

# Comprendre la progression des maladies vasculaires grâce à la simulation numérique

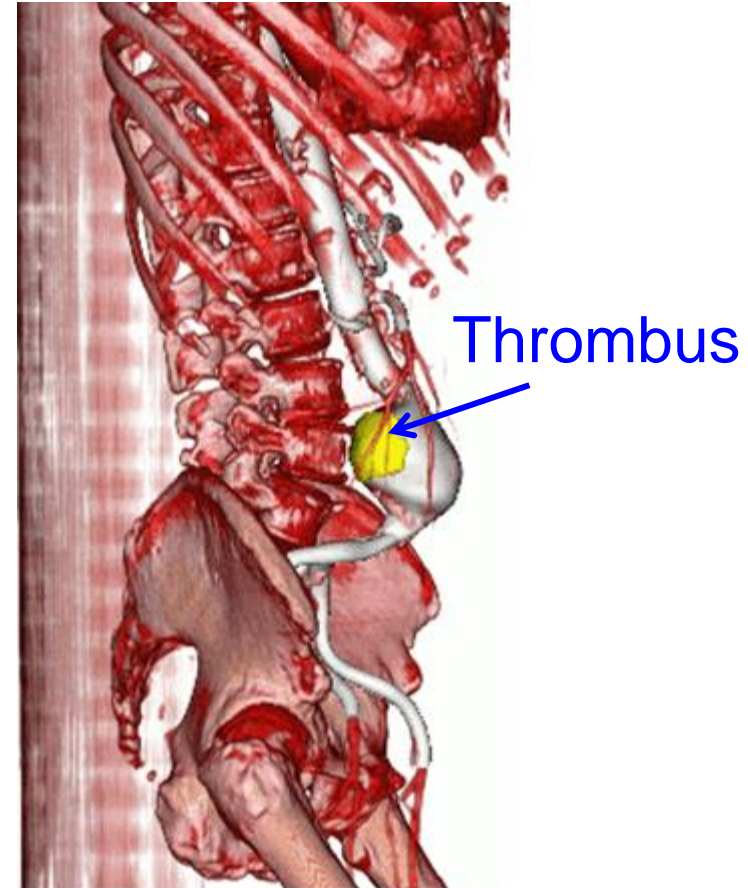
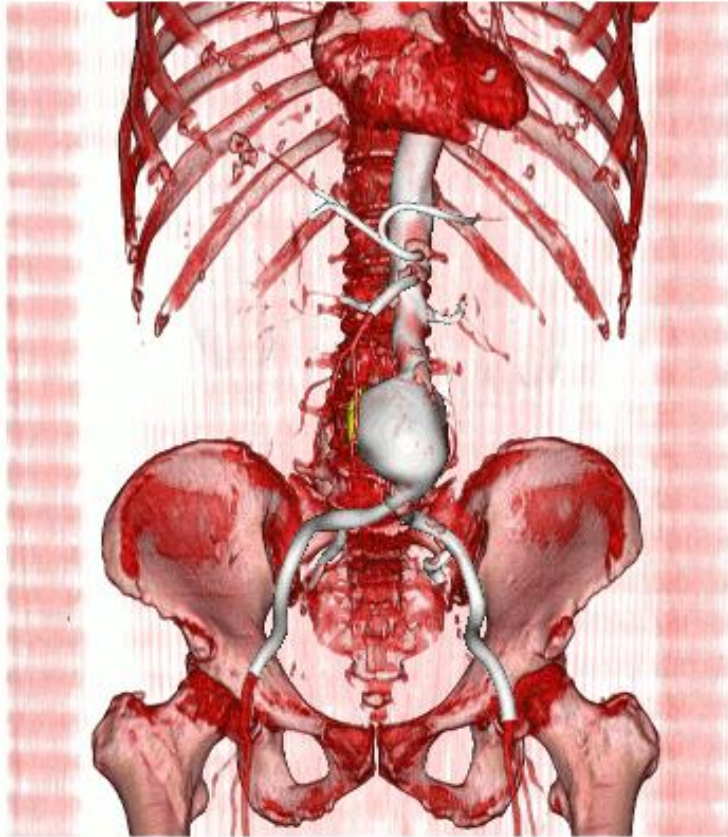
(Understanding vascular disease progression via advanced computational modeling)

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Vascular Biology and Therapeutics Program,  
Yale University, USA

