

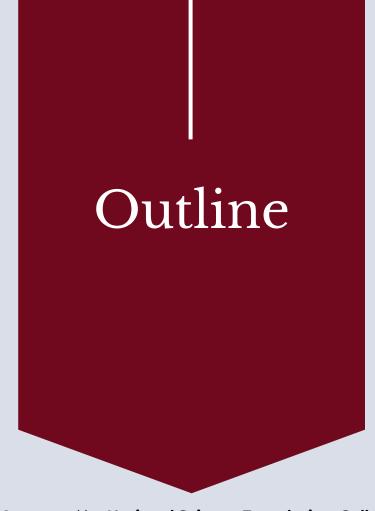






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Rome Sept 10th, 2025





Clinical Background

What is Radiofrequency Ablation (RFA)?



Multiphysics Model

Why do we need multiphysics modeling? What physics are we including? How do we implement it?



Domain Decomposition

How do we handle multiple domains?
Why do we need Domain Decomposition?



Results

What did we find?



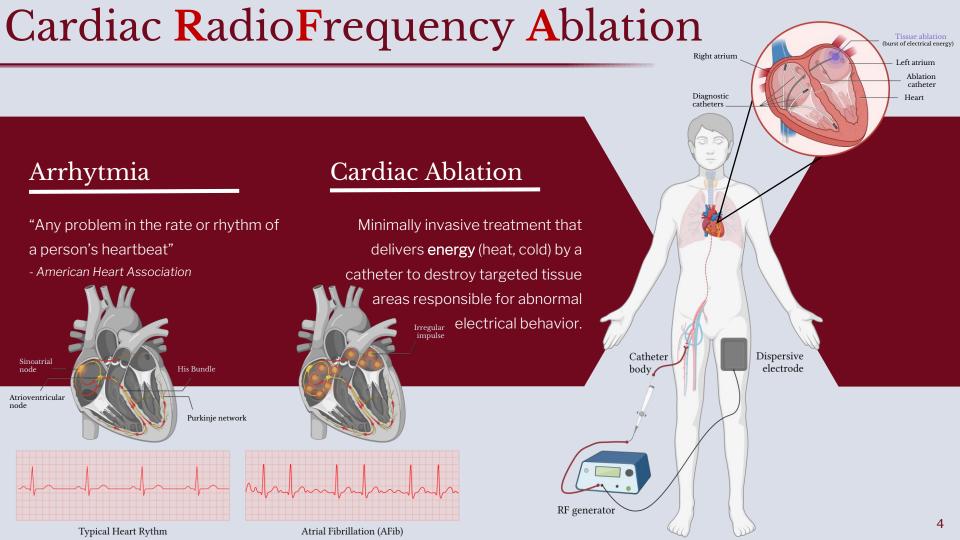
Conclusion & Future work

How can we improve our framework?



