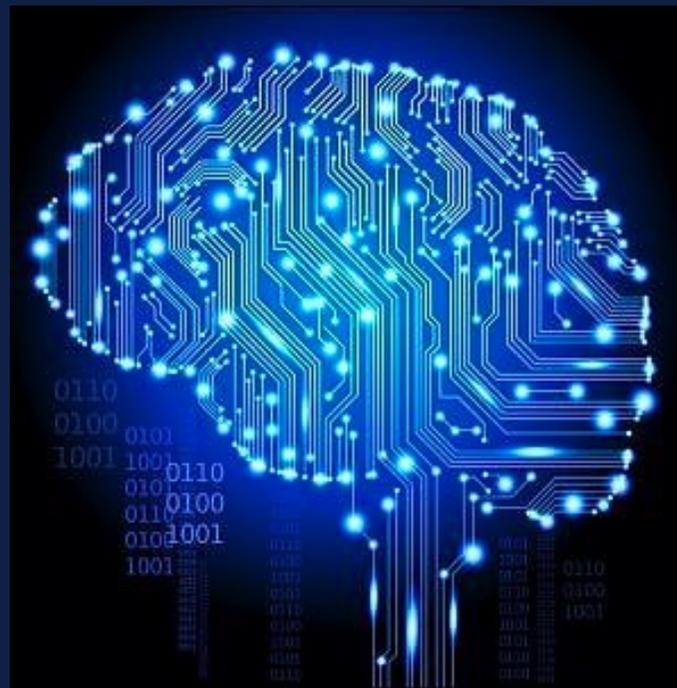


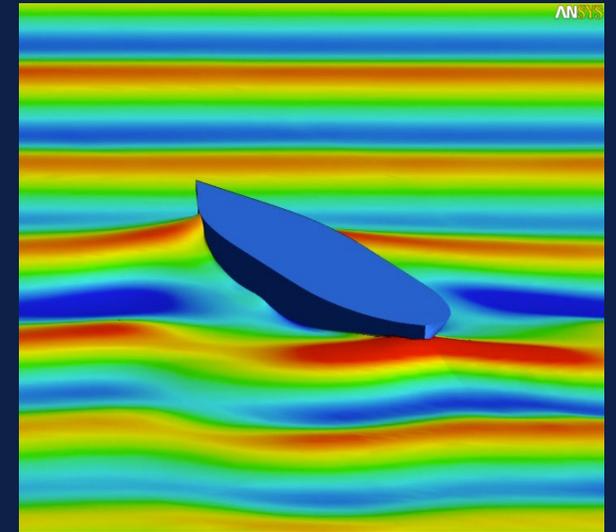
Intelligenza Artificiale (IA)

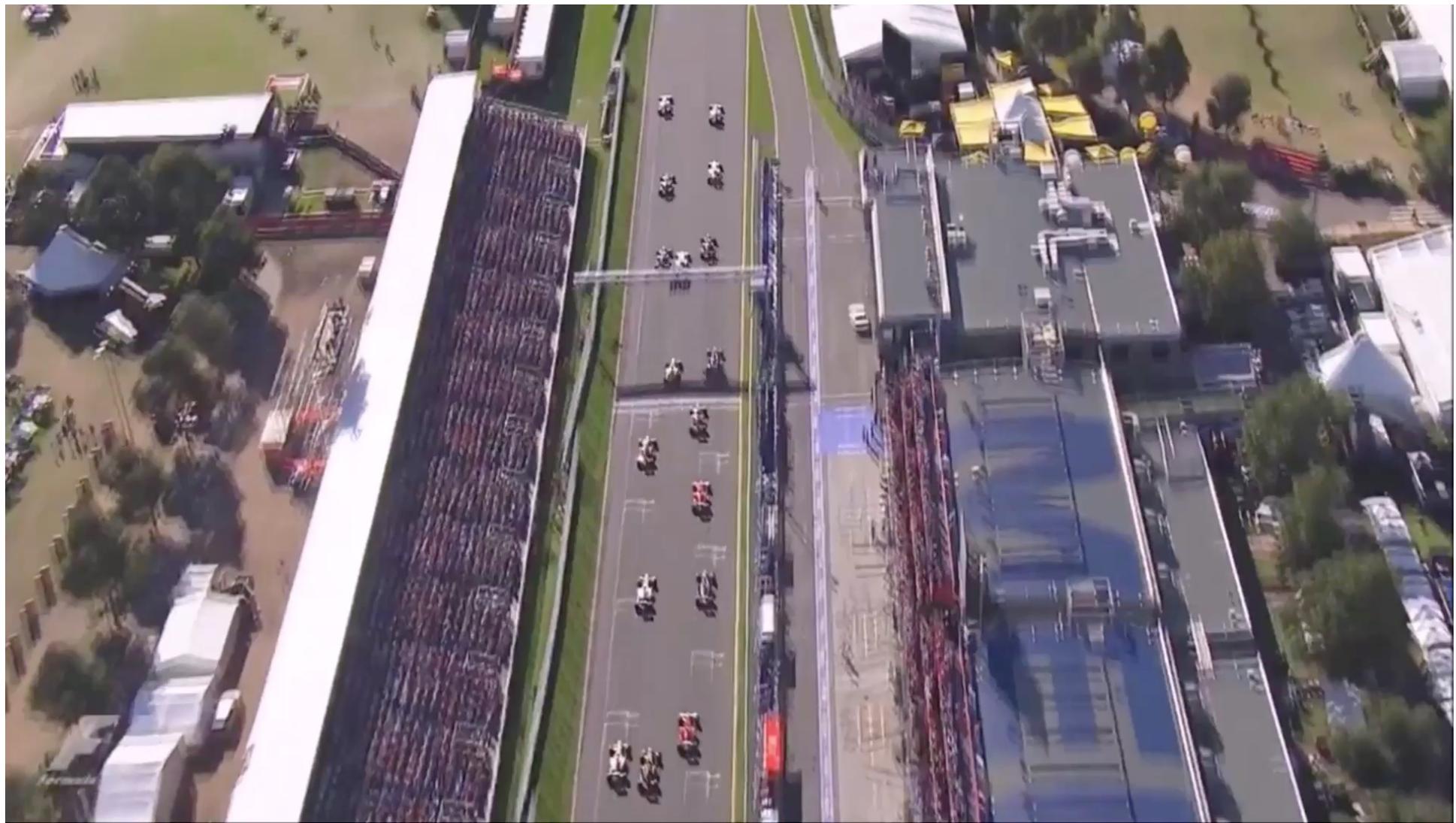
Parte 4 : Modelli Matematici (Physics Based)

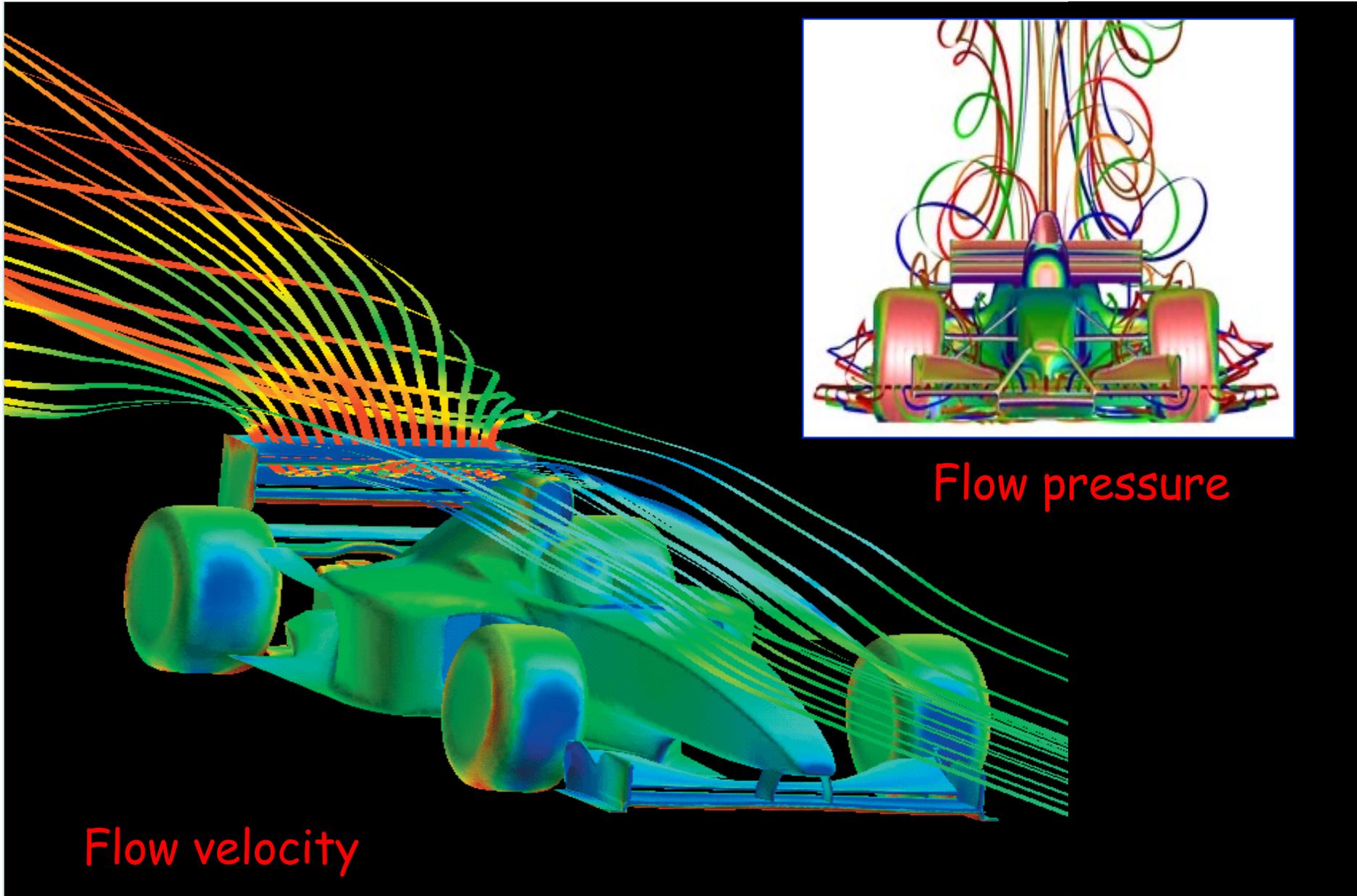




MATH for SPORTS







A Brief History of the America's Cup

☒

- Held by the U.S.A. for 132 years since 1851, the cup was then won sporadically by few other Countries (New Zealand and Australia)
- Switzerland competing for the first time in 2003 with EPFL engaged as scientific consultant



- **The Swiss Team Alinghi won the 2003 and 2007 consecutive editions**
- Switzerland still remains the only European country to have won the America's Cup



Our Main Achievements Thanks to Math Modelling





Math Model for America's Cup

Air
Phase

$$\frac{\partial(\rho_a \mathbf{u}_a)}{\partial t} + \nabla \cdot (\rho_a \mathbf{u}_a \otimes \mathbf{u}_a) - \nabla \cdot \mathbf{T}_a(\mathbf{u}_a, p_a) = \rho_a \mathbf{g}$$

$$\nabla \cdot \mathbf{u}_a = 0$$

Interface
Conditions

$$\mathbf{u}_a = \mathbf{u}_w \quad \text{on } \Gamma$$

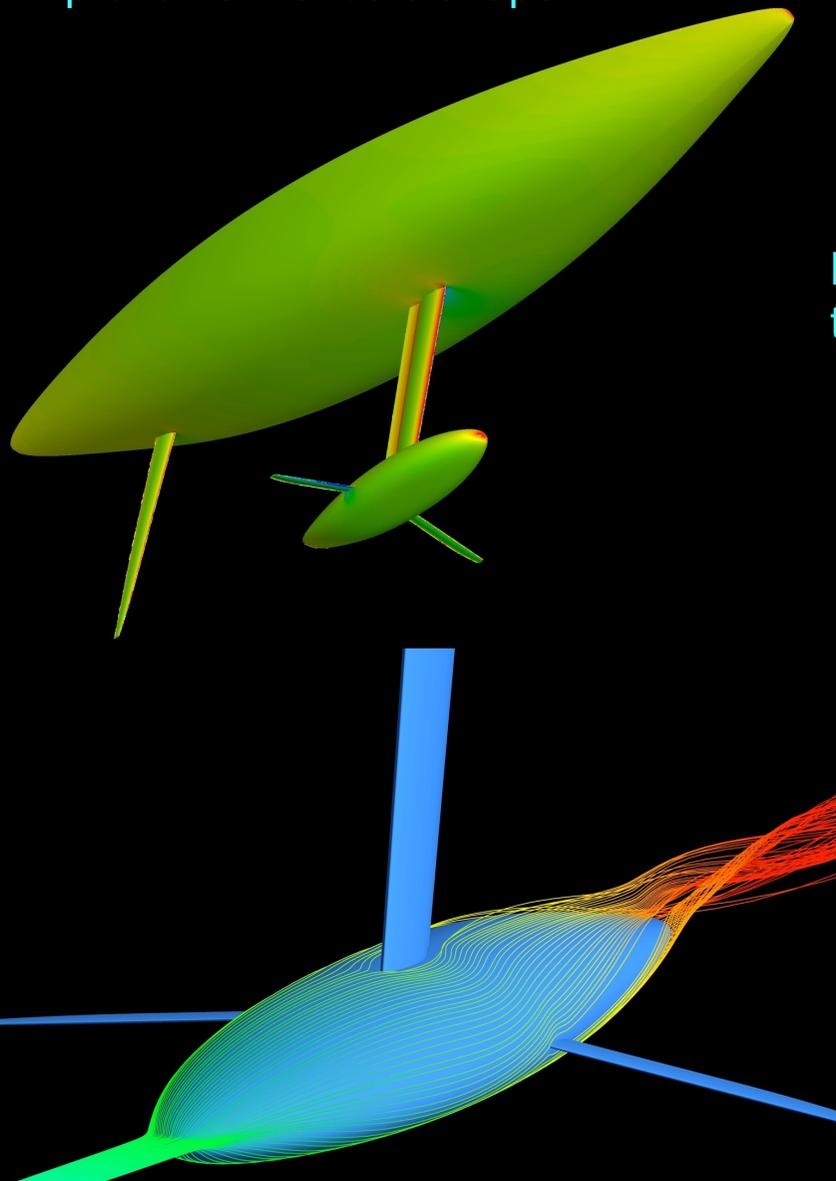
$$\mathbf{T}_a(\mathbf{u}_a, p_a) \cdot \mathbf{n} = \mathbf{T}_w(\mathbf{u}_w, p_w) \cdot \mathbf{n} + \kappa \sigma \mathbf{n} \quad \text{on } \Gamma$$

$$\frac{\partial(\rho_w \mathbf{u}_w)}{\partial t} + \nabla \cdot (\rho_w \mathbf{u}_w \otimes \mathbf{u}_w) - \nabla \cdot \mathbf{T}_w(\mathbf{u}_w, p_w) = \rho_w \mathbf{g}$$