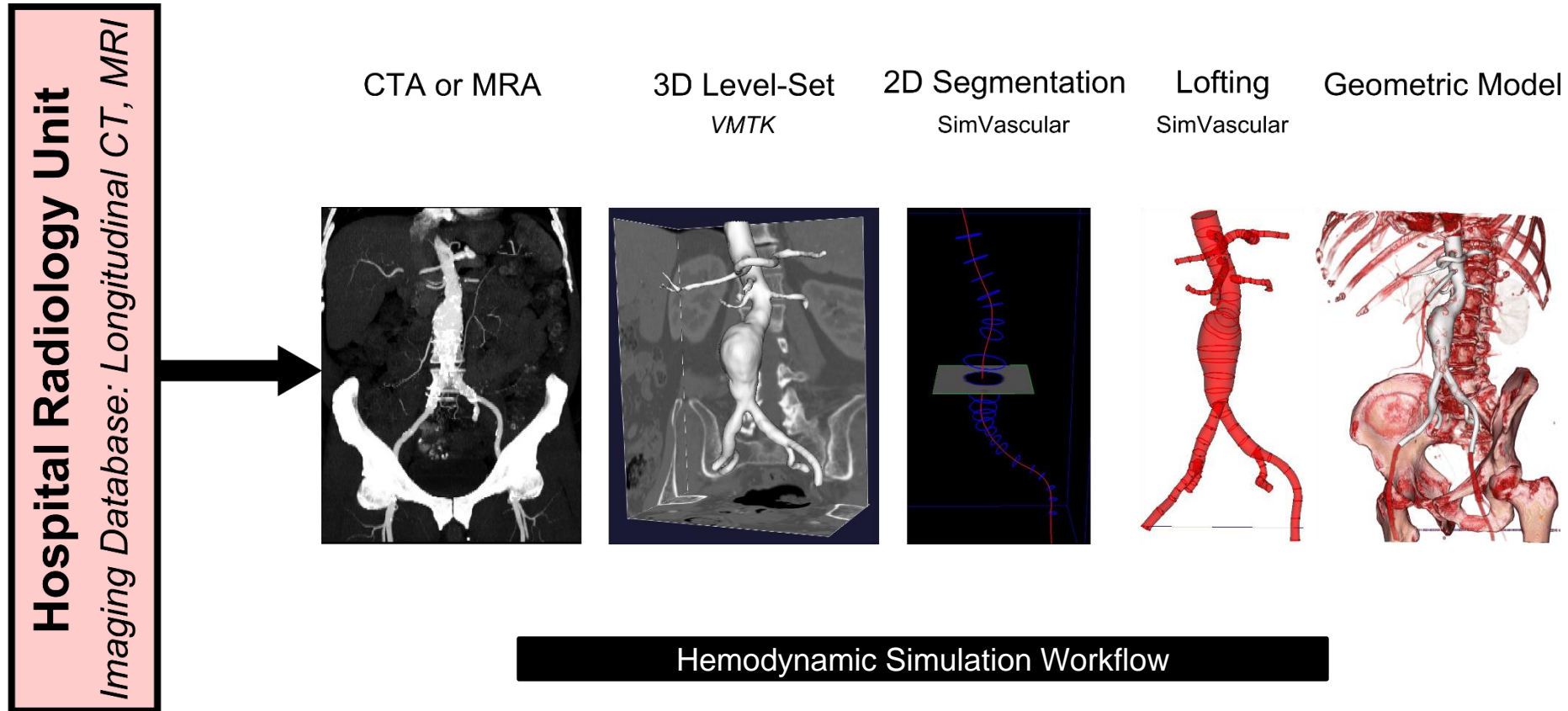
TERATEC 2017, Paris, France

# The Clinical Dilemma - Abdominal Aortic Aneurysm (AAA)



AAA2 (6 cm maximum diameter)