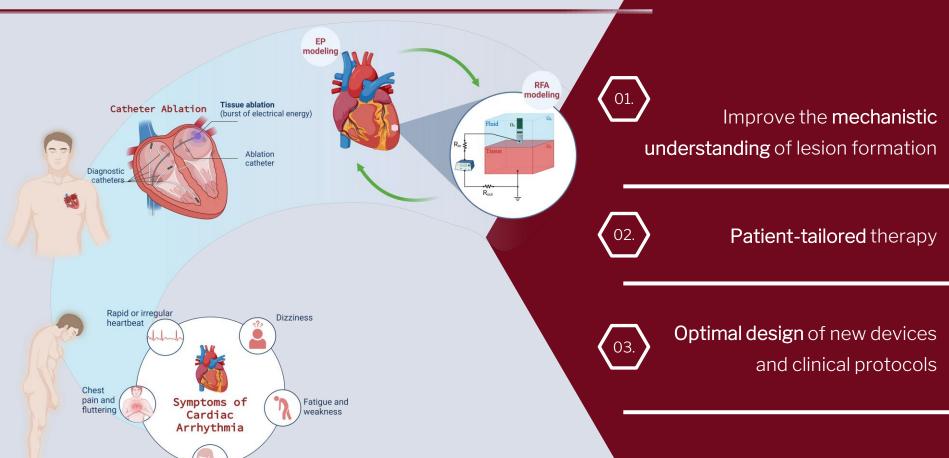


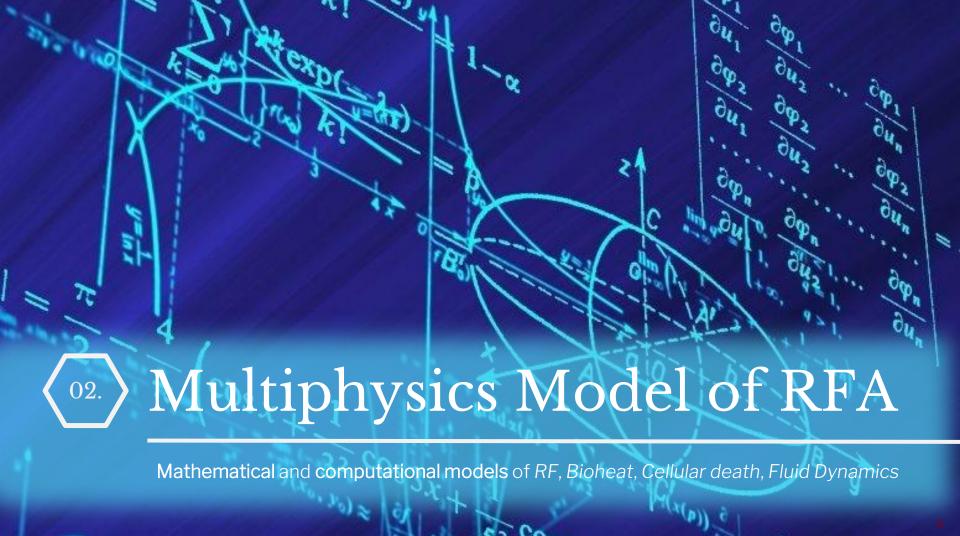
Cardiac Arrhytmias, Radiofrequency Ablation and Motivation to Modeling



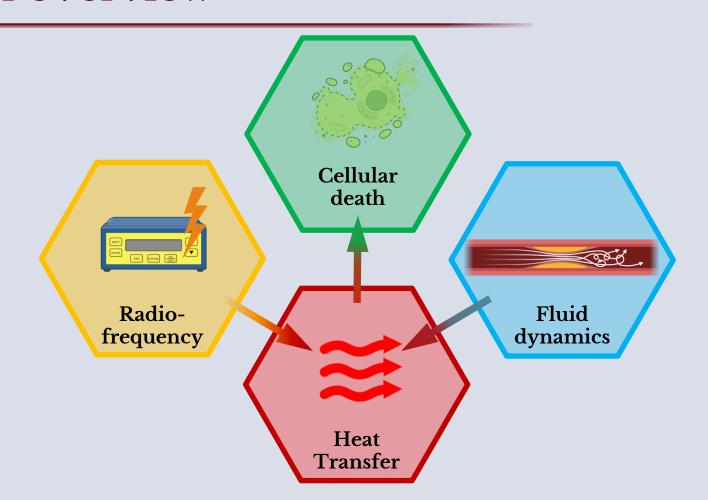
Why do we need modeling?

Wheezing and shortness of breath





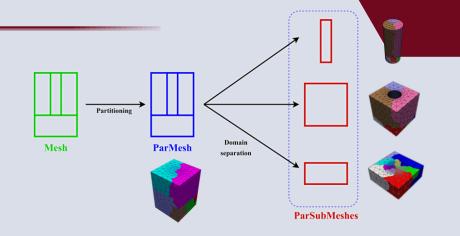
Model overview

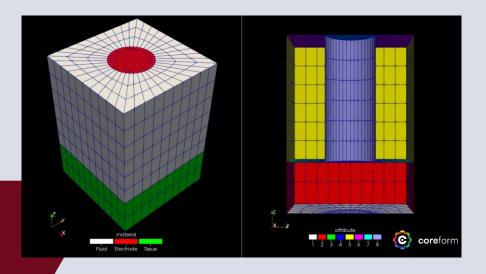


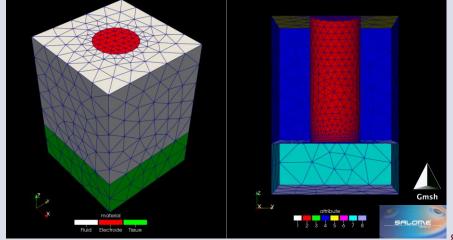
Geometry

Idealized geometry (...for now)

