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Bidomain Model Coupled To A Realistic Multi-Electrode Device In Cardiac Electrophysiology

SimCardioTest European Project

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WP2 of SimCardioTest focuses on Pacing Leads and Catheters, especially on **energy required to trigger a cardiac beat** with a pacemaker.



(a) BOREA[™]pacemaker



(b) Lead implanted in the septum



(c) Simulation of electrical field



Electrophysiological propagation models already exist, but we need to describe the stimulation

- Three discontinuous parts by stimulation ⇒ Device circuit must be modeled Structural analysis of electric circuits and consequences for MNA,
 D. Estévez Schwarz, C. Tischendorf, 2000
- Presence of a reactive layer at electrode boundaries ⇒ Layer must be modeled too Existence and Uniqueness for Electrode Models for Electric Current Computed Tomography, E. Somersalo, M. Cheney and D. Isaacson, 1992



(a) Geometric setup



(b) BOREA[™]

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+ Contacts +



Principle & Notations - I

- An oriented graph $\mathcal{G} = (S, B)$ with
 - $N_{
 m n}$ nodes,
 - $N_{\rm b}$ branches (dipoles)
- Potentials U in Volt and currents J in Ampere
- Incidence matrix $A = (a_{s,b})_{s \in \mathcal{S}, b \in \mathcal{B}}$ of size $N_{ ext{n}} imes N_{ ext{b}}$

$$a_{s,b} = \left\{ egin{array}{cc} -1 & ext{if s is an end point of b} \\ +1 & ext{if s is a start point of b} \\ 0 & ext{elsewhere} \end{array}
ight.$$

• Kirchhoff's current law AJ = 0 and voltages $V := A^{\mathsf{T}}U$.





Principle & Notations - II

- Only contains capacitors, resistors and voltage sources
- Split graph and matrix accordingly: $B = B_c \cup B_g \cup B_s$ and $A = [A_c, A_g, A_s]$

•
$$AJ = 0 \Rightarrow A_c J_c + A_g J_g + A_s J_s = 0$$

•
$$V = A^{\mathsf{T}}U \Rightarrow V_{\mathsf{c}} = A^{\mathsf{T}}_{\mathsf{c}}U, V_{\mathsf{g}} = A^{\mathsf{T}}_{\mathsf{g}}U$$
 and $V_{\mathsf{s}}(t) = A^{\mathsf{T}}_{\mathsf{s}}U$

- Dipole laws: $J_c = C \frac{dV_c}{dt} = C \frac{dA_c^{\mathsf{T}}U}{dt}$ and $J_g = GV_g = GA_g^{\mathsf{T}}U$
- Differential algebraic equations (DAE)

$$(U, J_{\rm s}) \quad \text{ solution of } \quad \begin{cases} A_{\rm c} C \frac{dA_{\rm c}^{\mathsf{T}} U}{dt} + A_{\rm g} G A_{\rm g}^{\mathsf{T}} U + A_{\rm s} J_{\rm s} = 0 \\ A_{\rm s}^{\mathsf{T}} U = V_{\rm s} \end{cases}$$





- Graph (S, B) oriented with L + 1 specialized nodes
- Potentials split as $U=(U^\circ,\gamma U)$ of size $N_{
 m n}$, internal potentials U°
- Dirichlet boundary condition, $\gamma_0 U = 0$ (ground)
- Coupling boundary potentials $\gamma_1 U, \ldots, \gamma_L U$
- Coupling output currents $\gamma J = (\gamma_1 J, \dots, \gamma_L J)$



solution of DAE
$$\begin{cases} A_c C \frac{dA_c^{\intercal} U}{dt} + A_g G A_g^{\intercal} U + A_s J_s + \gamma A \gamma J = 0 \\ A_s^{\intercal} U = V_s \end{cases}$$

with $\gamma \mathbf{A} \coloneqq [0, \mathsf{Id}_L]^{\mathsf{T}}$.

 (U, J_s)



Generic Models Equivalent Models for each electrode

On each electrode $\ell = 1, \dots, L$ (without voltage sources)

- Defined spatially: $\forall x \in \Gamma_{\ell}$
- Potentials split as $u_\ell = (u_\ell^\circ, \gamma u_\ell)$ of size N_ℓ
- Coupling boundary potentials $\gamma_1 u_\ell, \gamma_2 u_\ell$
- Coupling output currents $\gamma j_\ell = (\gamma_1 j_\ell, \gamma_2 j_\ell)$



solution of *x*-ODE
$$\boxed{A_{\ell,c}C_{\ell}\frac{\partial A_{\ell,c}^{\mathsf{T}}u_{\ell}}{\partial t} + A_{\ell,g}G_{\ell}A_{\ell,g}^{\mathsf{T}}u_{\ell} + \gamma A_{\ell}\gamma j_{\ell} = 0} \qquad \text{on } \Gamma_{\ell}$$

with $\gamma \mathbf{A}_{\ell} \coloneqq [0, \mathsf{Id}_2]^{\intercal}$.