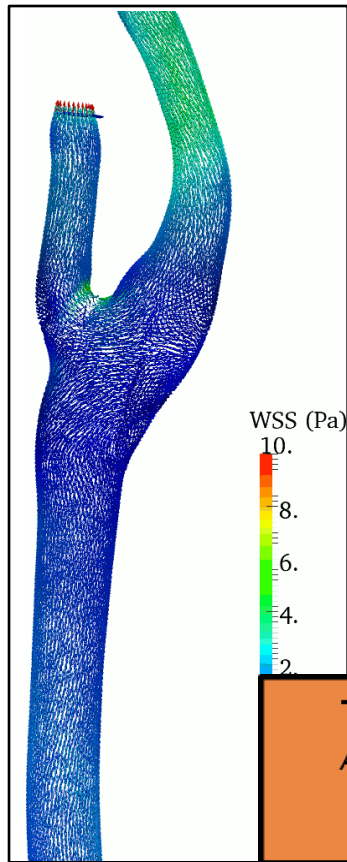
# Computational Model Construction



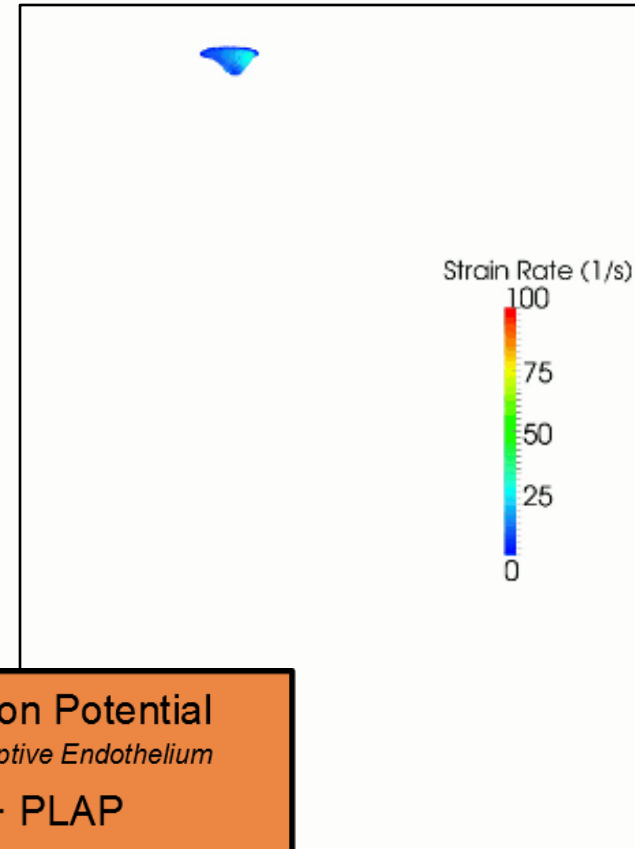
*data collection*

# The Thrombus Formation Potential (TFP)

*Carotid Bifurcation*



*Abdominal Aortic Aneurysm*



**Thrombus Formation Potential**

*Activated Platelets on Receptive Endothelium*

$$\text{TFP} = \text{ECAP} \cdot \text{PLAP}$$

**Endothelial Cell Activation Potential**

*Low and Oscillatory Shear Stress Regions*

$$\text{ECAP} = \frac{\text{OSI}}{\text{TAWSS}}$$

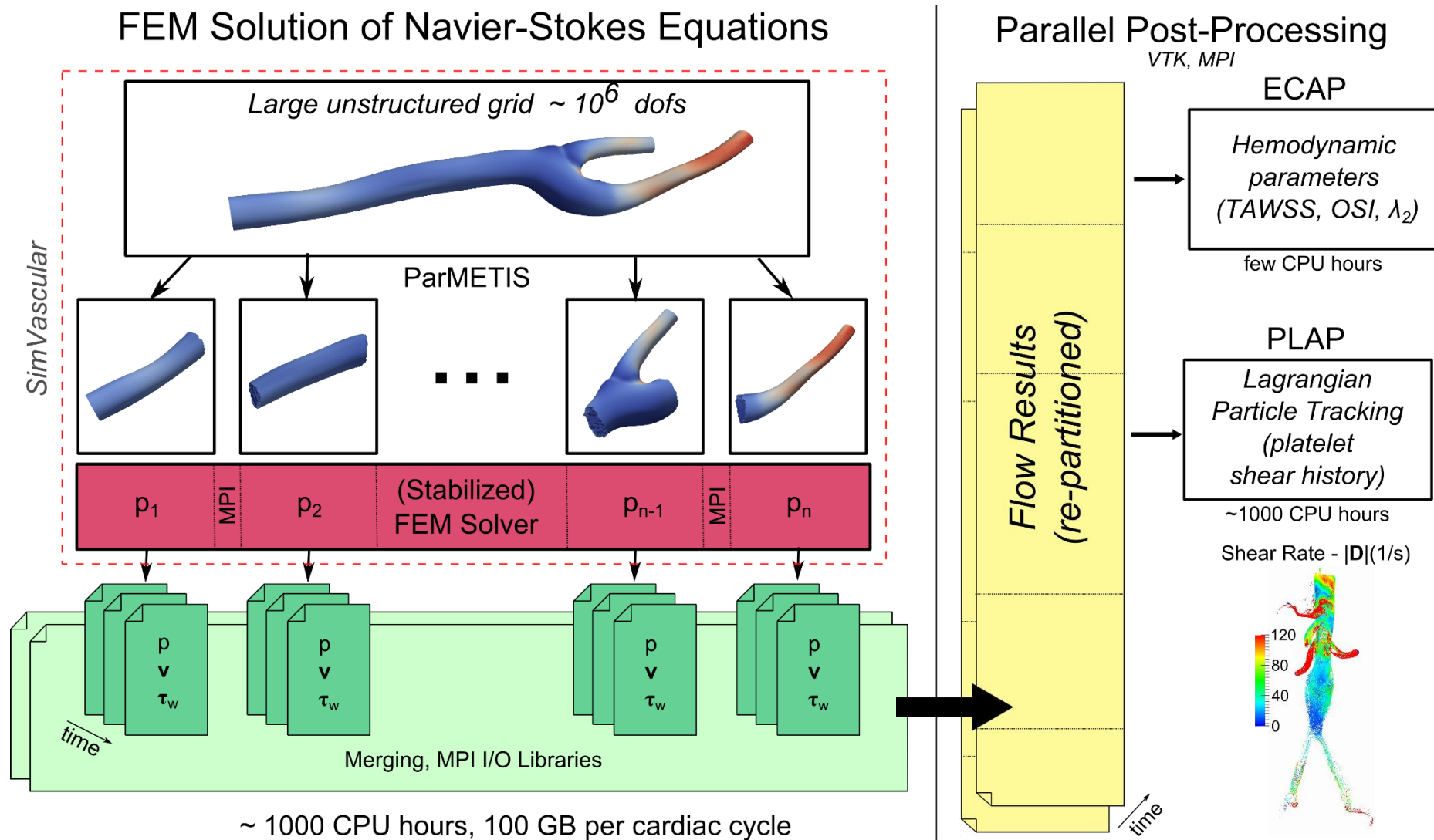
**Platelet Activation Potential**

*Regions of Endothelium Exposed to Activate Platelets*

$$\text{PLAP} = \int_{t-2\tau}^t |\mathbf{D}(\mathbf{x}, t)| \, dt$$