- Three physical domains: Tissue, Fluid, Electrode
- Available both as *structured* and *unstructured* meshes



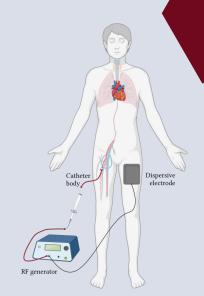




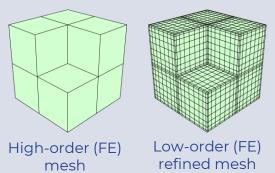
Radiofrequency source

Quasi-static Maxwell's equation

$$\nabla \cdot (\boldsymbol{\sigma} \nabla \Phi) = 0$$
 in Ω_f , Ω_t



- Numerical implementation
 - Spatial discretization: Q_4 elements
 - Linear solver: CG preconditioned with Low-Order-Refined¹ (LOR) coupled to AMG (PA)



t = Tissue

f = Fluid e - Electrode

Heat Transfer

Pennes' bioheat equation

$$\begin{split} \rho_t c_t \frac{\partial T}{\partial t} &= \nabla \cdot (\textbf{k}_t \nabla T) + Q_m + Q_p + Q_s & \text{in } \Omega_t \\ \rho_f c_f \frac{\partial T}{\partial t} &= \nabla \cdot (\textbf{k}_f \nabla T) - \textbf{u} \cdot \nabla T & \text{in } \Omega_f \\ \rho_e c_e \frac{\partial T}{\partial t} &= \nabla \cdot (\textbf{k}_e \nabla T) & \text{in } \Omega_e \\ Q_s &= \boldsymbol{\sigma} \cdot |\textbf{E}|^2 \quad [\text{W/m}^3] \end{split}$$

$$Q_m = \text{metabolic (constant)}$$

 $Q_p = -w_b \rho_b c_b (T_t - T_{bp})$

 T_{bp} constant temperature of blood perfusing the tissue

 w_b is a perfusion coefficient depending on the cell viability so that $0 < w_b(N) \le w_{b,max}$

Numerical implementation

- Spatial discretization: Q_2 elements
- Time discretization: two-step third-order Singly-diagonal implicit Runge-Kutta scheme (SDIRK23)
- Linear solver: GMRES preconditioned with matrix-free Jacobi smoother (PA)

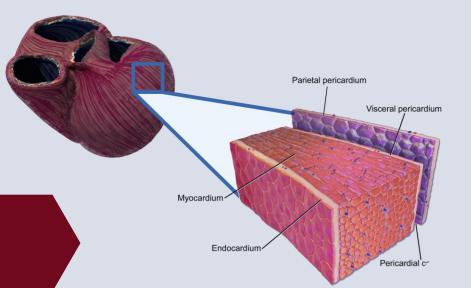
e - Electrode

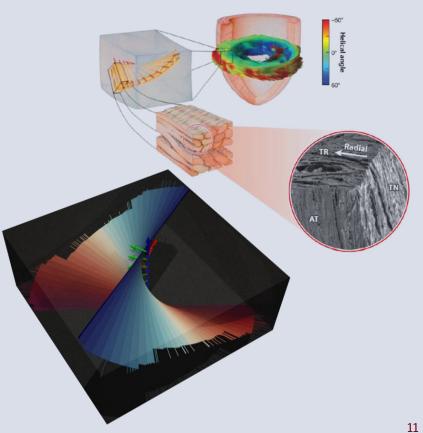
Anisotropy

Anisotropic thermo-electrical conductive properties in the tissue due to fiber microstructure