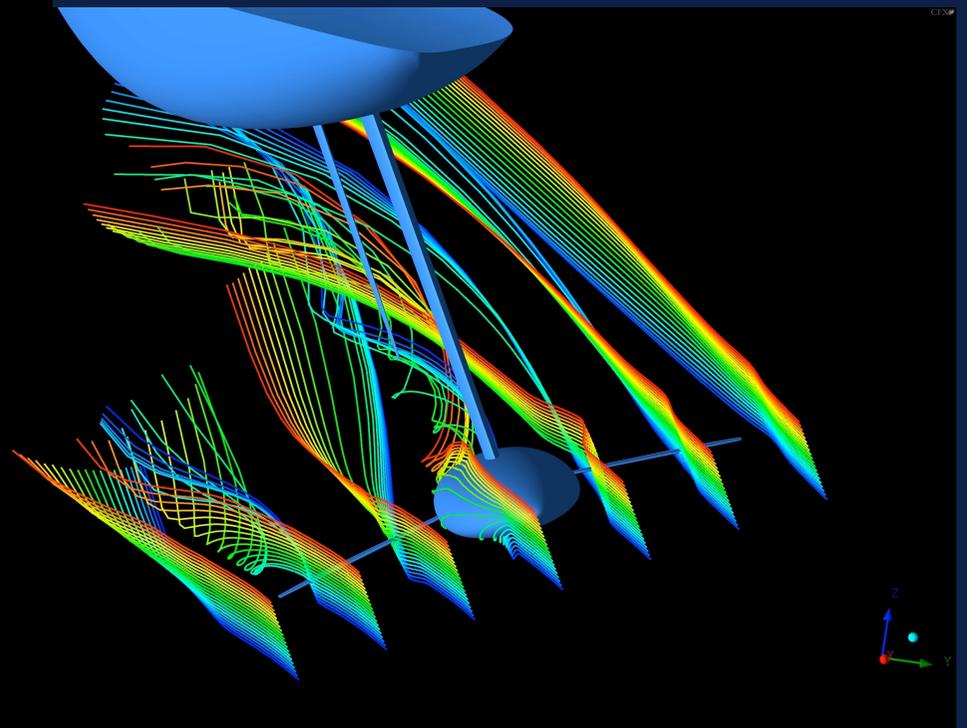
$$\nabla \cdot \mathbf{u}_w = 0$$

Water
Phase

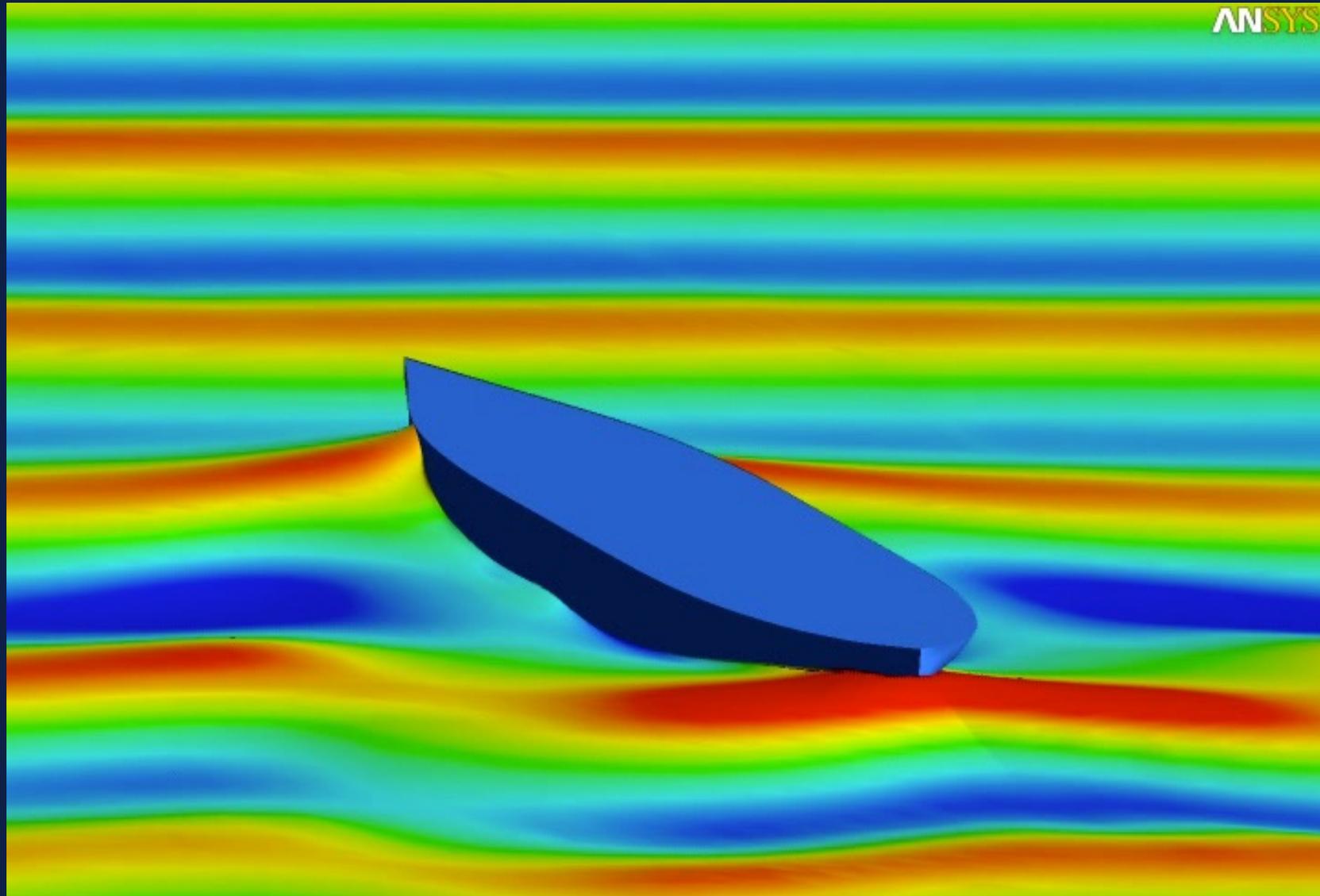
Improvement of bulb shape



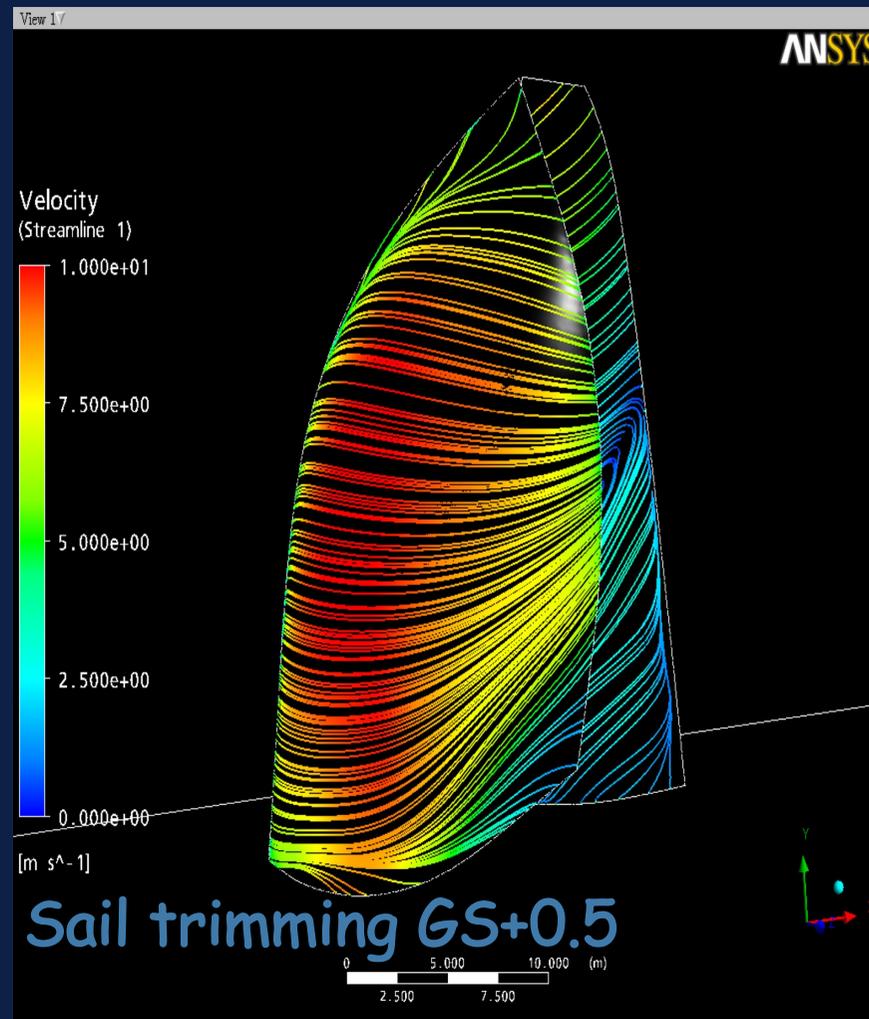
Better dislocation of winglets on the bulb



Minimisation of the wet surface on hydrostatic regime

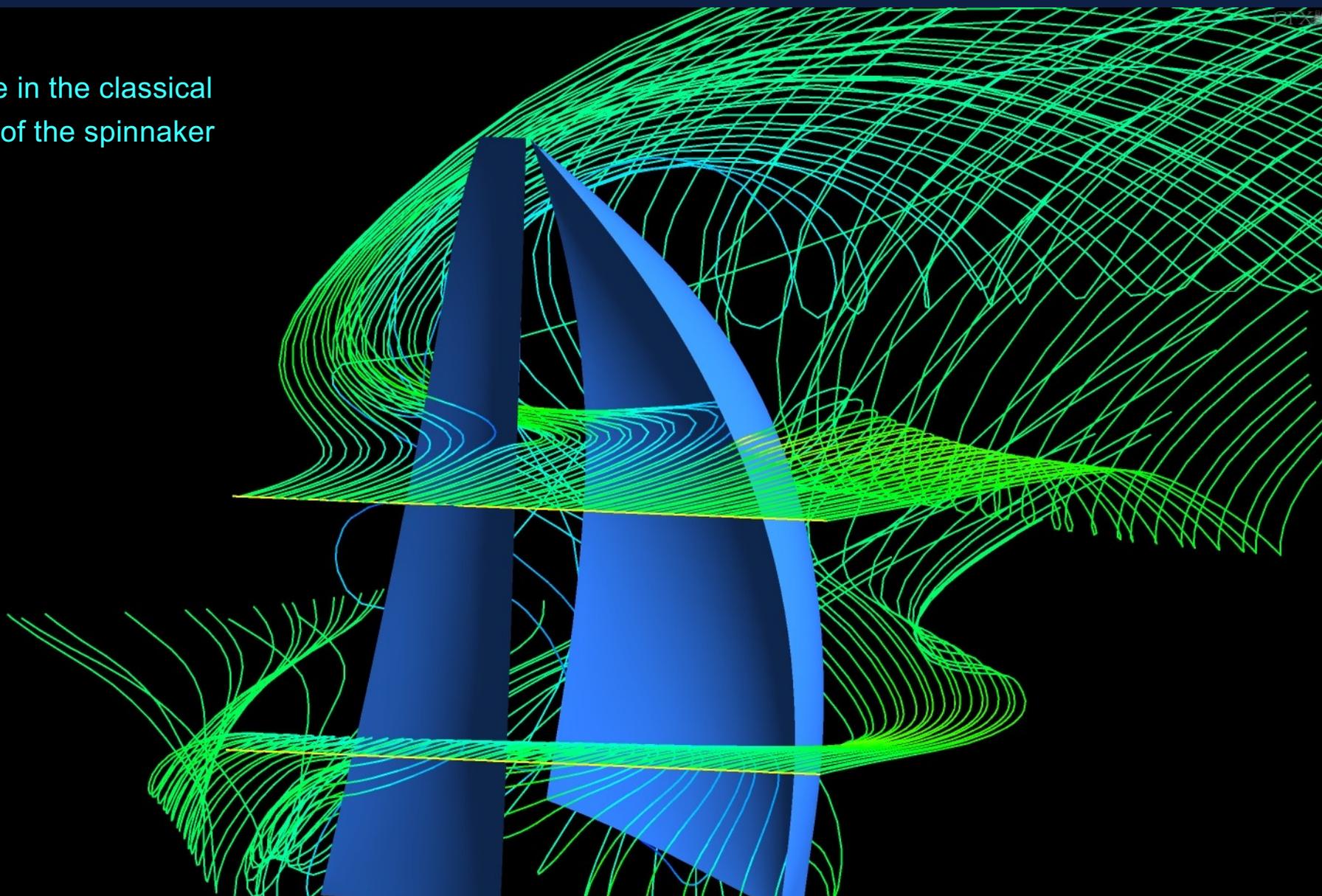


Sail trimming to maximize flow reattachment

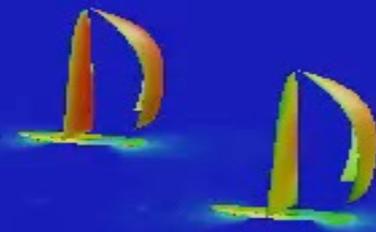


Sail trimming GS-1

Change in the classical
design of the spinnaker



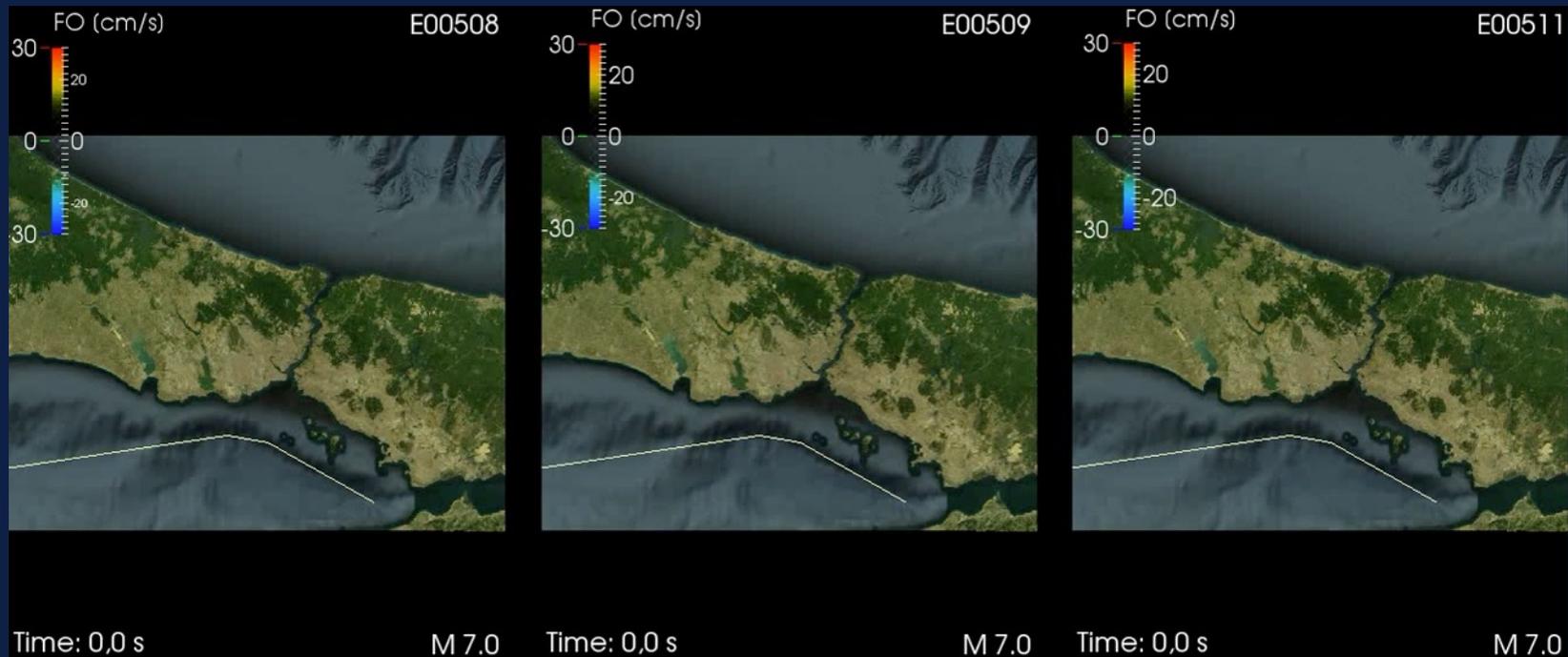
Maximise the shadow region in
downstream leg