*Shadden and Hendabadi 2013 - BMMB*

# Large-scale Computational (Hemodynamic) Simulations



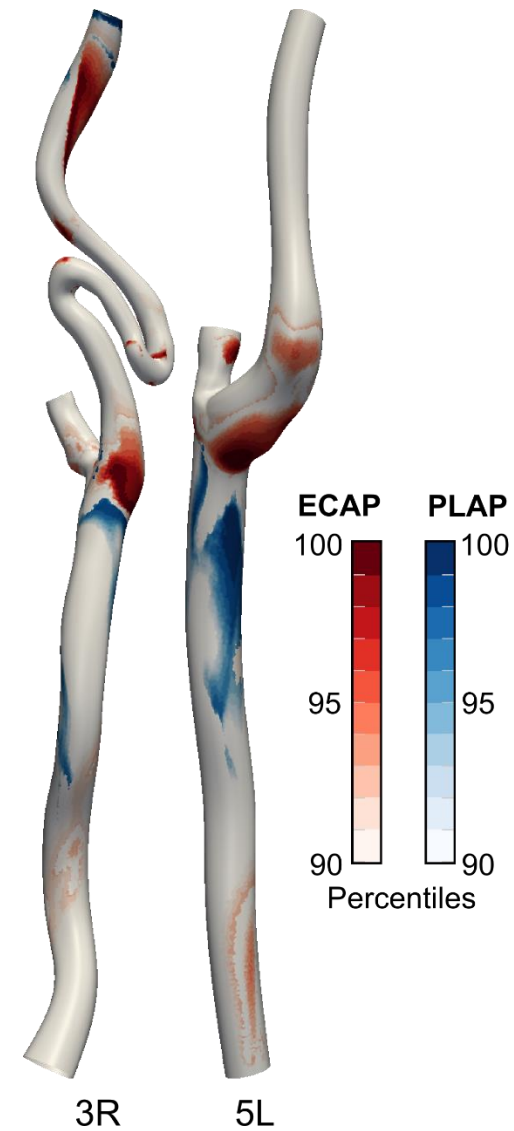
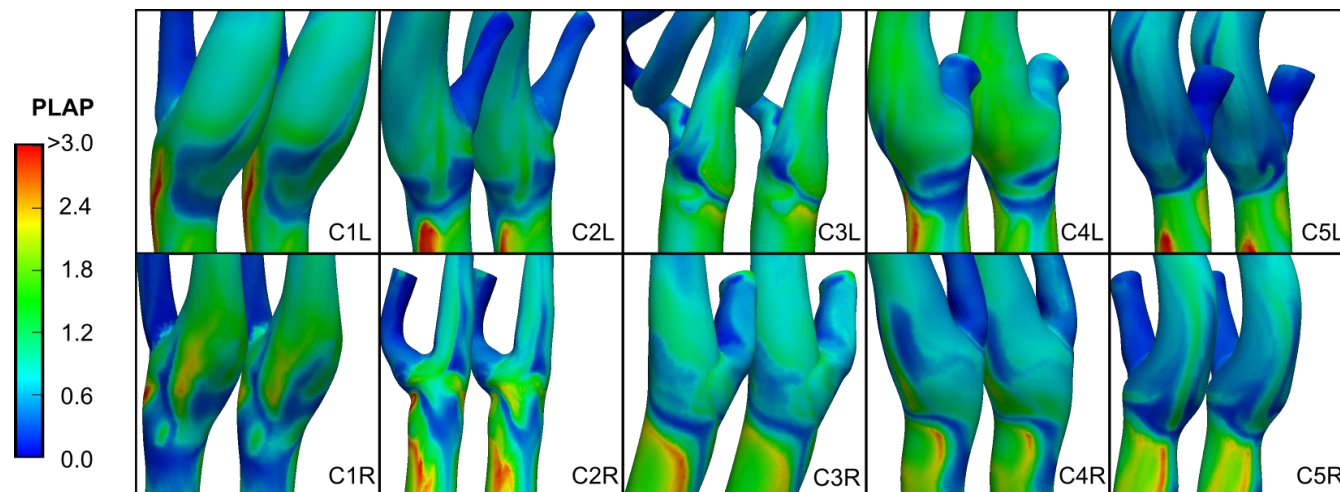
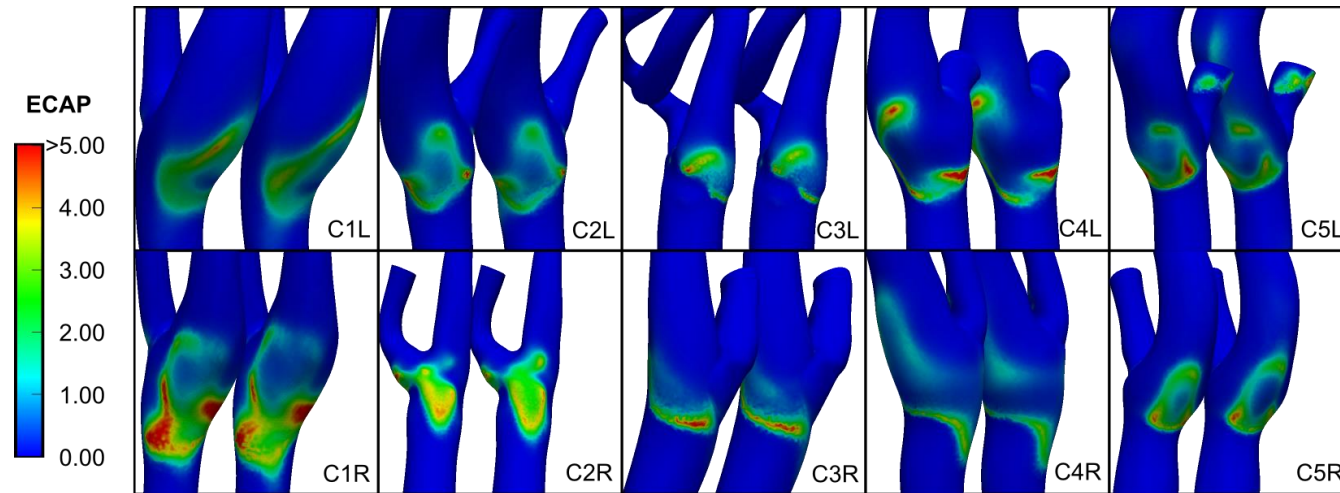
# Typical Computational Workload

## Typical Blood Flow Simulation (5 heart beats)

- $5 \cdot 10^6$  degrees of freedom (patient-specific model)
- $10^4$  time steps (unsteady Navier-Stokes)
- 256 cores (max limit for linear scaling of our solver)
- $5 \cdot 10^3$  CPU hours =  $4 \cdot 10^3$  CPU hours for solution +  $10^3$  for parallel post-processing (particle tracking, advection diffusion)
- 50 GB storage/simulation *after* reduction (elimination of transients, efficient storage in HDF5 files)