$$\boldsymbol{\sigma} = \mathbf{R} \begin{bmatrix} \sigma_f & 0 & 0 \\ 0 & \sigma_{\perp}^y & 0 \\ 0 & 0 & \sigma_{\perp}^z \end{bmatrix} \mathbf{R}^{\mathsf{T}} \qquad \qquad \mathbf{k} = \mathbf{R} \begin{bmatrix} k_f & 0 & 0 \\ 0 & k_{\perp}^y & 0 \\ 0 & 0 & k_{\perp}^z \end{bmatrix} \mathbf{R}^{\mathsf{T}}$$

$$R = R_z(\alpha) R_y(\beta) R_x(\gamma)$$





Cellular death

Vulnerable (Unfolded)

Alive (Native)

$$N \xrightarrow{\upsilon_1} U \xrightarrow{\upsilon_2} D \xrightarrow{\text{Dead}}_{\text{(Denatured)}}$$

Numerical implementation

• Eigenvalue method to solve

$$\frac{dX}{dt} = \mathcal{A} X, \quad X^T = [N, U, D]^T$$
$$X = \mathbf{P} e^{\int_0^t diag(\lambda)d\tau} \mathbf{P}^{-1} \mathbf{x}(\mathbf{0})$$

Three-state cell-death model

$$\begin{split} \frac{dN}{dt} &= -\upsilon_1 N + \upsilon_3 U \\ \frac{dU}{dt} &= \upsilon_1 N - \upsilon_2 U - \upsilon_3 U & \text{ in } \Omega_t \\ \frac{dD}{dt} &= \upsilon_2 U \\ \upsilon_i(T) &= A_i \bar{e}^{\Delta E/RT} \\ N + U + D &= 1 \end{split}$$

$$\mathcal{A} = \begin{bmatrix} -v_1 & v_3 & 0 \\ v_1 & -(v_2 + v_3) & 0 \\ 0 & v_2 & 0 \end{bmatrix} = P \Lambda P^{-1} \qquad X_{n+1} = P e^{\Lambda \Delta t} P^{-1} X_n$$

• Symbolic evaluation of eigenpairs $\{\Lambda, P\}$

Fluid Dynamics

Incompressible Navier Stokes equation

$$\frac{\partial \mathbf{u}}{\partial t} + \mathbf{u}^* \cdot \nabla \mathbf{u} - \nabla \cdot \boldsymbol{\tau} + \nabla \mathbf{p} = \mathbf{f}$$
$$\nabla \cdot \mathbf{u} = 0$$

$$u^* = \sum_{i=0}^{p} (\beta_i u_{n+1-i} \cdot \nabla) u_{n+1}$$
 extrapolated velocity (semi-implicit convection)

$$au$$
 viscous stress tensor $au = egin{cases}
u \nabla u & \text{Stiff strain} \\
u (\nabla u + \nabla u^T) & \text{Viscous strain}
\end{cases}$

Numerical implementation

ALgebraic splitting Time ADaptive solver for Incompressible Navier-Stokes (ALADINS)¹

- Time discretization: BDF up to order 3
- Spatial discretization: stable pair $Q_2 Q_1$
- Time adaptivity (Pressure correction- based)
- Algebraic system:

$$\mathcal{A} = \begin{bmatrix} \mathcal{C} & \mathcal{G} \\ \mathcal{D} \end{bmatrix}, \quad \text{with } \mathcal{C} = \frac{\alpha}{\Delta t} \mathcal{M} + \nu \mathcal{K} + \mathcal{N}(\boldsymbol{u}), \quad \mathcal{G} = -\mathcal{D}^T$$

• FGMRES preconditioned with block preconditioner $\widehat{\mathcal{P}}$

Fluid Dynamic: Preconditioner

Navier-Stokes **Block Preconditioner** \hat{P}

Algebraic splitting pressure corrected preconditioner^{1,2}

Where we assumed:

