#### Immunology 101: Implications for Medical Device Failure

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# Learning Goals

• Understand innate and adaptive immunity

• Explain how innate and adaptive immunity work together to fight infections

• Discuss how the inflammatory response can be regulated

### **Basic Immunity**



# **Barriers to infection**

- Epithelial layers produce protective substances
  - S. aureus E. coli Inoculation 30 minutes – Antimicrobial peptides • Psoriasin **Fresh culture plates** Incubate E. coli S. aureus
- Why does our skin secrete psoriasin?

Acidic pH

# Immunity





#### Innate Pathogen Recognition



- Innate Immune System
  - <u>PAMPs</u> Pathogen-Associated Molecular Patterns
  - <u>**PRRs</u>** Pattern Recognition Receptors</u>
    - Germ-line encoded recognition molecules
  - Host-Pathogen "Arms" Race

## Inflammatory Response



## Hallmarks of Inflammation



## Phagocytosis



## Neutrophil - Phagocytosis

![](_page_9_Picture_1.jpeg)

https://www.youtube.com/watch?v=Z\_mXDvZQ6dU

#### Adaptive Pathogen Recognition

![](_page_10_Picture_1.jpeg)

Antibodies (products of B cells)

- Adaptive Immune System
  - Randomly generated (B and T cell receptors)
    - These bind to very specific antigens, rather than generic molecules found on many pathogens

![](_page_11_Figure_0.jpeg)

#### **Adaptive Memory**

- Memory is the hallmark of adaptive immunity
  - Reason why vaccines work!

![](_page_12_Figure_3.jpeg)

### Teamwork is Key!

- Innate and adaptive immunity work cooperatively
  - Activation of innate immune responses produces <u>cytokines</u> which communicates with the adaptive immune system

![](_page_13_Figure_3.jpeg)

![](_page_14_Picture_0.jpeg)

## Implantable Biomaterials and Biocompatibility

![](_page_15_Picture_1.jpeg)

![](_page_15_Picture_2.jpeg)

![](_page_15_Picture_3.jpeg)

![](_page_15_Picture_4.jpeg)

![](_page_15_Picture_5.jpeg)

![](_page_16_Figure_0.jpeg)

![](_page_16_Figure_1.jpeg)

Inflammatory Cells

Pro-inflammatory Cytokines Reactive oxygen species (ROS)

![](_page_16_Picture_4.jpeg)

Blood Protein Adsorption

Slee JB et al. Polymers 2014

## Biocompatibility

The condition of being compatible with living tissue or a living system by not being toxic or injurious and not causing immunological rejection

## Resveratrol

![](_page_18_Picture_1.jpeg)

![](_page_18_Figure_2.jpeg)

- Produced in plants during environmental stress
- Consuming red wine in high amounts (20-30 g per day) can lower the risk of coronary heart disease, a type of CVD, by at least 40%

# Resveratrol and Endothelial Cell Morphology

![](_page_19_Figure_1.jpeg)

1 μM Resveratrol

100 μM Resveratrol

Greco G. & Slee JB. In Prep.

## **Resveratrol and Cell Stress**

![](_page_20_Picture_1.jpeg)

Blue: Nucleus Green: F-actin

- Tumor Necrosis Factor-  $\alpha$  (TNF-  $\alpha$ ) was used to induce cell stress
- Induces actin stress fiber accumulation in BAECs, mimicking the inflammatory response

#### **Resveratrol and Cell Stress**

![](_page_21_Figure_1.jpeg)

+ Resveratrol + TNF-  $\alpha$ 

Greco G. & Slee JB. In Prep.

# Wound Healing

![](_page_22_Picture_1.jpeg)

- Resveratrol facilitates the activation of vascular endothelial growth factor (VEGF) expression
- VEGF promotes angiogenesis

#### Wound Healing

![](_page_23_Figure_1.jpeg)

Greco G. & Slee JB. In Prep.

### **THP-1** Adhesion Assay

![](_page_24_Figure_1.jpeg)

#### **THP-1** Adhesion Assay

![](_page_25_Figure_1.jpeg)

Greco G. & Slee JB. In Prep.

# Chandler Loop Assay

![](_page_26_Picture_1.jpeg)

#### Chandler Loop Assay

![](_page_27_Figure_1.jpeg)

10 µM

**Resveratrol Treatment Concentration** 

1 μM

Control

\* represents p<0.01. P values determined using ANOVA with Tukey Post-Hoc Test

Greco G. & Slee JB. In Prep.

# Summary

- Pathogens and foreign objects trigger an inflammatory response in the body.
- Innate Immunity is the "non-specific" way the body deals with eliminating pathogens and foreign objects from the body.
- Adaptive Immunity is the "specific" way the body deals with eliminating pathogens from the body.
- Resveratrol from red wine and grapes possesses antiinflammatory properties that are worth investigating futher.

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**Thank You!** 

**Questions?** 

![](_page_29_Picture_1.jpeg)

**GH** The Children's Hospital *of* Philadelphia<sup>®</sup> Hope lives here.