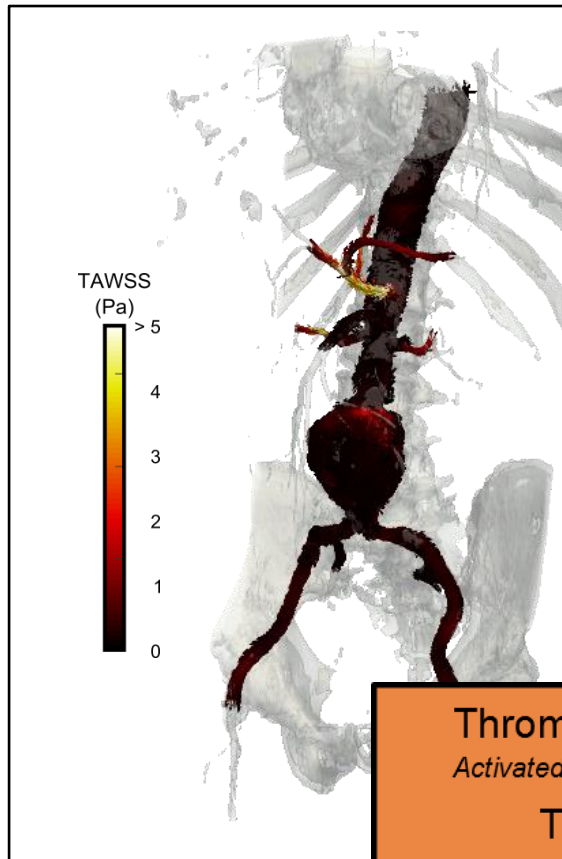


# ECAP & PLAP do not Co-localize in Carotids

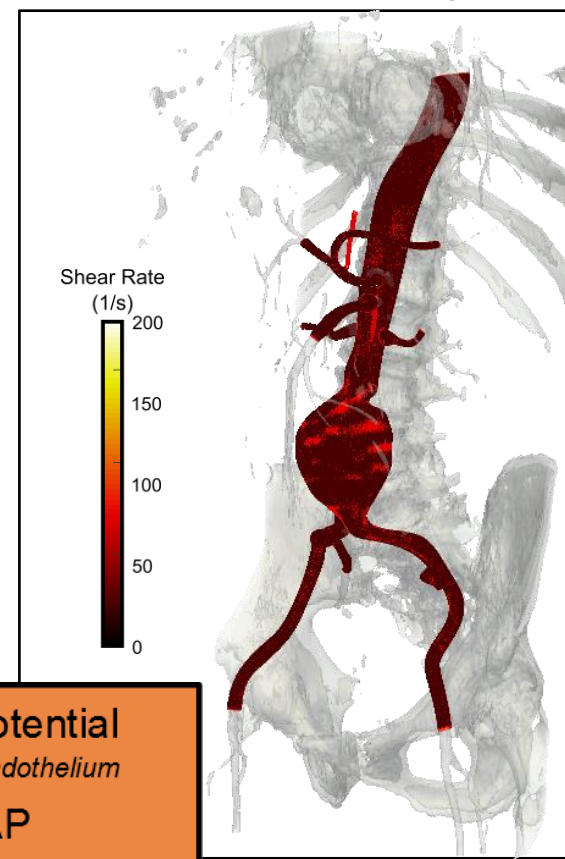


# Thrombus Formation Potential for AAAs

Computational Fluid Dynamics



Particle Tracking



**Thrombus Formation Potential**  
*Activated Platelets on Receptive Endothelium*

$$\text{TFP} = \text{ECAP} \cdot \text{PLAP}$$

**Endothelial Cell Activation Potential**

*Low and Oscillatory Shear Stress Regions*

$$\text{ECAP} = \frac{\text{OSI}}{\text{TAWSS}}$$

**Platelet Activation Potential**

*Regions of Endothelium Exposed to Activate Platelets*

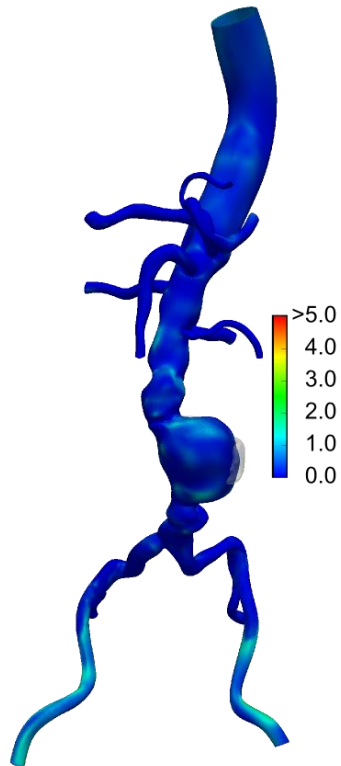
$$\text{PLAP} = \int_{t_1}^{t_2} |\mathbf{D}(\mathbf{x}, t)| \, dt$$