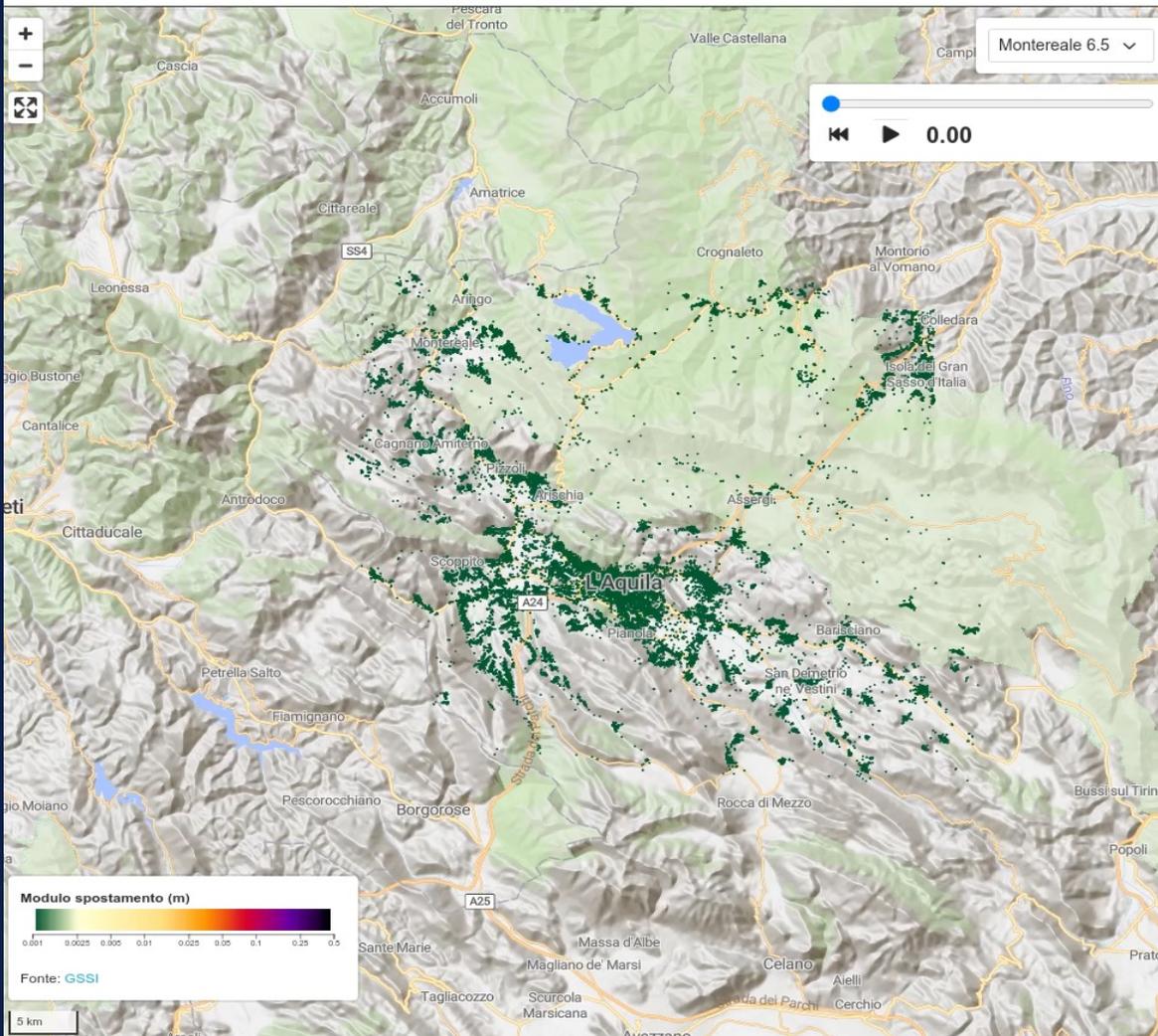




Alinghi

MATH for EARTHQUAKES







The Beijing metropolitan area

3D physics-based numerical modelling of Beijing



Beijing metropolitan area

Land: 16.801 km²

Population: 21.707.000

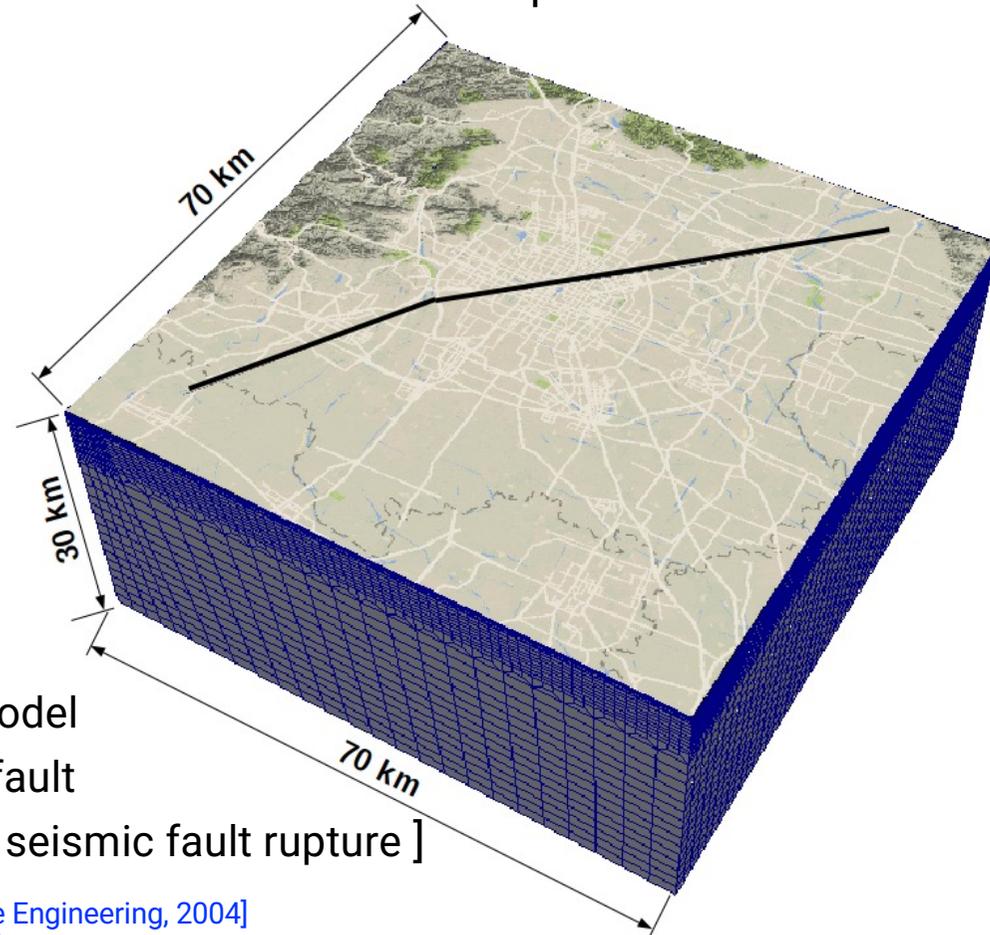
Density: 1.300/km²

- Topography and 3D basin model
- Shunyi-Qianmen-Liangxiang fault
- Kinematic description of the seismic fault rupture]

[Gao, Yu, Zhang, Wu, Conference on Earthquake Engineering, 2004]

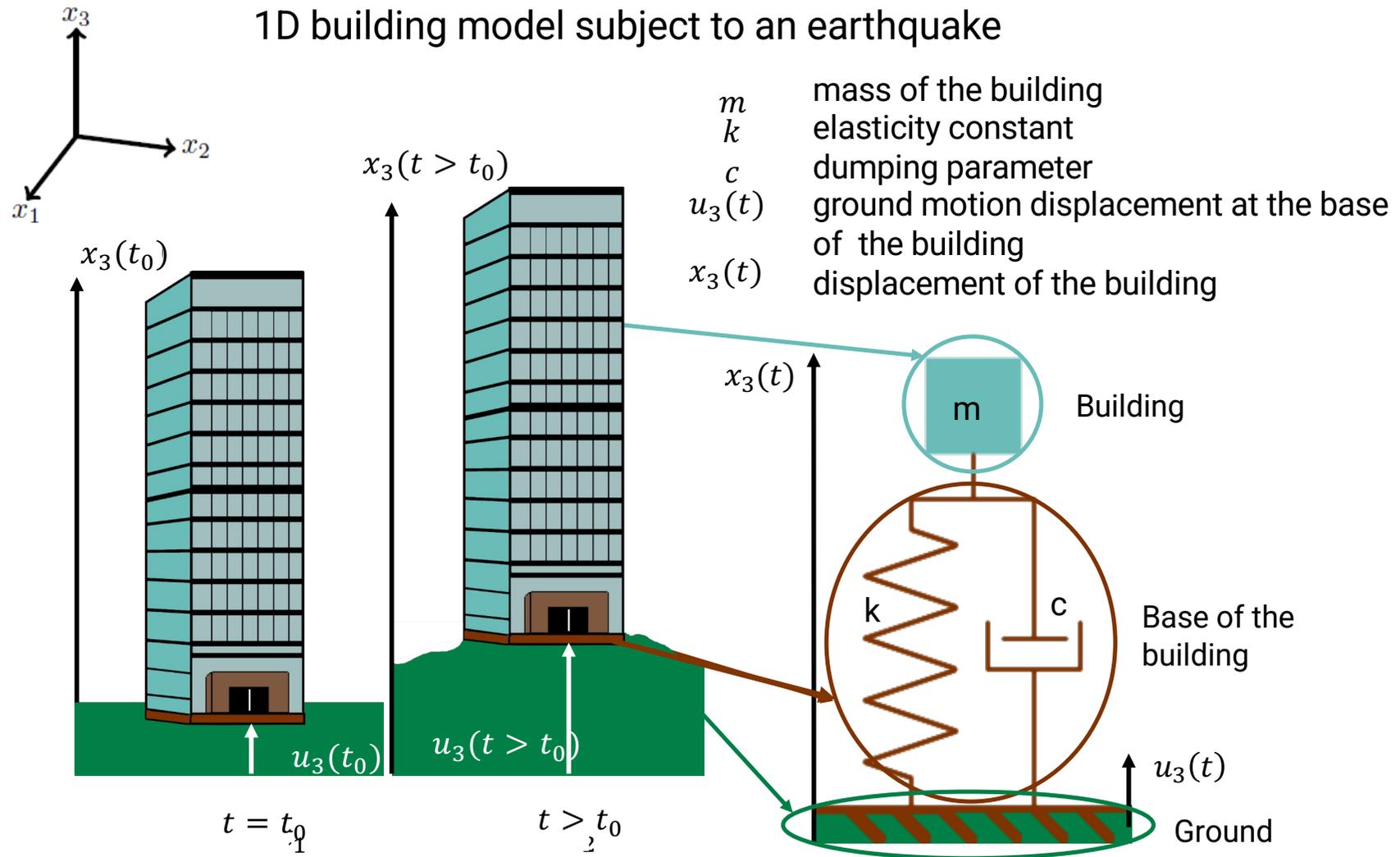
[Crempien, Archuleta, Seismol. Res. Lett., 2015]