- $\rightarrow \hat{S}$ preconditioner for the Schur Complement $\Sigma = \mathcal{DC}^{-1}\mathcal{G}$, $\hat{S} = \mathcal{D} \operatorname{diag}(\mathcal{M}_v)^{-1}\mathcal{G}$
- $\rightarrow \hat{\mathcal{C}}$ approximation to \mathcal{C} ,
- $\rightarrow \mathcal{H}_1, \mathcal{H}_2$ cheap approximation to \mathcal{C}^{-1}
- $\rightarrow Q$, \mathcal{R} pressure correction matrices, depend on the chosen factorization method.
- **Factorization method:** High-Order Yosida (HOY) $\mathcal{A}_{HOY} = \begin{bmatrix} \hat{\mathcal{C}} \\ \mathcal{D} \\ -\hat{\mathcal{S}} \end{bmatrix} \begin{bmatrix} I & \mathcal{C}^{-1}\mathcal{G} \\ 0 \end{bmatrix}$ (momentum preserving)

$$\mathcal{A}_{HOY} = \begin{bmatrix} \hat{\mathcal{C}} & \\ \mathcal{D} & -\hat{\mathcal{S}} \end{bmatrix} \begin{bmatrix} I & \mathcal{C}^{-1}\mathcal{G} \\ & \mathcal{Q} \end{bmatrix}$$

$$\mathcal{A} \approx \begin{bmatrix} \hat{\mathcal{C}} & \\ \mathcal{D} & -\hat{\mathcal{S}} \end{bmatrix} \begin{bmatrix} \mathbf{I} & \mathcal{H}_2 \mathcal{G} \mathcal{R} \\ & \mathcal{Q} \end{bmatrix}$$

$$\begin{split} \mathcal{H}_1 &= (\frac{\alpha}{\Delta t}\mathcal{M})^{-1}, \quad \mathcal{H}_2 = \mathcal{C}^{-1} \\ \mathcal{R} &= I, \quad \mathcal{Q} = (\mathcal{D}\mathcal{H}_1\mathcal{C}\;\mathcal{H}_1\mathcal{G})^{-1}\hat{\mathcal{S}} \\ \boldsymbol{D}_k &= -\mathcal{D}(-\mathcal{H}_1\mathcal{F})^k\mathcal{G}, \quad k>0 \qquad \mathcal{F} = \nu\mathcal{K} + \mathcal{N}(\boldsymbol{u}) \end{split}$$

Pressure correction step: $y = Q^{-1}x = \begin{cases} \hat{\delta}z_1 = D_1z_0 \\ \hat{\delta}z_2 = D_2z_1 + D_1z_0 \end{cases}$

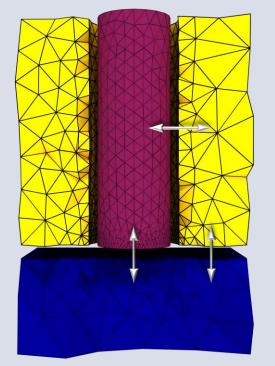
Time adaptivity:

Adaptive selection of time step based on a posteriori pressure-correction based error estimator

- $\rightarrow \alpha$ Safety factor for time adaptivity
- $\rightarrow \epsilon$: **Tolerance** for time adaptivity
- $\rightarrow z_a$ pressure correction of order q

$$\Delta t = \chi \, \Delta t_{old}$$

$$\chi = \min(\max\left(\alpha \cdot (\frac{\epsilon \, \Delta t_{old}}{\|\mathbf{z}_q\|})^{\frac{1}{q}}, \chi_{min}\right), \chi_{max})$$