*Shadden and Hendabadi 2013 - BMMB*

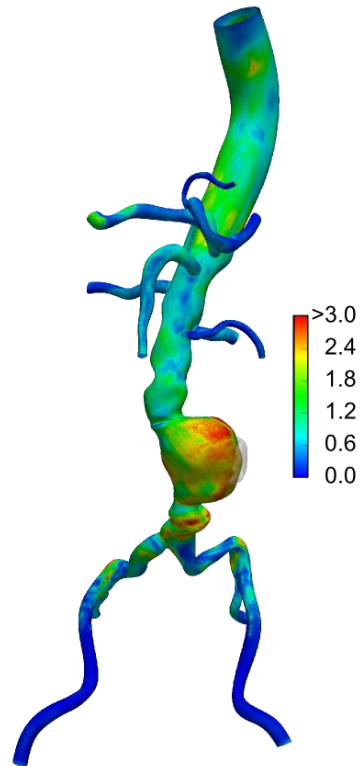


# ECAP & PLAP Co-localize in AAAs – Promote Thrombus

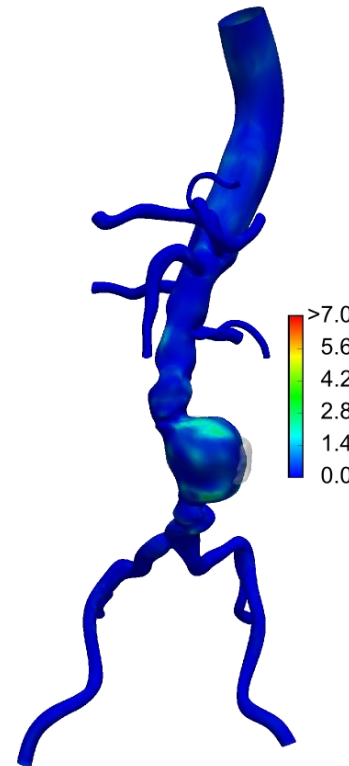
ECAP



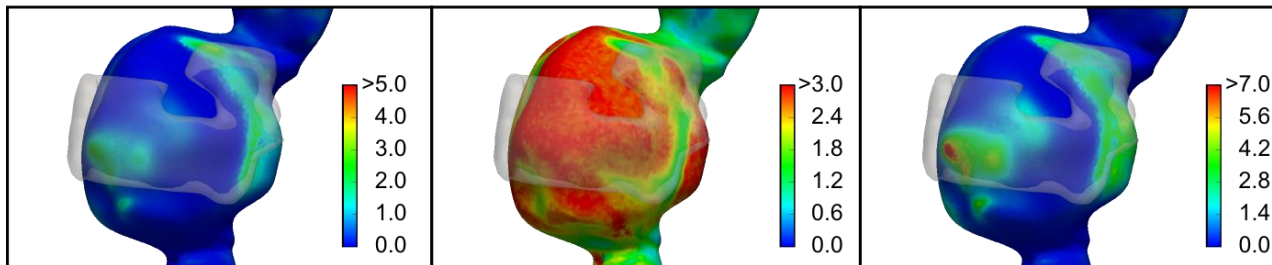
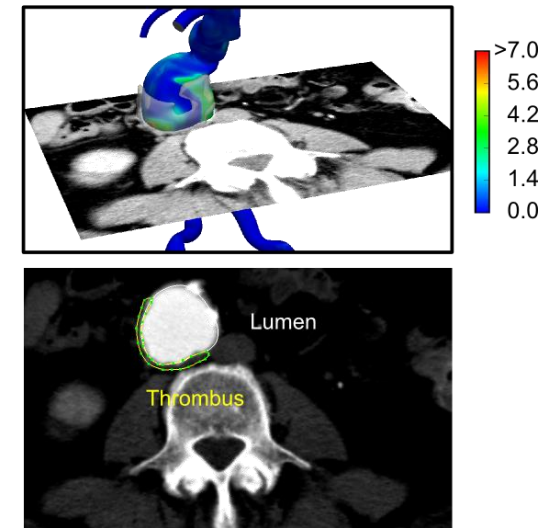
PLAP



TFP



TFP/Thrombus Colocalization



# Hemodynamic Metrics of Thrombus Formation

## Thrombus Formation Potential *initiation*

**Endothelial cells**

$$TFP(x) = \frac{OSI(x)}{TAWSS(x)} \cdot \overline{PLAP}(x),$$

**Platelets**

## Thrombus Deposition Potential *propagation*

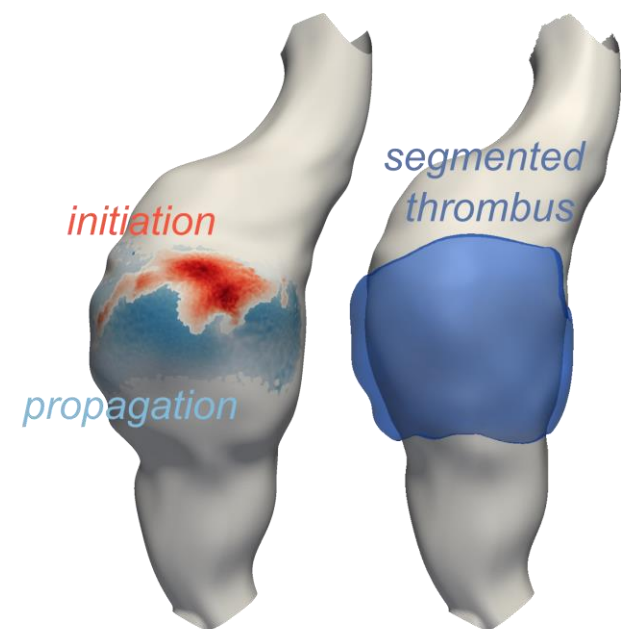
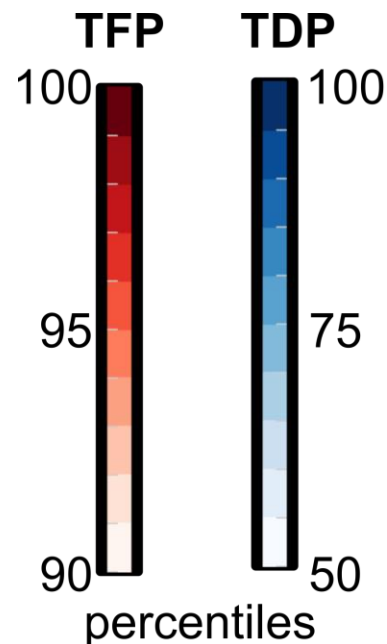
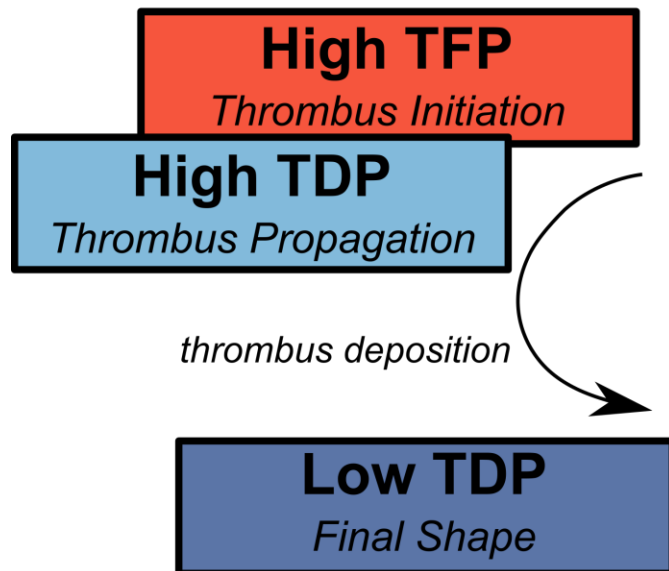
**Thrombogenic**