Main features of the computational model

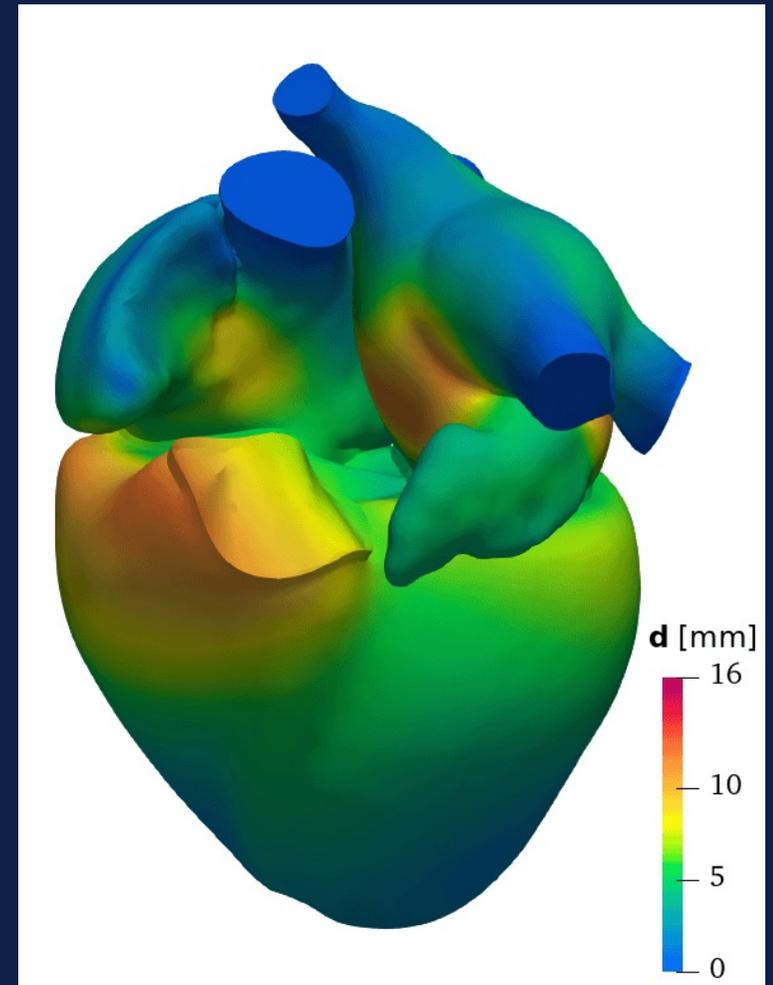


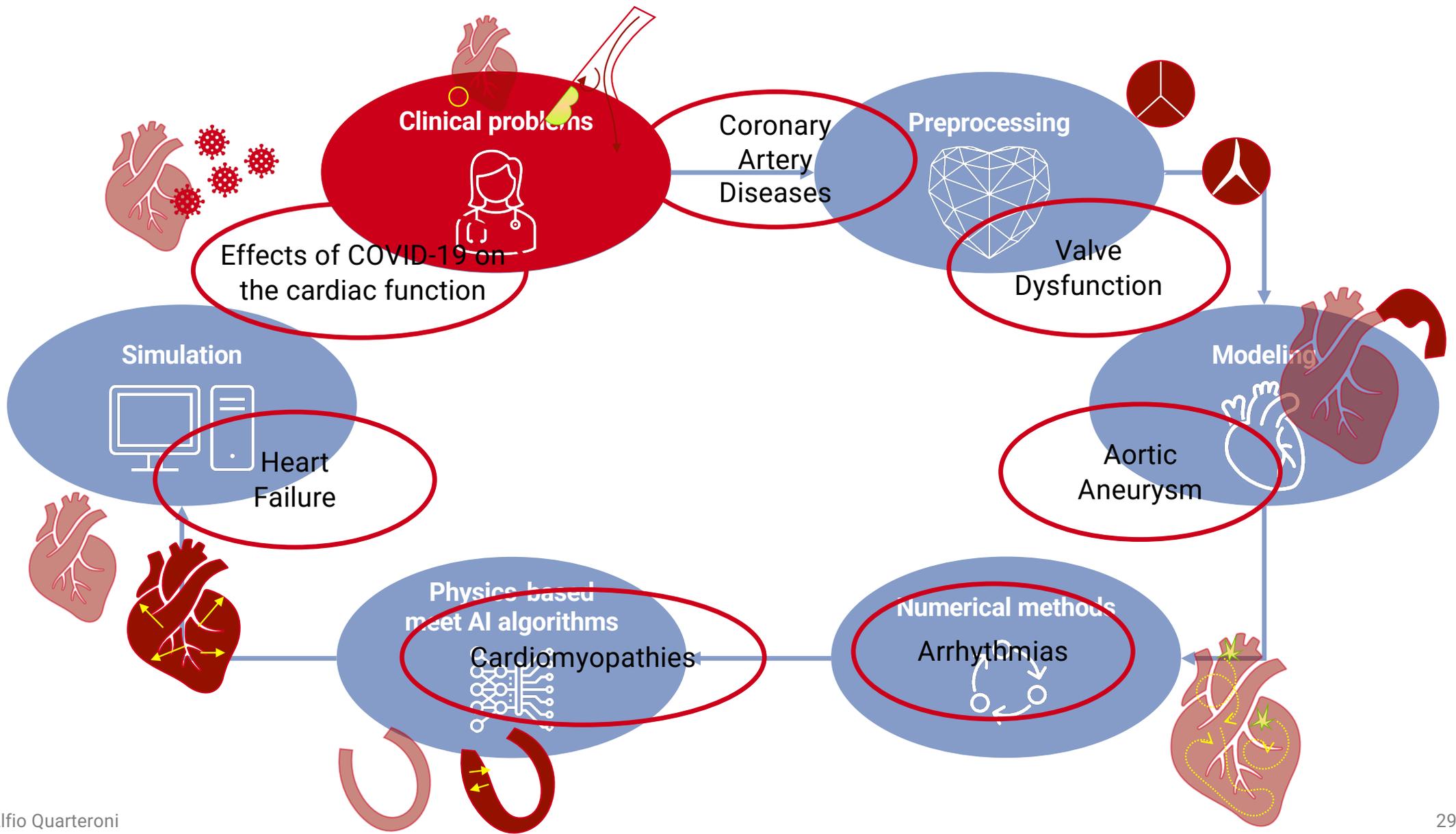
Ground motion parameters: spectral displacement

1D building model subject to an earthquake



MATH for HEART





The Mathematical Heart

$$\begin{cases} \frac{\partial \mathbf{w}}{\partial t} = \mathbf{F}_{\text{ion}}^{\mathbf{w}}(v, \mathbf{w}) & \text{in } \Omega \\ \frac{\partial \mathbf{z}}{\partial t} = \mathbf{F}_{\text{ion}}^{\mathbf{z}}(v, \mathbf{w}, \mathbf{z}) & \text{in } \Omega \end{cases}$$

Cellular Ions' dynamics

$$\begin{cases} J\chi C_m \frac{\partial v}{\partial t} - \nabla \cdot (JF^{-1} D_i F^{-T} \nabla (v + v_e)) \\ + J\chi I_{\text{ion}}(v, \mathbf{w}, \mathbf{z}) = J\chi I_{\text{app}}(\mathbf{x}, t) & \text{in } \Omega \\ -\nabla \cdot (JF^{-1} D_i F^{-T} \nabla v) - \nabla \cdot (JF^{-1} (D_i + D_e) \nabla v_e) = 0 & \text{in } \Omega \end{cases}$$

Cardiomyocytes Contraction

Electrophysiology

$$\mathbf{F}_{\text{circ}} \left(\mathbf{c}, \frac{d\mathbf{c}}{dt}, t \right) = \mathbf{0}$$

External Circulation

$$\begin{cases} \frac{\partial \mathbf{s}}{\partial t} = \mathbf{F}_{\text{act}} \left(\mathbf{s}, [\text{Ca}^{2+}]_i, \text{SL}, \frac{\partial \text{SL}}{\partial t} \right) & \text{in } \Omega \\ \text{SL} = \text{SL}_0 \sqrt{I_{4f}} & \text{in } \Omega \end{cases}$$

$$P_{\text{act}}(\mathbf{d}, \mathbf{s}) = T_{\text{act}} \left(n_f \frac{F\mathbf{f}_0 \cdot \mathbf{f}_0}{\sqrt{I_{4f}}} + n_s \frac{F\mathbf{s}_0 \cdot \mathbf{s}_0}{\sqrt{I_{4s}}} + n_n \frac{F\mathbf{n}_0 \cdot \mathbf{n}_0}{\sqrt{I_{4n}}} \right)$$

$$T_{\text{act}} = T_{\text{act}}(\mathbf{s}, \text{SL}), \quad I_{4i} = F\mathbf{i}_0 \cdot F\mathbf{i}_0 \quad i \in \{\mathbf{f}, \mathbf{s}, \mathbf{n}\}$$

$$\rho_f \left[\frac{\partial \mathbf{u}}{\partial t} + ((\mathbf{u} - \mathbf{u}_{\text{ALE}}) \cdot \nabla) \mathbf{u} \right] - \nabla \cdot \sigma_f(\mathbf{u}, p) + \mathcal{R}(\mathbf{u}, \mathbf{u}_{\text{ALE}}) = \mathbf{0}$$

$$\mathcal{R}(\mathbf{u}, \mathbf{u}_{\text{ALE}}) = \sum_{k \in \mathcal{V}} \frac{R_k}{\varepsilon_k} \delta_{\varepsilon_k} (\varphi_k^t(\mathbf{x})) (\mathbf{u} - \mathbf{u}_{\text{ALE}} - \mathbf{u}_{\Gamma_k})$$

Valves Dynamics

$$\begin{cases} \text{NS}(\mathbf{u}, p) = \mathbf{0} & \text{in } \Omega_{\text{cor}} \\ \mathbf{K}_i^{-1} \mathbf{u}_{\text{myo},i} + \nabla p_{\text{myo},i} = \mathbf{0}, \quad i = 1, 2, 3 & \text{in } \Omega_{\text{myo}} \\ \nabla \cdot \mathbf{u}_{\text{myo},1} = \sum_{j=1}^J \frac{\chi_{\Omega_{\text{myo}}^j}}{|\Omega_{\text{myo}}^j|} \int_{\Gamma_j^{\text{coro}}} \mathbf{u} \cdot \mathbf{n} - \beta_{1,2}(p_{\text{myo},1} - p_{\text{myo},2}) & \text{in } \Omega_{\text{myo}} \\ \nabla \cdot \mathbf{u}_{\text{myo},2} = -\beta_{2,1}(p_{\text{myo},2} - p_{\text{myo},1}) - \beta_{2,3}(p_{\text{myo},2} - p_{\text{myo},3}) & \text{in } \Omega_{\text{myo}} \\ \nabla \cdot \mathbf{u}_{\text{myo},3} = -\gamma(p_{\text{myo},3} - p_{\text{veins}}) - \beta_{3,2}(p_{\text{myo},3} - p_{\text{myo},2}) & \text{in } \Omega_{\text{myo}} \end{cases}$$

Myocardial Perfusion

$$\begin{cases} -\nabla \cdot P_{\text{ALE}}(\mathbf{d}_{\text{ALE}}) = \mathbf{0} & \text{in } \hat{\Omega} \\ \mathbf{d}_{\text{ALE}} = \mathbf{d} & \text{on } \hat{\Sigma} \end{cases} \quad \mathbf{u}_{\text{ALE}} = \frac{\partial \mathbf{d}_{\text{ALE}}}{\partial t}$$

$$\begin{cases} \rho_f \left[\frac{\partial \mathbf{u}}{\partial t} + ((\mathbf{u} - \mathbf{u}_{\text{ALE}}) \cdot \nabla) \mathbf{u} \right] - \nabla \cdot \sigma_f(\mathbf{u}, p) = \mathbf{0} & \text{in } \Omega \\ \nabla \cdot \mathbf{u} = 0 & \text{in } \Omega \end{cases}$$

$$\sigma_f(\mathbf{u}, p) = \mu (\nabla \mathbf{u} + \nabla \mathbf{u}^T) - pI$$

Blood Dynamics, Contraction & Relaxation

$$\rho_s \frac{\partial^2 \mathbf{d}}{\partial t^2} - \nabla \cdot P_s(\mathbf{d}, \mathbf{s}) = \mathbf{0} \quad \text{in } \Omega$$

$$P_s(\mathbf{d}, \mathbf{s}) = P_{\text{pas}}(\mathbf{d}) + P_{\text{act}}(\mathbf{d}, \mathbf{s})$$

$$P_{\text{pas}}(\mathbf{d}) = \frac{\partial W}{\partial F} \quad F = I + \nabla \mathbf{d}$$

ECG Reconstruction

$$\begin{cases} \text{EP}(v, v_e) = 0 & \text{in } \Omega_H \\ -\nabla \cdot (D_T \nabla v_T) = 0 & \text{in } \Omega_{\text{torso}} \\ v_T = v_e & \text{on } \Sigma \\ D_T \nabla v_T \cdot \mathbf{n}_H = D_e \nabla v_e \cdot \mathbf{n}_H & \text{on } \Sigma \end{cases}$$

Whole Heart electrophysiology simulation

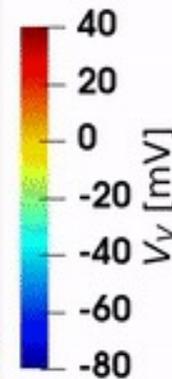
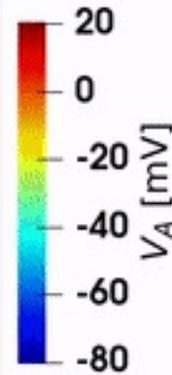
Time evolution of the ventricula (V_V) and atria (V_A) transmembrane potential