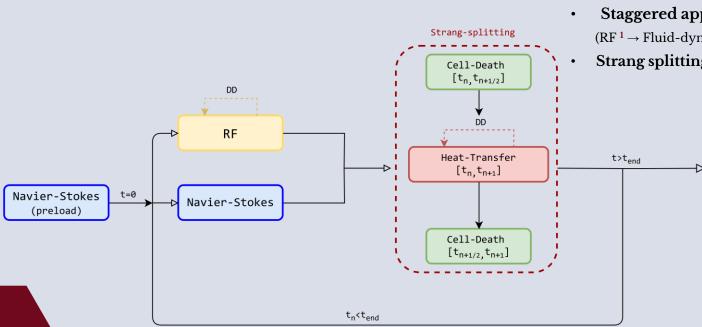




(03.) Domain Decomposition

Modeling interface between tissue, electrode and fluid domains

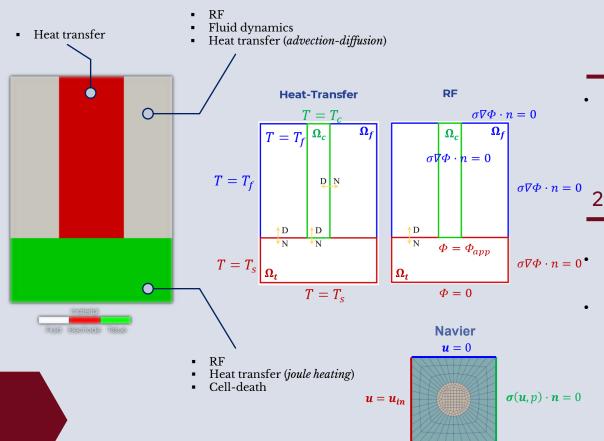
Two-level segregation approach



1. Multiphysics segregation

- Staggered approach for multiphysics
 (RF¹→ Fluid-dynamics →Heat transfer¹→Cell-death)
- Strang splitting for HT/CD problems $O(\Delta t^2)$

Two-level segregation approach



1. Robin-Robin implemented and tested for RF subproblem

 $\mathbf{u} = 0$

1. Multiphysics segregation

Staggered approach for multiphysics
(RF¹ → Fluid-dynamics → Heat transfer¹ → Cell-death)

- 2. Substructuring by (physical) subdomains
 - Nonoverlapping domain decomposition

Based on physical domains

Dirichlet-Neumann, Robin-Robin coupling (DN, RR)¹

For transmission conditions at the physical interfaces

 $\begin{aligned} \mathsf{RF} & \quad \nabla \cdot \sigma_1 \, \nabla \phi_1^{n+1} = f_1 & \text{in } \Omega_1 \\ \phi_1^{n+1} = \phi_2^n & \text{on } \Gamma_{12} \end{aligned} \\ & \quad \nabla \cdot \sigma_2 \, \nabla \phi_2^{n+1} = f_2 & \text{in } \Omega_2 \\ & \quad \sigma_2 \nabla \phi_2^{n+1} \cdot \boldsymbol{n} = \sigma_1 \nabla \phi_1^{n+1} \cdot \boldsymbol{n} & \text{on } \Gamma_{12} \end{aligned}$





Solvers verifications, Domain Decomposition convergence, RFA

Solvers verification: Heat transfer

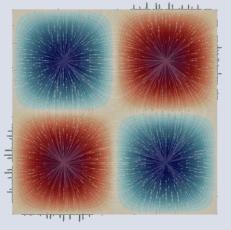
Convergence (MMS on squared domain) +

+ toy problems

Advection-diffusion

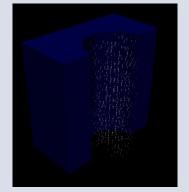
Time-dependent square plate heating

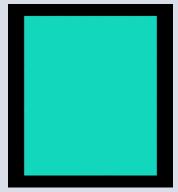




| Heat-Transfer ($o = 2$) | | | | | |
|---------------------------|---------|-------------|------------|-------------|------------|
| DOFs | h | L^2 error | L^2 rate | H^1 error | H^1 rate |
| 81 | 0.25 | 0.01445 | 0 | 0.4044 | 0 |
| 289 | 0.125 | 0.001933 | 2.902 | 0.102 | 1.987 |
| 1089 | 0.0625 | 0.0002451 | 2.979 | 0.02553 | 1.998 |
| 4225 | 0.03125 | 3.075e-05 | 2.995 | 0.006383 | 2 |
| 16641 | 0.01562 | 3.847e-06 | 2.999 | 0.001596 | 2 |