$$TDP(x) = 0.5 (1 + [FBG](x))$$

$$-k_1^2 \overline{OSI}(x) - k_2^2 \overline{TAWSS}(x)$$

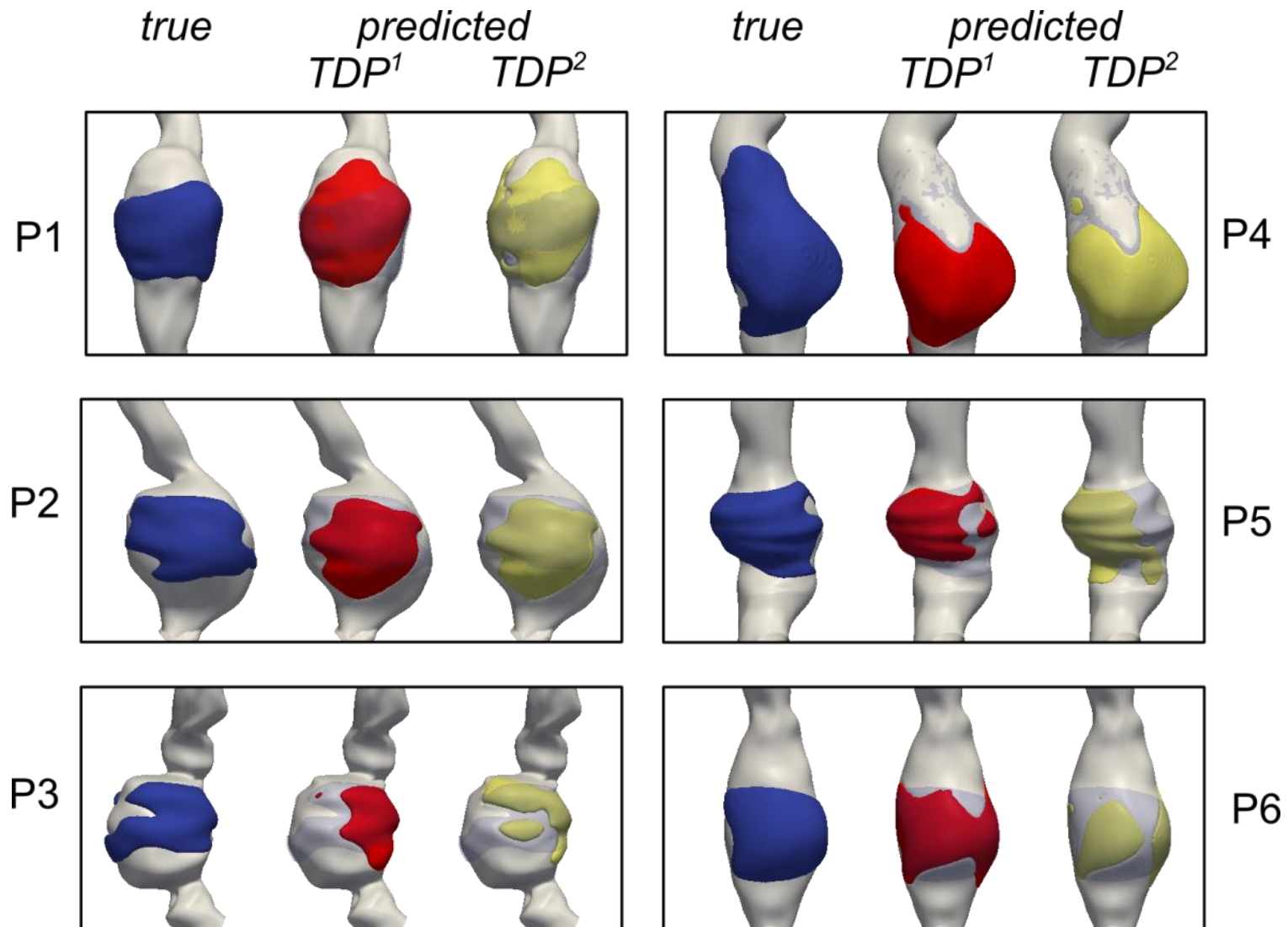
**Friction**

### Proposed Mechanism





# Predicted Thrombus Deposition



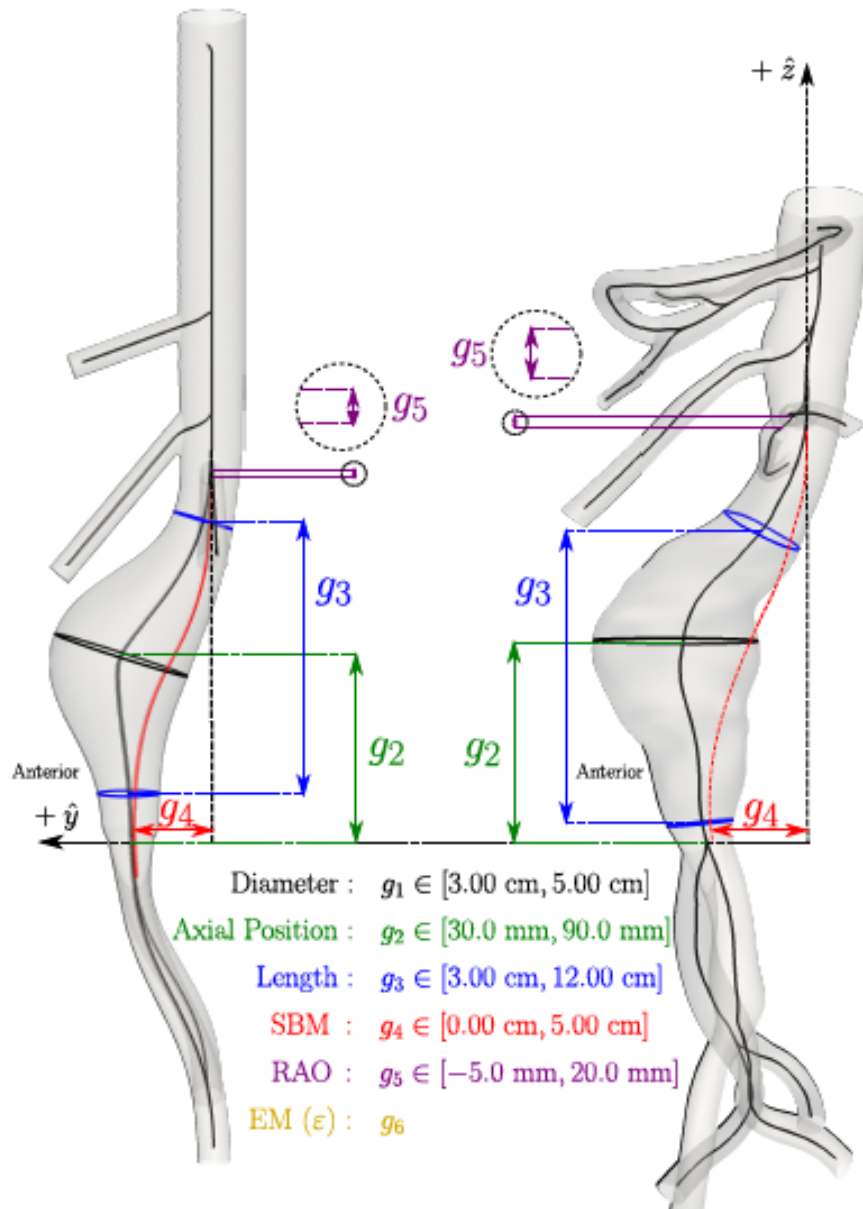
# HPC Resources Utilized



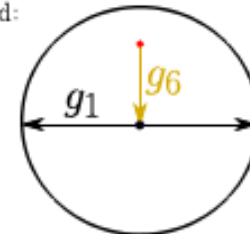
- Yale Center for Research Computing
  - Omega HPC Cluster (HP ProLiant blades) –  $3 \cdot 10^6$  CPU/hrs
  - Grace HPC Cluster (IBM/Lenovo NeXtScale) –  $0.5 \cdot 10^6$  CPU/hrs
- XSEDE Consortium
  - TACC LoneStar (Dell/Cray) –  $0.35 \cdot 10^6$  CPU/hrs
  - TACC Stampede (Dell/Mellanox) –  $0.3 \cdot 10^6$  CPU/hrs
  - SDSC Comet (Dell) –  $0.1 \cdot 10^6$  CPU/hrs
  - LSU SuperMic (Dell) –  $0.1 \cdot 10^6$  CPU/hrs







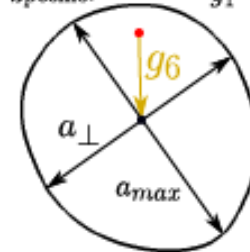
Idealized:



Anterior

Patient-Specific:

$$g_1 = \frac{a_{max} + a_{\perp}}{2}$$



Anterior

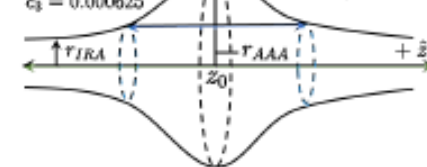
## Lesion Geometry

$$r(z) = e^{-\gamma(\theta)^{2.48}|z-z_0|^{1.75}} \left[ r_{AAA} - r_{IRA} - c_3 \left( \frac{z^2}{r_{IRA}} \right) \right] + r_{IRA}$$