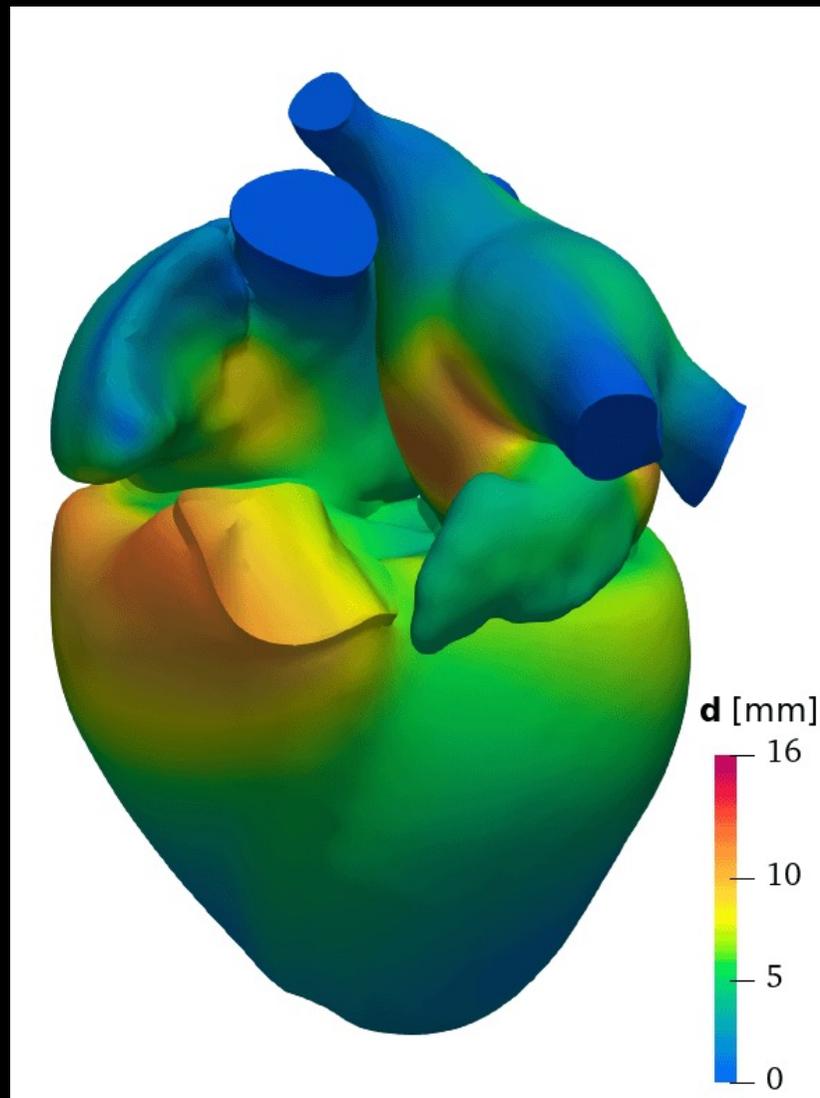
Time: 0.0016s



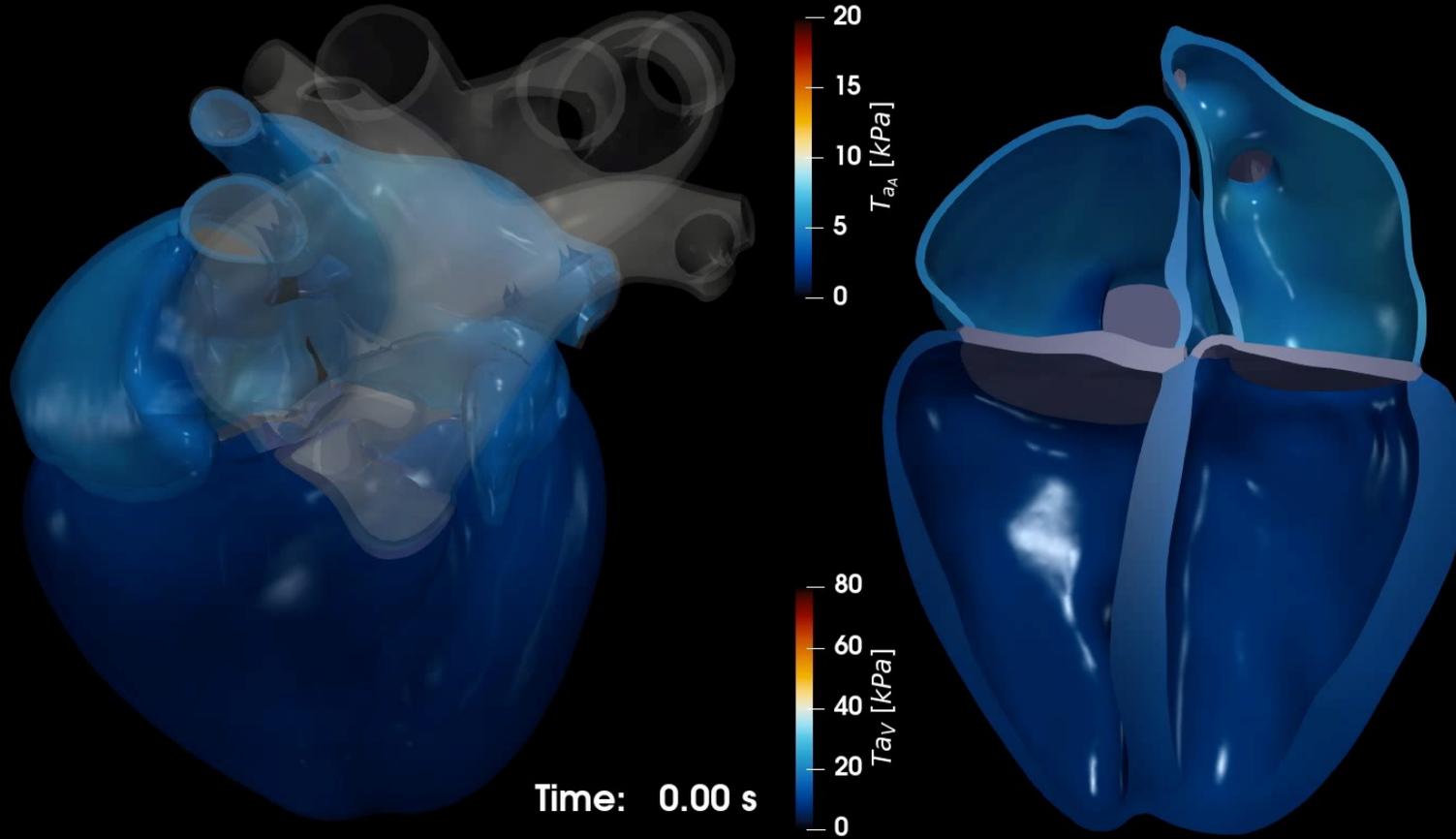
Electrical wave propagation in the heart



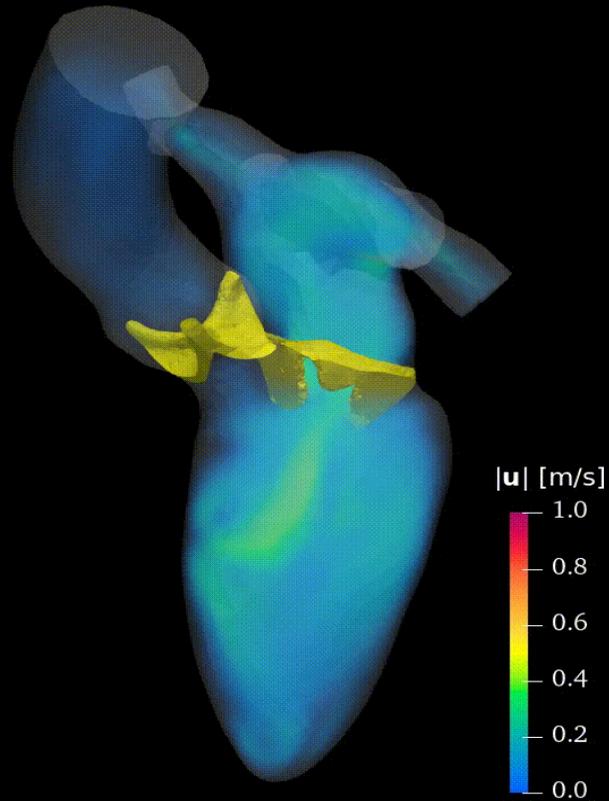
Displacement (Contraction and Relaxation)



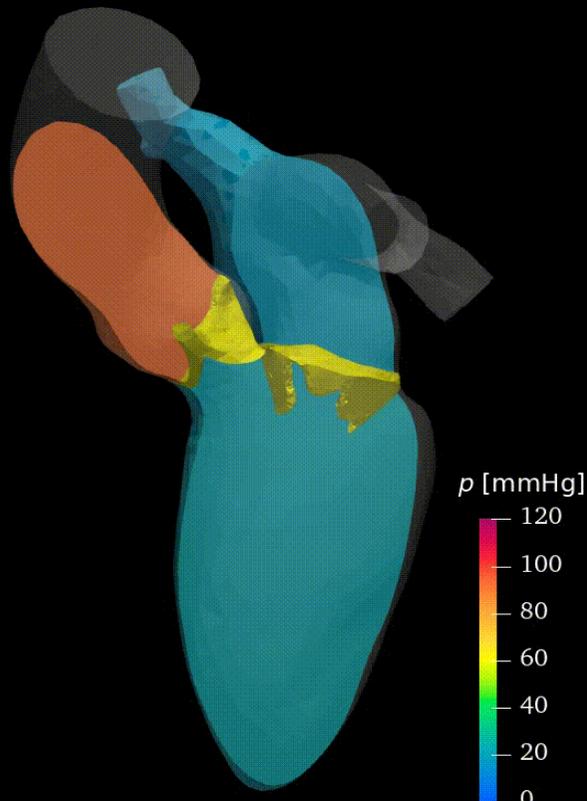
Active Tension at Cardiomyocytes



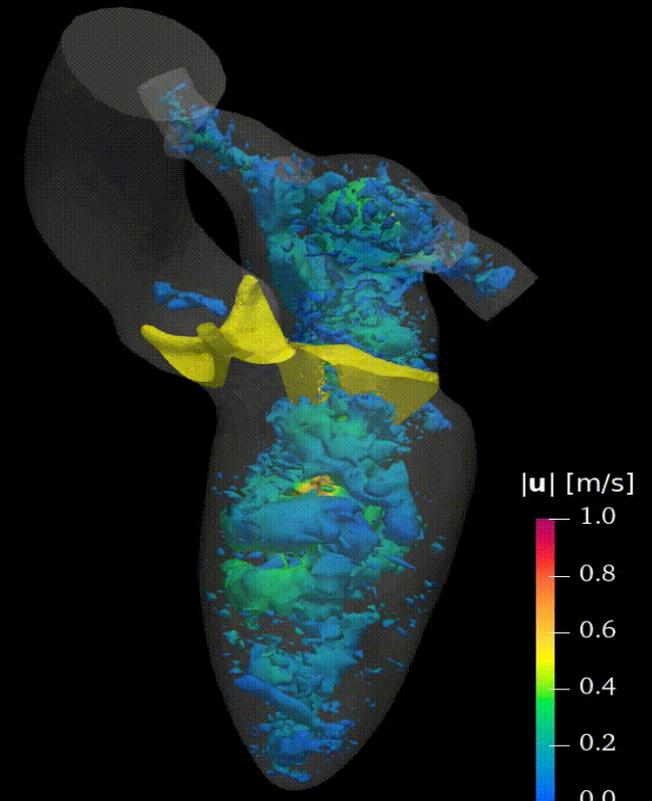
Blood velocity



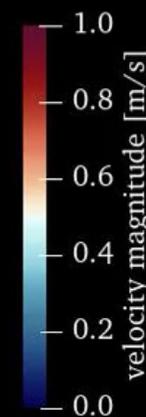
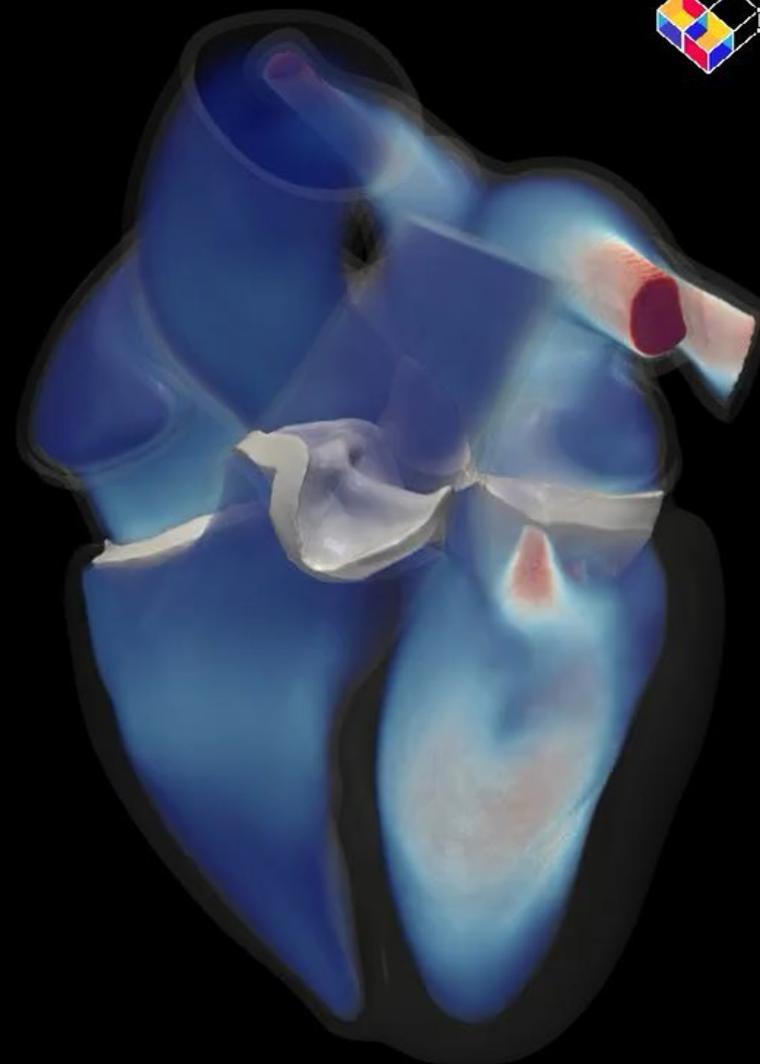
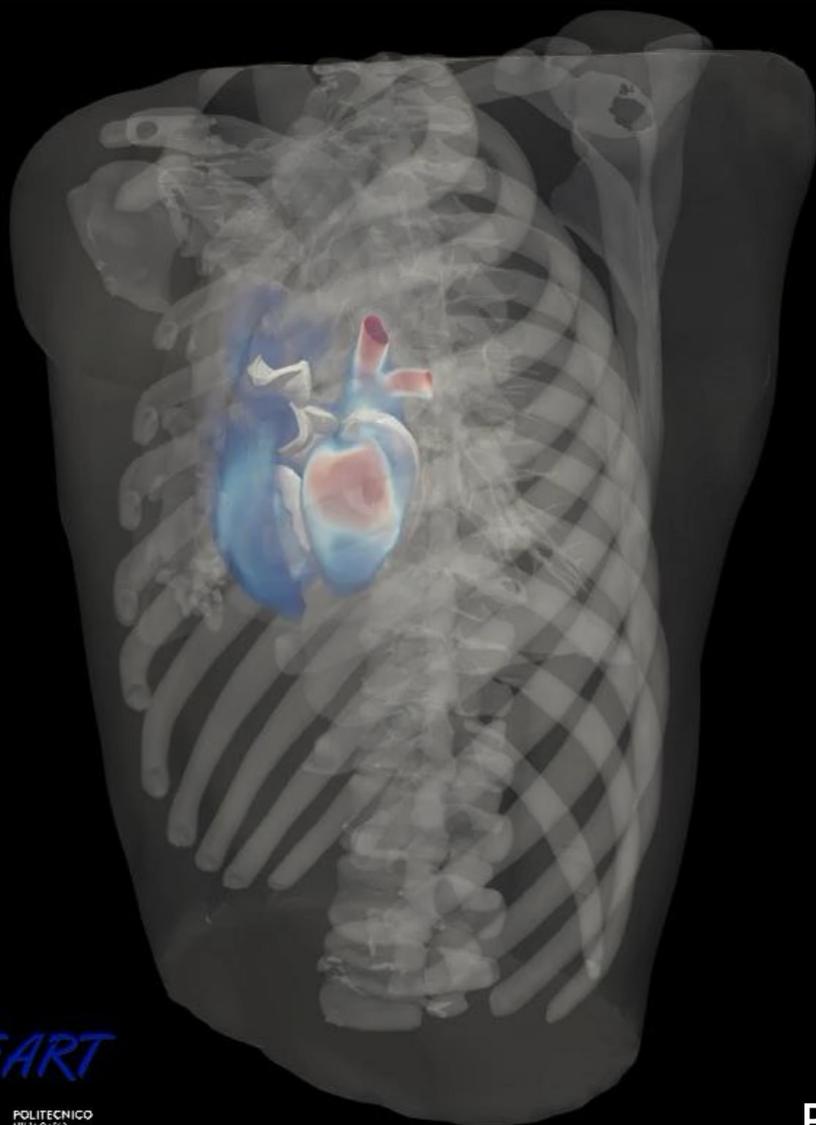
Blood pressure



Blood vorticity

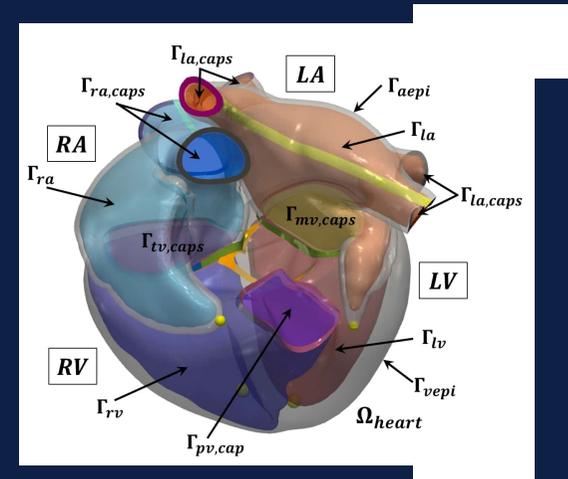


Fluid dynamics of the left heart (A.Zingaro)



Blood dynamics of the whole heart (M.Bucelli)

A Few Selected Clinical Applications





Risk assessment for AAA

(Abdominal Aortic Aneurisms)