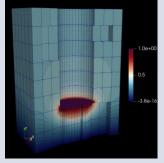


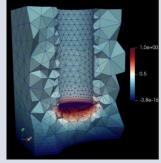
Convective cooling of sphere

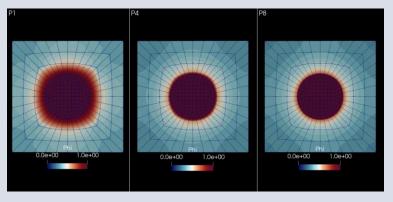
Solvers verification: RF

Convergence (MMS on squared domain) + toy problems

| RF | | | | | | |
|-------|---------|-------------------------|------------|------------------------|------------|--|
| Dofs | h | L^2 error | L^2 rate | H^1 error | H^1 rate | |
| 289 | 0.2686 | 2.425×10^{-5} | 0 | 0.001126 | 0 | |
| 1089 | 0.1343 | 7.761×10^{-7} | 4.965 | 7.144×10^{-5} | 3.979 | |
| 4225 | 0.06716 | 2.443×10^{-8} | 4.99 | 4.478×10^{-6} | 3.996 | |
| 16641 | 0.03358 | 8.309×10^{-10} | 4.878 | 2.802×10^{-7} | 3.998 | |



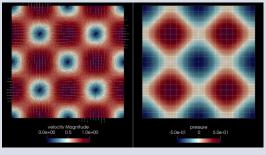




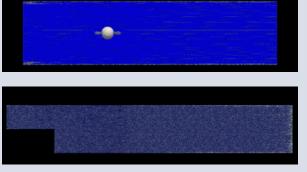
Solvers verification: Navier-Stokes

Convergence (MMS on squared domain) + toy problems

| Navier-Stokes | | | | | | |
|---------------|--------|--------|--------------------------|--------------------------|----------------|----------------|
| DOFs u | DOFs p | h | L^2 error (u) | L^2 error (p) | L^2 rate (u) | L^2 rate (p) |
| 242 | 36 | 0.2 | 1.99644×10^{-2} | 0.110601 | 0 | 0 |
| 882 | 121 | 0.1 | 5.26485×10^{-3} | 0.0185581 | 1.92297 | 2.57525 |
| 3362 | 441 | 0.05 | 7.36945×10^{-4} | 0.00379613 | 2.83676 | 2.28945 |
| 13122 | 1681 | | 5.4351×10^{-5} | | 3.76118 | 2.0196 |
| 51842 | 6561 | 0.0125 | 4.3819×10^{-6} | 2.99535×10^{-4} | 3.63268 | 1.64413 |





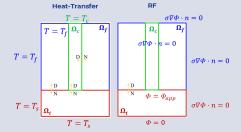




Domain Decomposition convergence

Subiteration convergence

Dirichlet-Neumann scheme exhibits linear convergence for Heat/RF problems.

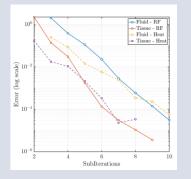


• h-refinement

Consistent results for DD convergence across different levels of **mesh refinement** For RF, RR conditions did not lead to significant improvements (but we did not analyze the optimal parameters derived form Fourier Analysis)

• p-refinement

Slower convergence rates for the DD algorithm



| h-Refinement | | | | | | |
|--------------|---------|--------|-------|----------------|--|--|
| Ref | Dofs RF | Dofs H | It-RF | It-Heat | | |
| 0 | 90k | 19k | 10 | 7.4 ± 0.70 | | |
| 1 | 673k | 150k | 9 | 7.4 ± 0.52 | | |
| 2 | 5M | 1.1M | 9 | 7.7 ± 0.67 | | |

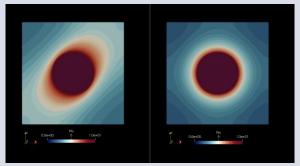
(a) Iteration count for DD convergence ($tol = 10^{-6}$) after h-refinement (uniform refinement). For heat problem, iterations taken over the first 10 timesteps. H = Heat, RF = Radio-Frequency.

| p-Refinement | | | | | | |
|----------------------|---------|--------|-------|-----------------|--|--|
| Order (H-RF) | Dofs RF | Dofs H | It-RF | It-Heat | | |
| $\overline{Q_1-Q_2}$ | 13k | 3k | 10 | 7.5 ± 0.53 | | |
| $Q_2 - Q_4$ | 90k | 15k | 10 | 7.4 ± 0.70 | | |
| $Q_4 - Q_8$ | 673k | 150k | 15 | 12.9 ± 2.92 | | |

(b) Iteration count for DD convergence ($tol = 10^{-6}$) after **p-refinement**. For heat problem, iterations taken over the first 10 timesteps. H = Heat, RF = RadioFrequency.

