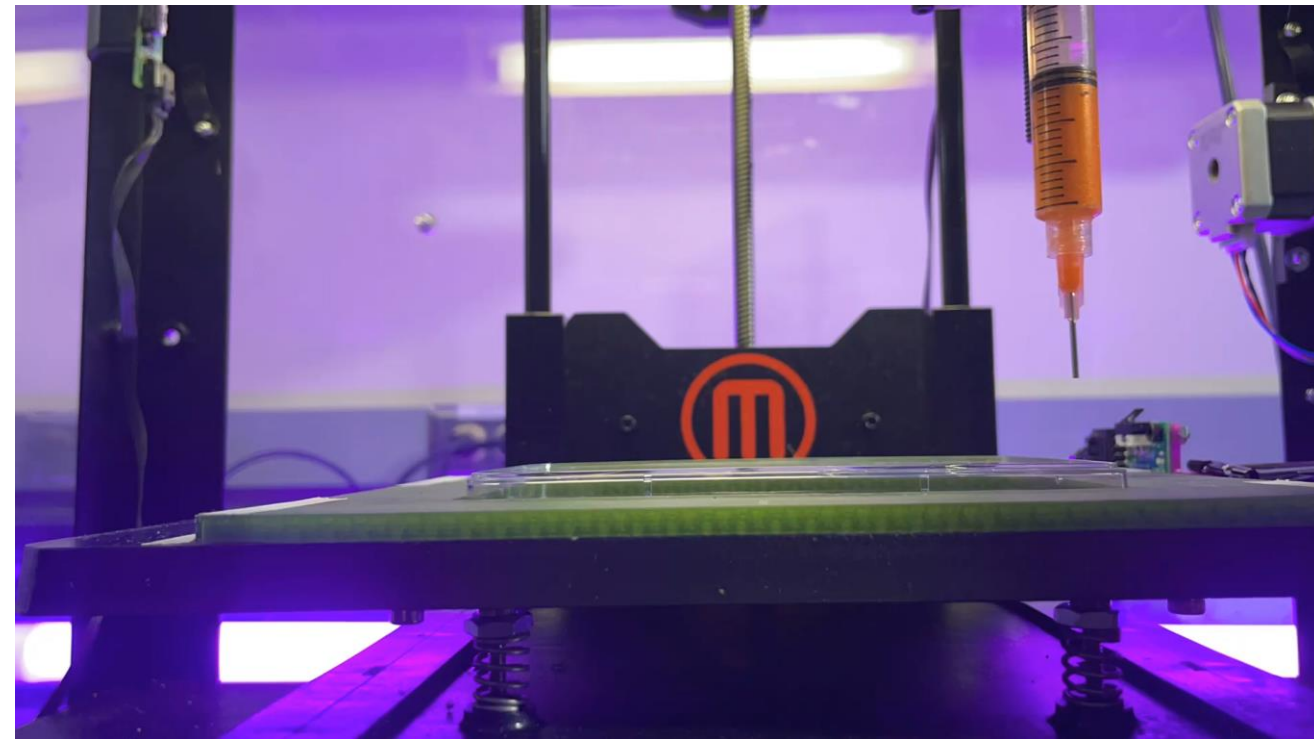
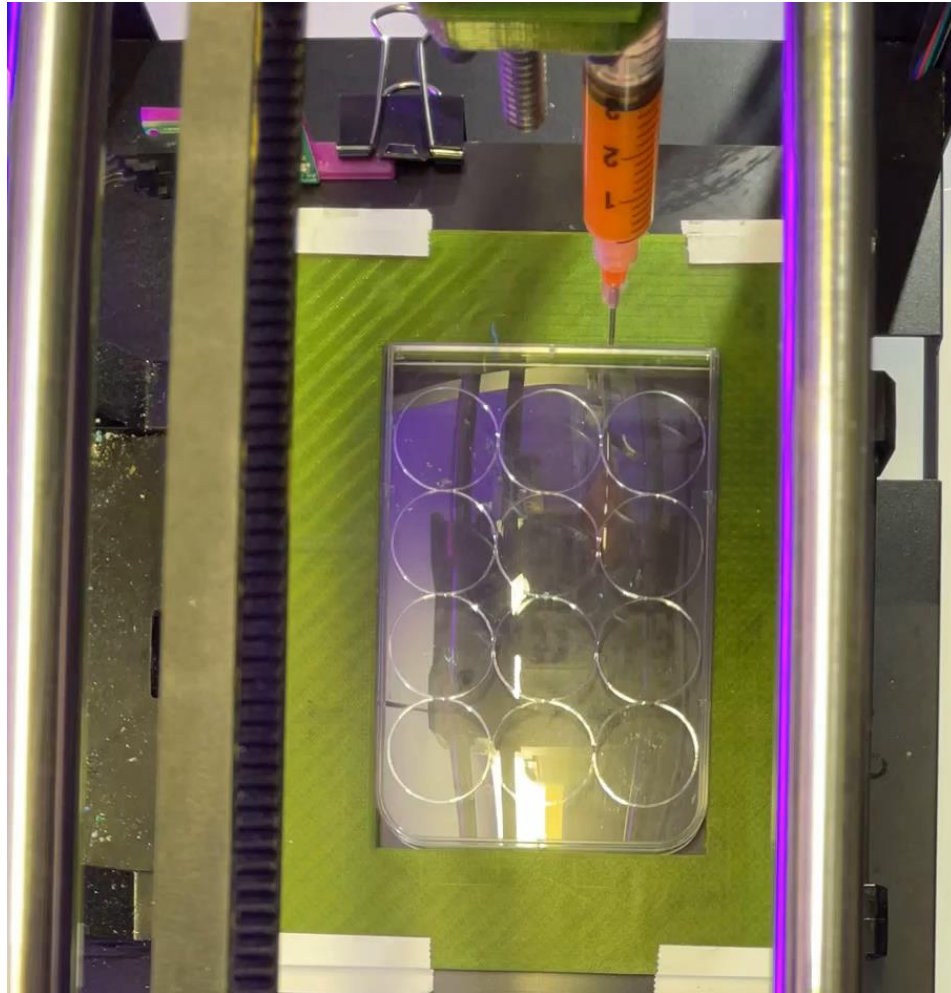
Vibrantly colored
bioinks