$$c_1 = 0.2$$

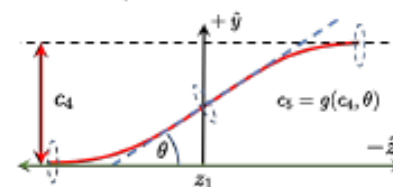
$$c_3 = 0.000625$$

$$c_2 = f(l_{AAA}, r_{AAA})$$



## Spinal Bend Geometry

$$CL_y(z) = c_4 \text{erf}(c_5(z - z_1))$$



# Acknowledgments

## Colleagues

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## Current & Former Students / Fellows

S. Baek, C. Cyron, **P. Di Achille**, S. Kyriacou, **S. Sankaran**, P. Seshaiyer, A. Valentin, L. Virag, J. Wilson

## Key References:

Wilson JS et al. (2013) *J Biomech Engr*;  
Di Achille P et al. (2014) *Pro R Soc A*;  
Virag L et al. (2015) *Annl Biomed Engr*;  
Di Achille P et al. (2016) *IJNMBBE*

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# Additional Resources

- Visualization Hardware
  - UT Maverick (HP/Nvidia)
  - Visualization Nodes on Stampede
  - High-End Workstations in Continuum Biomechanics Lab
- Data Transfer/Storage
  - Globus, GridFTP
  - TACC Ranch (500 GB)
  - Local Storage in Continuum Biomechanics Lab (~20 TB)