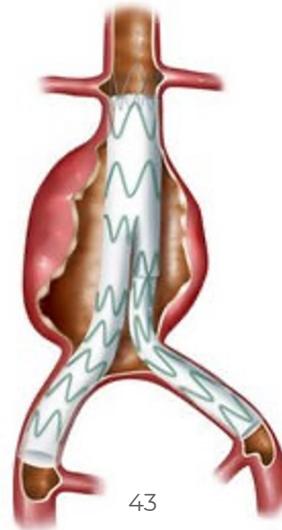
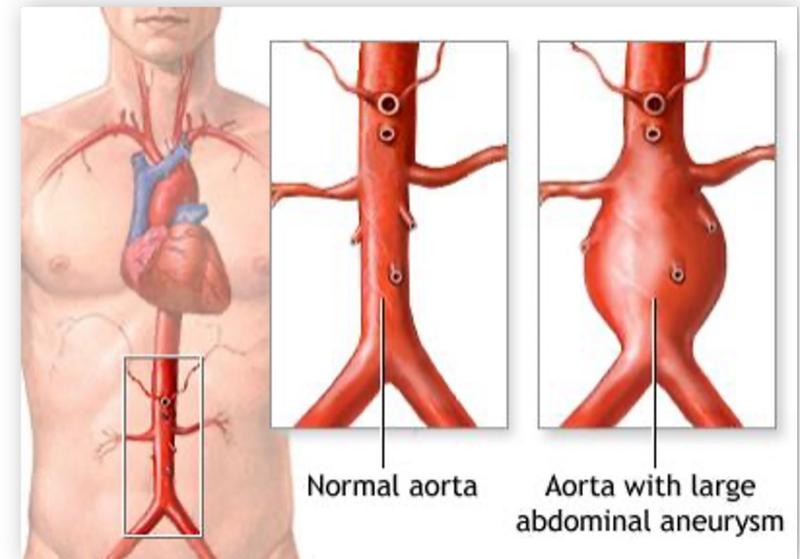
Risk assessment of abdominal aortic aneurysm (AAA)

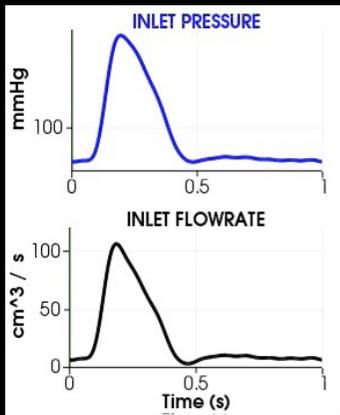
Problem:

- Affects 8% of males over the age of 65
- Mortality if ruptured is 85-90%
- Surgical intervention if AAA diameter > 5 cm
- Diameter manually measured
- Proven not to be the best predictive choice

Methods:

- Semi-automatic CT segmentation
- Mesh generation and fluid/mechanical simulations

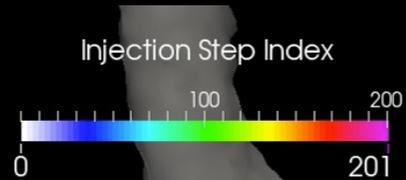




Patient : p057
 Age : 64
 Sex : M
 Pressure : 140/85
 ILT : none
 History : NA

aXurge

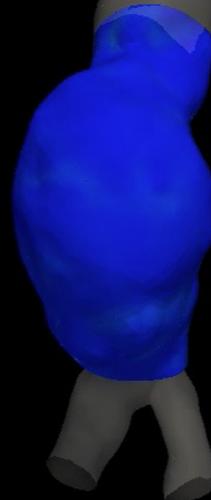
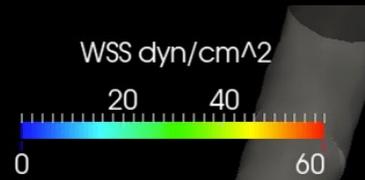
@cmcs-epfl



Inlet section



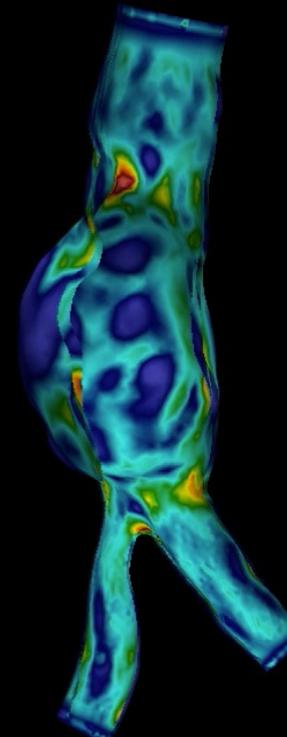
Time: 0.00 s



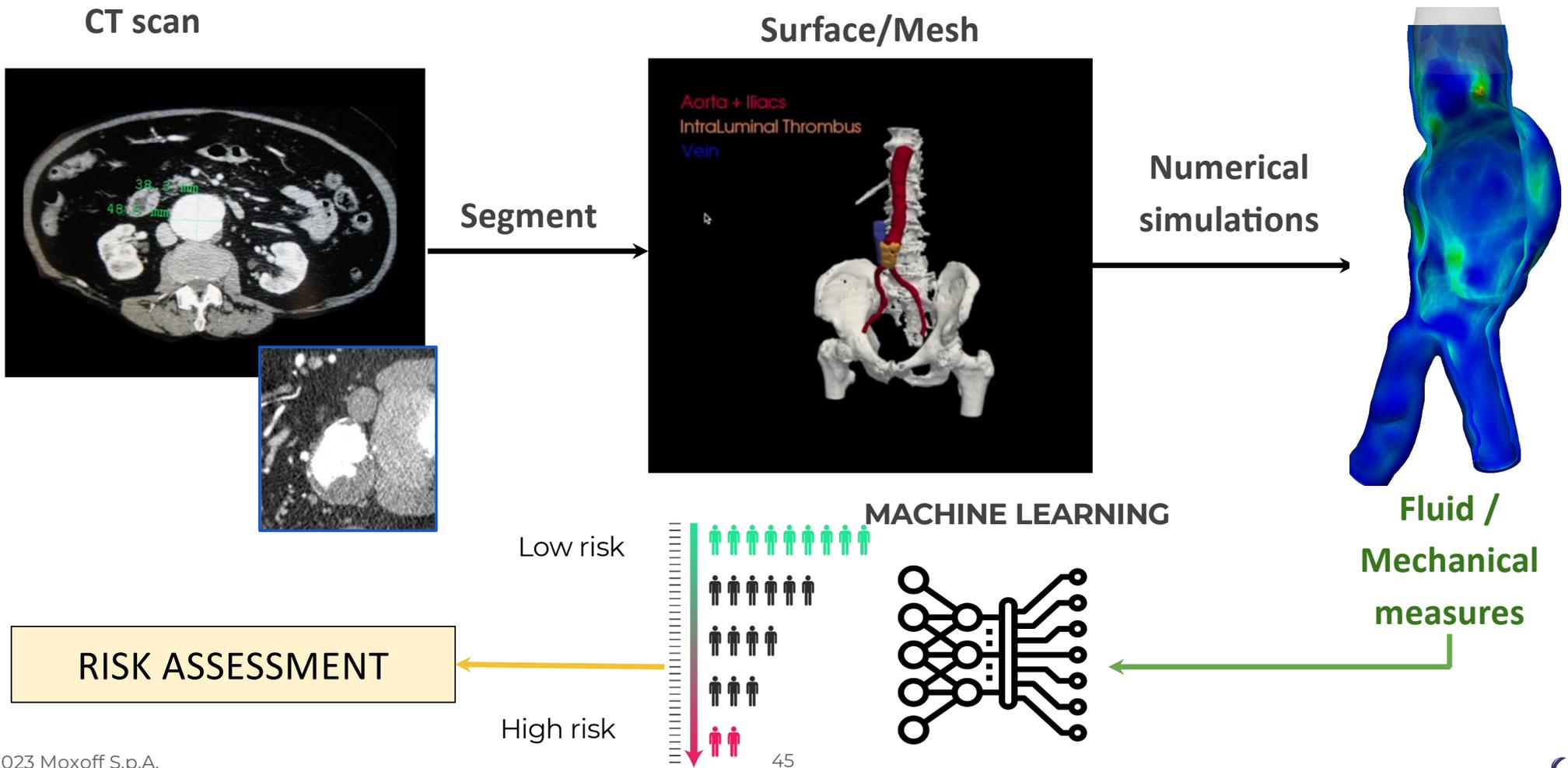
Systolic peak

Time: 0.2 s

VonMises Stress N/cm²



Risk assessment of abdominal aortic aneurysm (AAA)



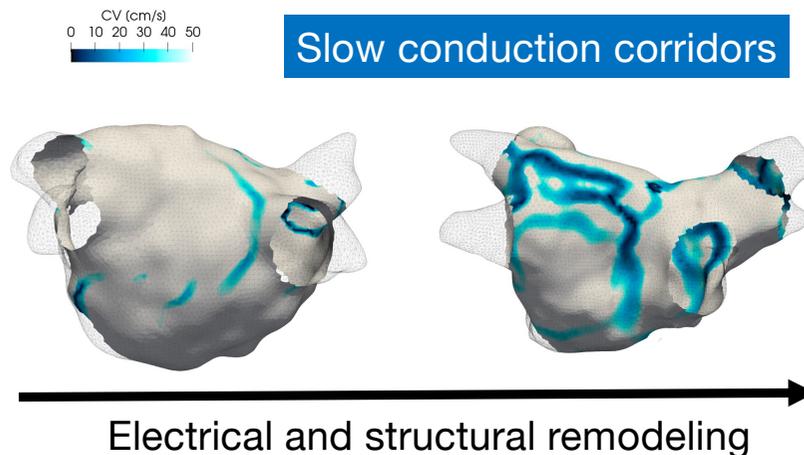
Atrial fibrillation (AF)



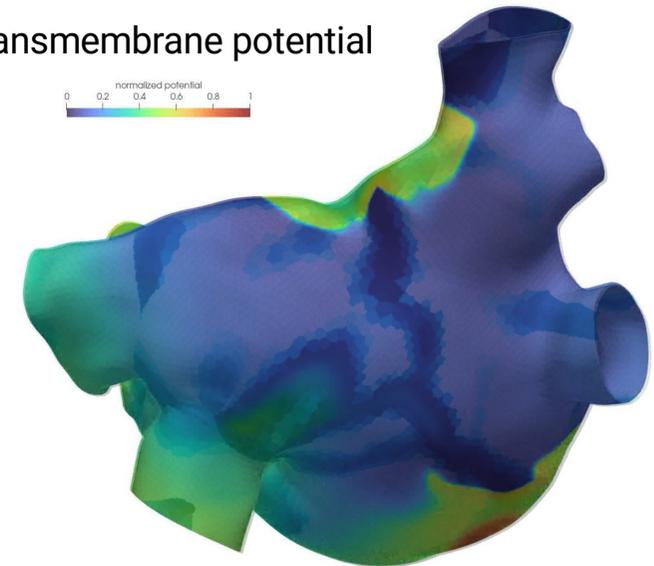
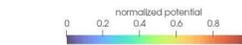
Clinical question: which are the mechanisms behind AF progression?

IRCCS
HUMANITAS
RESEARCH HOSPITAL

I.R.C.C.S. Ospedale
San Raffaele



Transmembrane potential



- **slow conduction corridors** and **pivot points** quantitatively characterize **AF progression**
- Numerical simulations confirm the role of **slow conduction corridors** in **AF sustainment** (localized reentry anchoring)

A. Frontera, S. Pagani, L.R. Limite et al., *JACC: Clinical Electrophysiology*, 2022

S. Pagani, L. Dede', A. Frontera et al., *Frontiers in Physiology*, 2021

Hypertrophic Cardiomyopathy (HCM)



Clinical question: can CFD simulations guide obstruction assessment and pre-operative design of septal myectomy?

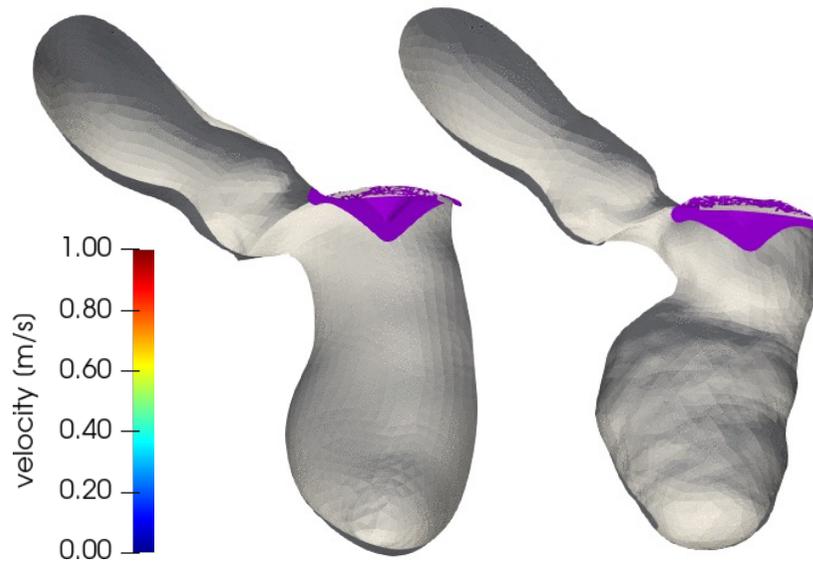


Simulation

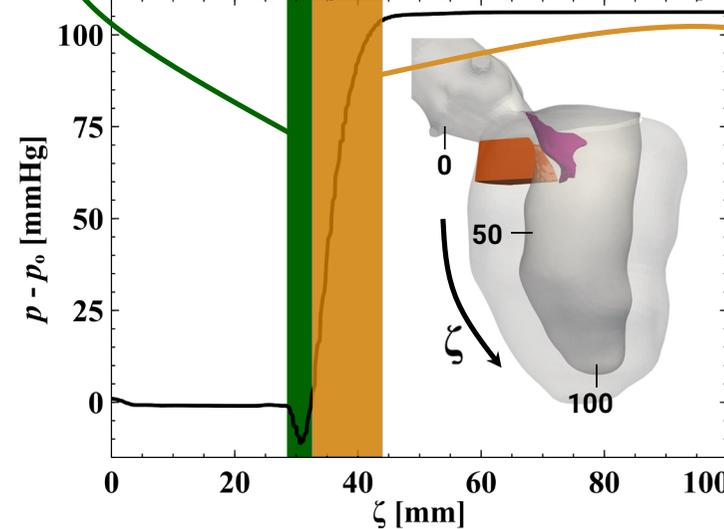
non-obstructive HCM

obstructive HCM

Obstructive HCM-induced **Venturi effect** causing systolic anterior motion



pressure along septum at systolic peak from coronary ostium down to the apex



obstruction severity: **extent** and **amplitude** of **subaortic pressure gradient**