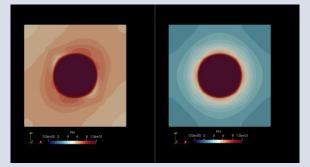
Anisotropy

RF, tissue only



Electric potential

RFA, three domains

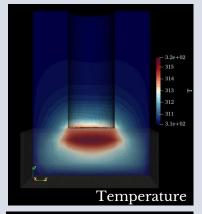


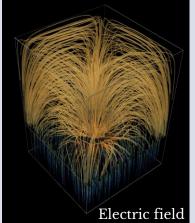
Electric field

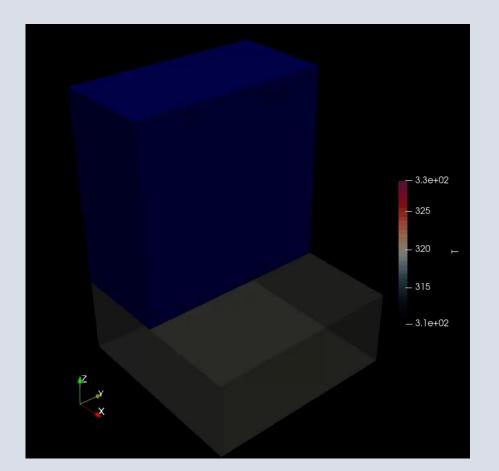
Electric potential

Temperature

(pseudo) RFA simulation



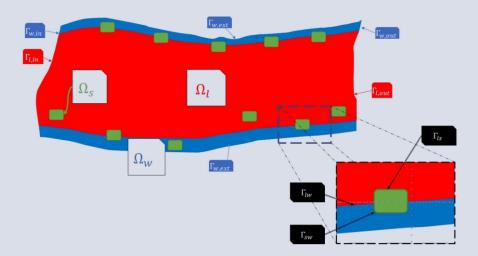






Other applications

- O Different ablation sources: MWA, HIFU, Laser
- **Different ablated tissues:** Liver, Intestine, Prostate, Bone
- Erosion of bioresorbable stents



Future plan

Multiphysics model

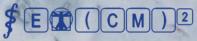
 $\sum_{k \in T} (\mathcal{R}_{\phi}, \tau \mathcal{L}_{stab}(w)),$

- 1.1 Fluid dynamics:
 - Strongly consistent stabilization (SUPG, GLS, VMS), and turbulence model (EFR) $\mathcal{L}_{stab} = \begin{cases} a \cdot \nabla & SUPG \\ -a \cdot \nabla + \nu \Delta & GLS \\ -a \cdot \nabla \nu \Delta & VMS \end{cases}$
- 1.2 Heat transfer:
 - Extend tissue bioheat equation to **Non-Fourier models** (SPL, DPL) $\rho c \frac{\partial^2 T}{\partial t^2}$, $\rho c \nabla \cdot (k \frac{\partial \nabla T}{\partial t})$
- 1.3 Mechanics:
 - Develop quasi-incompressible, hyperelastic, anisotropic **model for tissue mechanics**
 - Develop **electrode-tissue contact model** (leverage **MFEM-Tribol** interface)
- Improved computational efficiency
 - High fidelity simulations: Extend PA to all solvers and enable GPU acceleration
 - Clinically-suitable simulations: investigate use of Reduced Order Modeling (ROM) techniques (libROM)
- Ultimate goal: Surgical planning/therapy optimization
 - Identification of optimality criteria
 - ROM techniques to accelerate identification of optimal solution
 - Ex-vivo and Pre-clinical Validation



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Scan me!

Thank you!