Images courtesy of Andrea Starr

Transforming our **bioink** recipe into **STEAM** educational tools

3D printing bioink art media



Videos courtesy of Graham Bourque

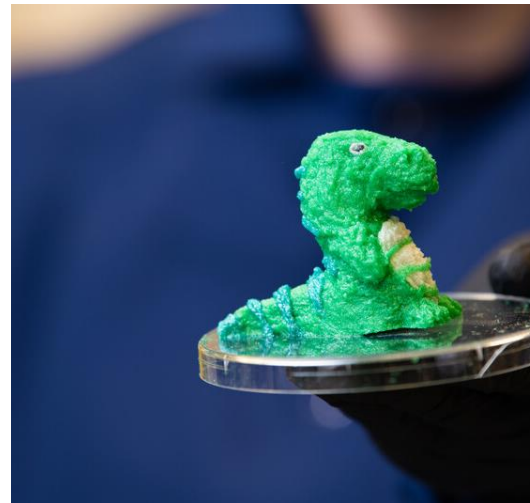
Transforming our **bioink** recipe into **STEAM** educational tools

3D printing bi



Videos courtesy of Graham Bourque Image courtesy of Andrea Starr

Transforming our **bioink** recipe into **STEAM** educational tools



Images courtesy of Andrea Starr

Our PNNL Team

- Janine Hutchison – Project Lead
- Shelby Phillips – New Hire
- Carson Bergstrom – New Hire
- Fatous Ndiaye – MSIP intern
- Becky Hess
- Sef Christ
- **Michelle Fenn – Junior Staff**
- Zack Kennedy
- Loreen Stromberg – New Hire
- Rebecca Erikson



A multidisciplinary team has all the roles it needs to design, build and operate a service

@ewebber

Thank you



Image courtesy of Andrea Starr