indication for surgical treatment: **portion to remove** by septal myectomy

I. Fumagalli, M. Fedele, C. Vergara, et al., *Computers in Biology and Medicine*, 2020

I. Fumagalli, P. Vitullo, C. Vergara, et al., *Frontiers in Physiology*, 2022

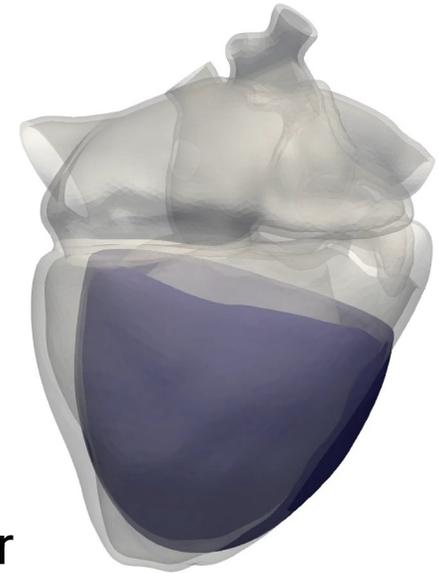
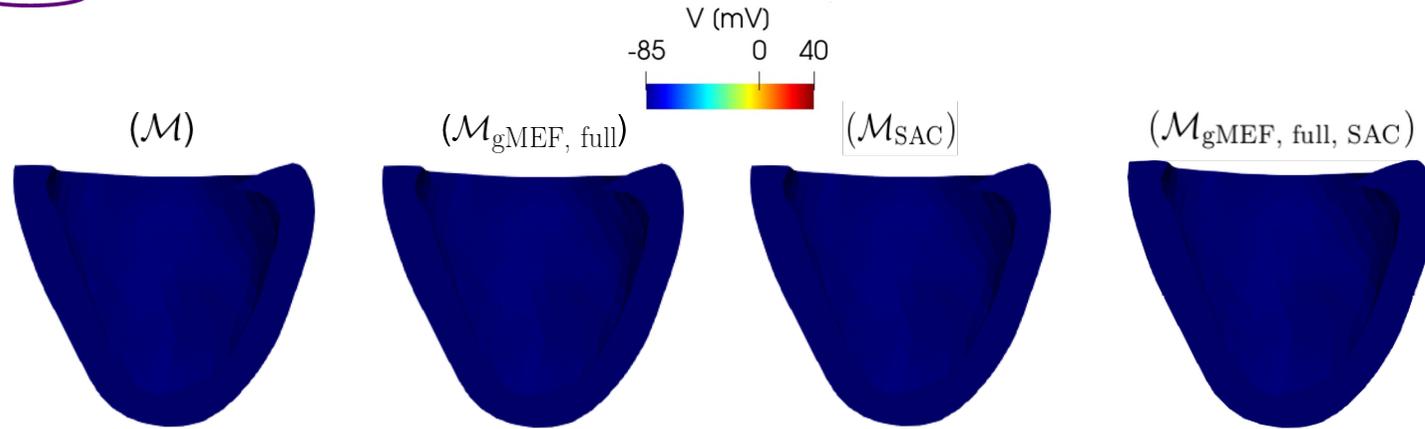
Ventricular tachycardia and fibrillation



Clinical question: are ventricular tachycardia and fibrillation better simulated by accounting for mechanical deformation?



JOHNS HOPKINS
UNIVERSITY



From **electrophysiology** to **electromechanics**
(geometry-mediated mechano-electric feedbacks and stretch-activated channels)

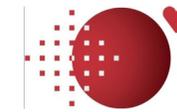
Electrophysiology and **electromechanics** simulations may differ in **conduction velocity**, **electric stability** and **hemodynamic stability**

M. Salvador, M. Fedele, P.C. Africa et al., *Computers in Biology and Medicine*, 2021
M. Salvador, F. Regazzoni, S. Pagani et al., *Computers in Biology and Medicine*, 2022

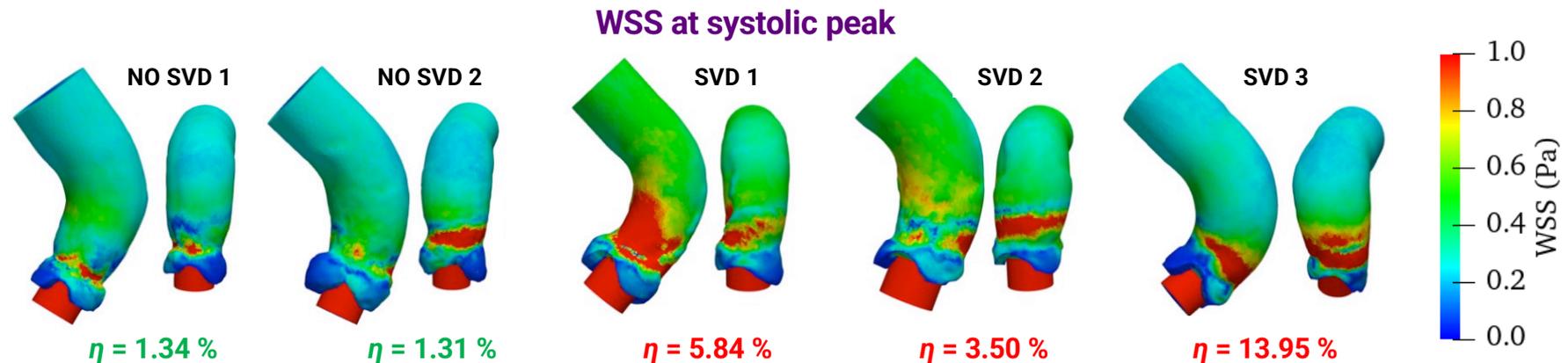
Transcatheter Aortic Valve Implantation (TAVI)



Clinical question: which are the predictive indicators of TAVI Structural Valve Deterioration (SVD)?



Centro Cardiologico
Monzino



- Analysis based on **pre-implantation** data only
- **WSS stronger** and more **persistent** in **SVD** cases
- η index discriminating SVD from NO-SVD, based on **Time-Averaged WSS (TAWSS) Critical Area (CA)**:

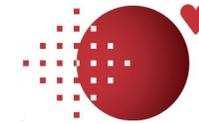
$$\eta = \frac{|CA|}{|\Gamma_{\text{wall}}|}, \text{ with } CA = \{\mathbf{x} \in \Gamma_{\text{wall}} : TAWSS(\mathbf{x}) > 0.5\text{Pa}\}$$

I. Fumagalli, R. Polidori, F. Renzi et al., *MOX Report*, Politecnico di Milano, 2022

Estimating cardiac blood flow maps

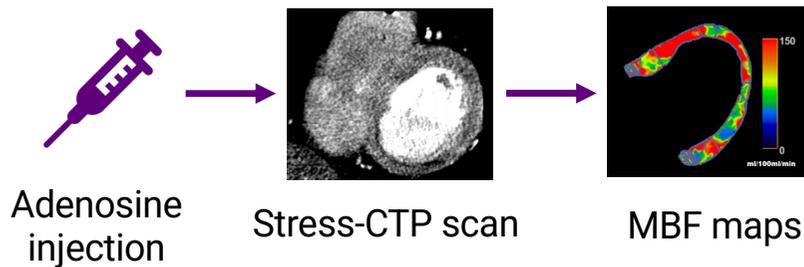


Clinical question: can we replace CT scans and stress protocols with a computational estimation of myocardial blood flow maps?



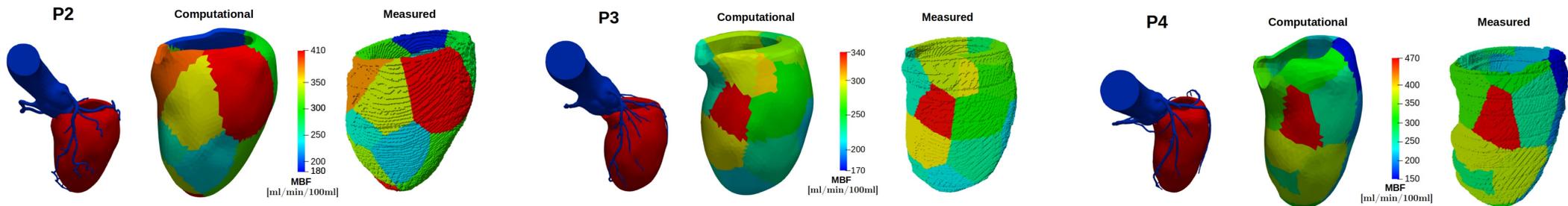
Centro Cardiologico
Monzino

Clinical pipeline



Consistency tests: calibration of perfusion model on available maps yields **excellent agreement**

Ongoing: calibration of patient-specific models based on **pressure data only** (no maps)



S. Di Gregorio, C. Vergara, G. Montino Pelagi et al., *European Journal of Nuclear Medicine and Molecular Imaging* 2022

Dealing with Complexity

How Fast Supercomputers Are?



FRONTIER: 1.6 Billion Billion FLOPS

8 Billion People on Earth

200 Million Operations per Second per Inhabitant

A complete simulation of a single heartbeat (I

Requires at least 20M degrees of freedom on 192 cores on a supercomputing facility

May last up to 48 hours

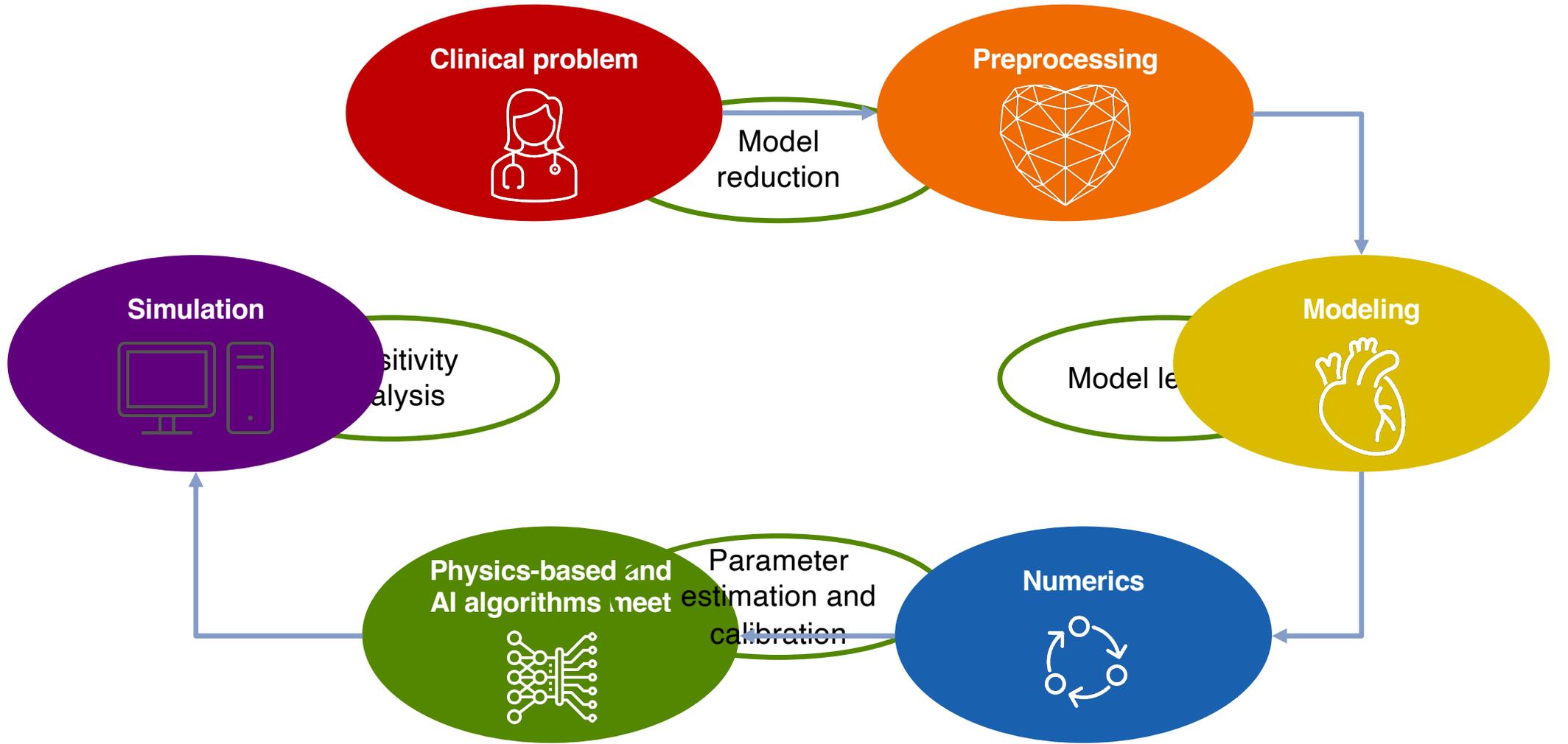
Costs about 2000 euros

Consumes 100kWh of energy

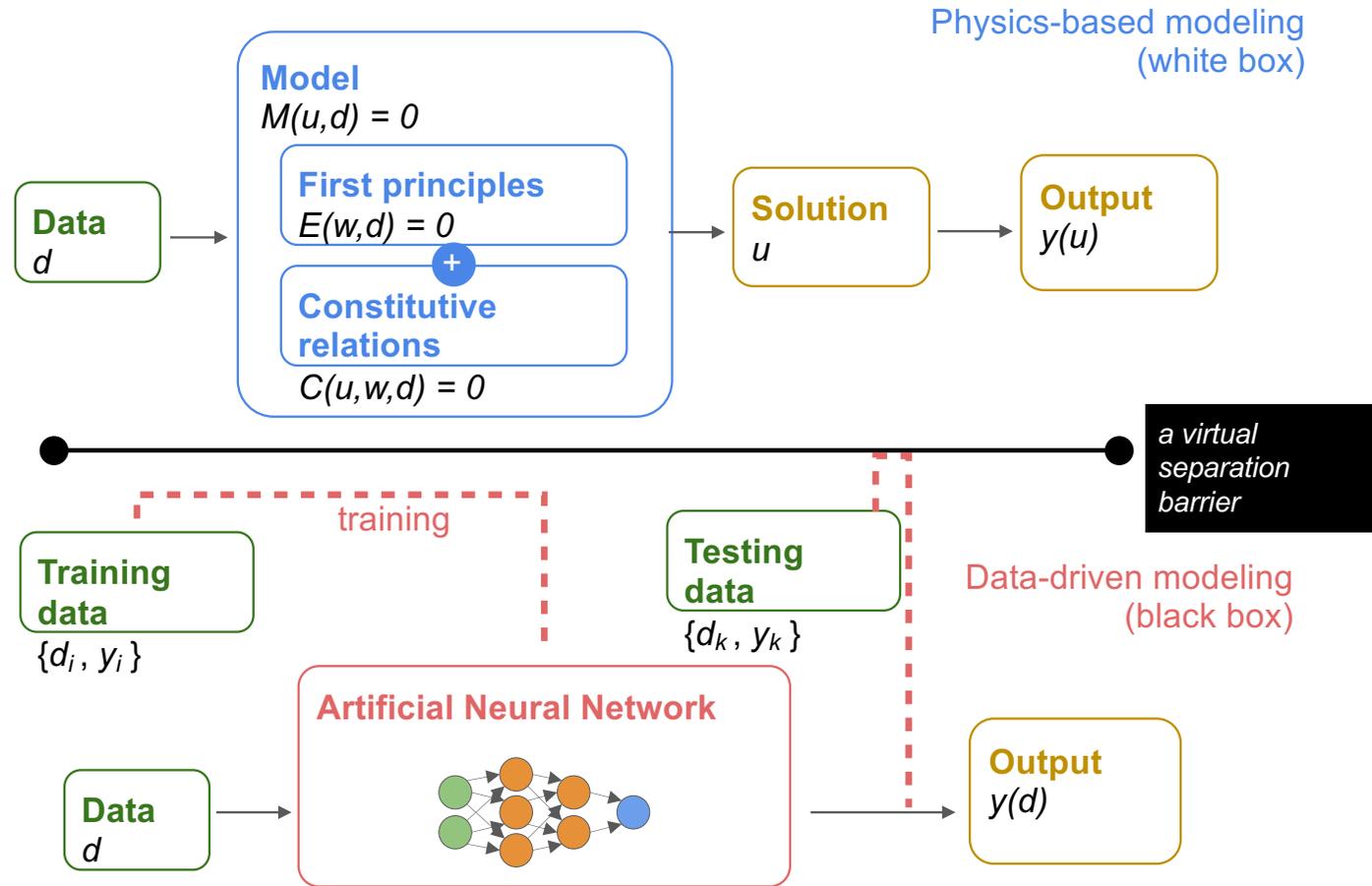
Produces 35kg of CO₂ (without accounting for the additional CO₂ produced for the cooling of the cluster)

Developing better models and more efficient and accurate numerical methods is of paramount importance

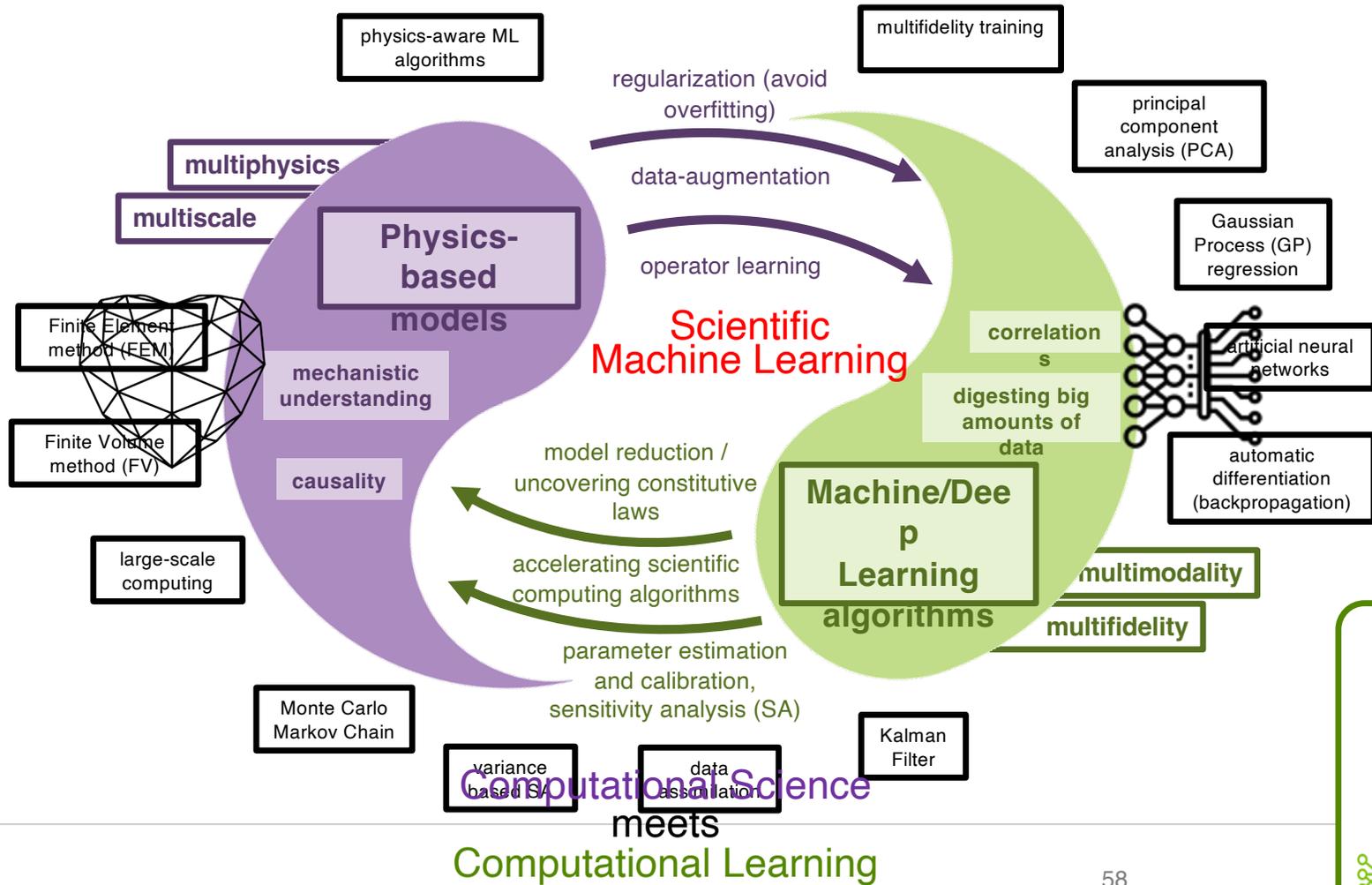
A new exig
in Engineer
Rapid and r



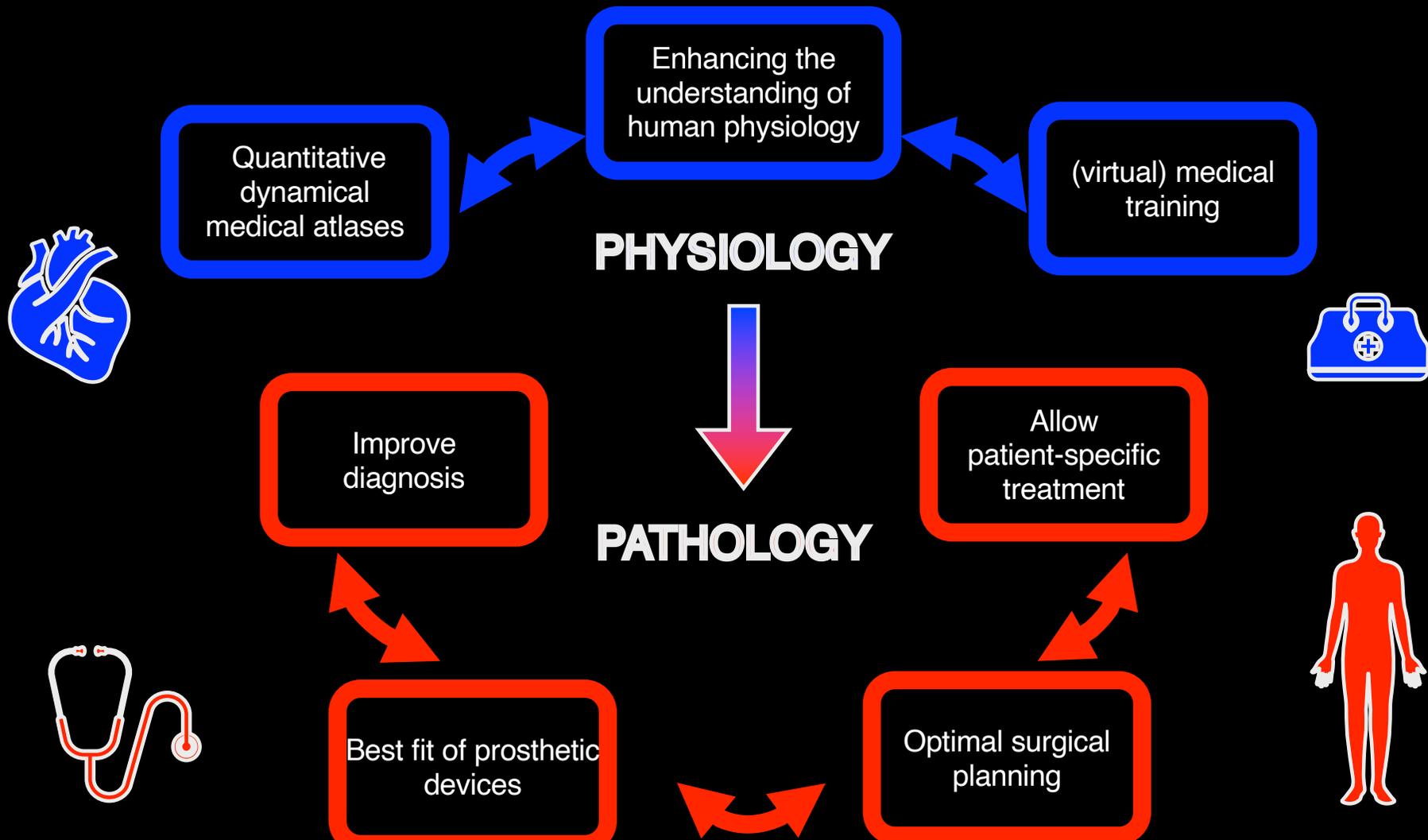
Physics-based vs Data-driven modeling



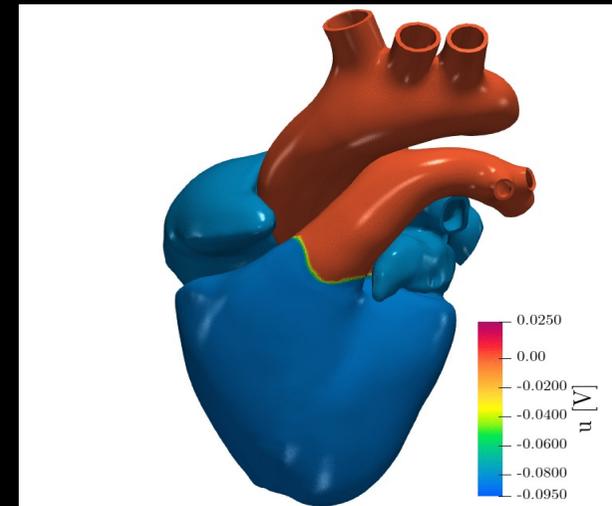
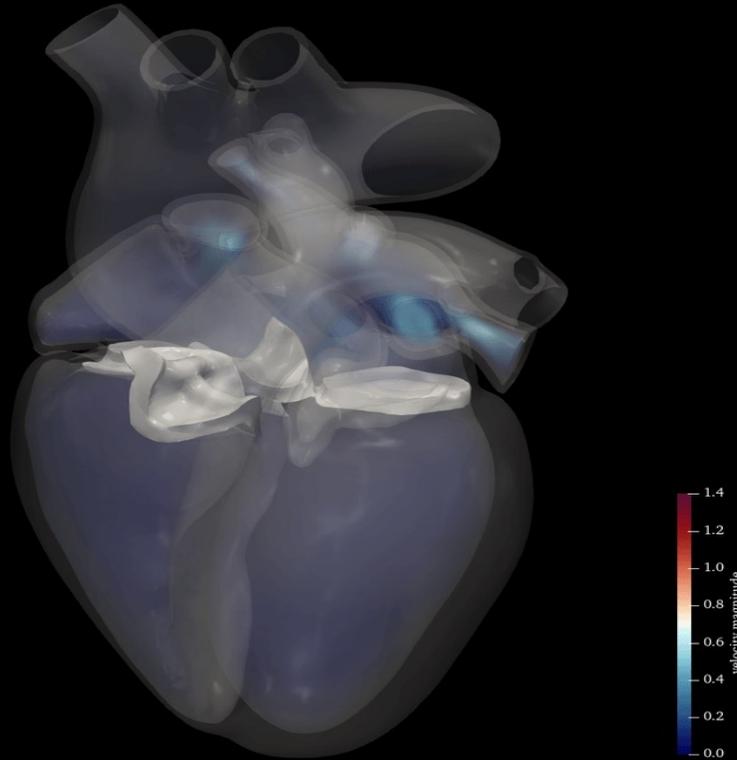
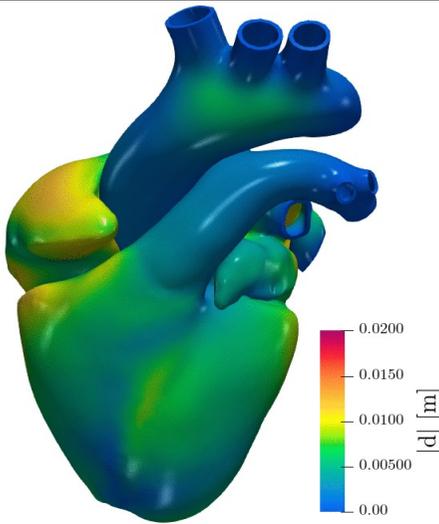
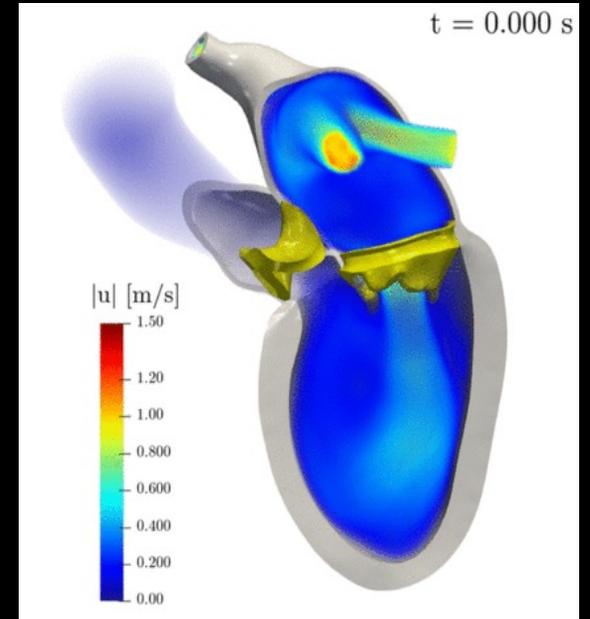
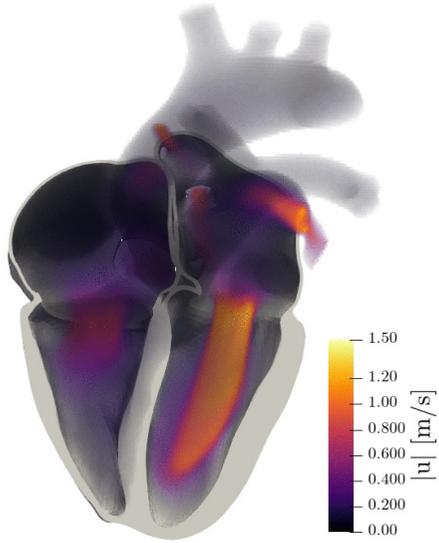
A cooperative game



TOWARD DIGITAL HEALTH



THANK YOU



THANK YOU