 u_{ℓ}



Generic Models Bidomain Model (PDE)

- Tissue Ω_{M} intracellular and extracellular
- Bath/Blood $\Omega_{\rm B}$ passive media
- Transmembrane voltage $[u] = u_{\rm i} u$
- Conductivity tensors σ and σ_i
- L electrodes at boundary $\bigcup_{\ell=1}^{L} \Gamma_{\ell} \subset \partial \left(\overline{\Omega_{M}} \cup \overline{\Omega_{B}} \right)$

•
$$\gamma_\ell u = u_{|\Gamma_\ell}$$
, and $-\sigma
abla u \cdot \mathbf{n} = j_{\Gamma_\ell}$ on Γ_ℓ





$$(u, u_{i}) \text{ solution of PDE} \begin{vmatrix} -\operatorname{div}\left(\sigma_{i}\nabla u_{i}\right) = -j_{m} \text{ in } \Omega_{M} \text{ and } \begin{cases} -\operatorname{div}\left(\sigma\nabla u\right) = j_{m} & \operatorname{in} \Omega_{M} \\ -\operatorname{div}\left(\sigma\nabla u\right) = 0 & \operatorname{in} \Omega_{B} \end{cases}$$

with membrane current density $j_{\mathrm{m}} = \chi \left(c_{\mathrm{m}} \partial_t \left[u \right] + I_{\mathrm{ion}}(\left[u
ight], h, t)
ight)$, and lonic Model on h.





 $\begin{array}{l} - \text{ Potentials} \\ \gamma_\ell U = \gamma_1 u_\ell(\mathbf{x}) \text{, for } \mathbf{x} \in \Gamma_\ell \\ \gamma_2 u_\ell(\mathbf{x}) = \gamma_\ell u(\mathbf{x}) \text{, for } \mathbf{x} \in \Gamma_\ell \end{array}$

 $\begin{aligned} &- \text{Currents} \\ &\gamma_\ell J + \int_{\Gamma_\ell} \gamma_1 j_\ell(\mathbf{x}) = 0 \\ &\gamma_2 j_\ell(\mathbf{x}) + j_{\Gamma_\ell}(\mathbf{x}) = 0 \text{, for } \mathbf{x} \in \Gamma_\ell \end{aligned}$



Sketch of the proof - I

Global variational formulation

For all $\phi = (\phi, \phi_i, \phi_1, \dots, \phi_L, \Phi) \in \mathcal{C}_c^{\infty}$ of $[0, T) \times \left\{\overline{\Omega}, \overline{\Omega_M}, \left(\overline{\Gamma_\ell}\right)^{N_\ell}, \mathbb{R}^{N_n}\right\}$, find $\boldsymbol{u} \coloneqq (\boldsymbol{u}, \boldsymbol{u}_i, \boldsymbol{u}_1, \dots, \boldsymbol{u}_L, \boldsymbol{U})$ such that $\gamma_0 \boldsymbol{U} = 0$ + transmission conditions on potentials, and J_s , verifying

$$\begin{split} \int_{0}^{T} \int_{\Omega_{\mathsf{M}}} \sigma_{\mathsf{i}} \nabla u_{\mathsf{i}} \cdot \nabla \phi_{\mathsf{i}} + \int_{0}^{T} \int_{\Omega} \sigma \nabla u \cdot \nabla \phi + \int_{0}^{T} \int_{\Omega_{\mathsf{M}}} \left(c_{\mathsf{m}} \frac{\partial \left[u \right]}{\partial t} \left[\phi \right] + f\left(\left[u \right], h, t \right) \left[\phi \right] \right) \\ &+ \int_{0}^{T} \left(C \frac{\mathrm{d} \mathbf{A}_{\mathsf{c}}^{\mathsf{T}} U}{\mathrm{d} t} \cdot \mathbf{A}_{\mathsf{c}}^{\mathsf{T}} \Phi + G \mathbf{A}_{\mathsf{g}}^{\mathsf{T}} U \cdot \mathbf{A}_{\mathsf{g}}^{\mathsf{T}} \Phi + \mathbf{A}_{\mathsf{s}} J_{\mathsf{s}} \cdot \Phi \right) \\ &+ \sum_{\ell=1}^{L} \int_{0}^{T} \int_{\Gamma_{\ell}} \left(C_{\ell} \frac{\partial \mathbf{A}_{\ell,\mathsf{c}}^{\mathsf{T}} u_{\ell}}{\partial t} \cdot \mathbf{A}_{\ell,\mathsf{c}}^{\mathsf{T}} \phi_{\ell} + G_{\ell} \mathbf{A}_{\ell,\mathsf{g}}^{\mathsf{T}} u_{\ell} \cdot \mathbf{A}_{\ell,\mathsf{g}}^{\mathsf{T}} \phi_{\ell} \right) = 0, \\ \mathbf{A}_{\mathsf{s}}^{\mathsf{T}} U = \mathbf{V}_{\mathsf{s}} \end{split}$$

+ IPP \Rightarrow weak solution



Sketch of the proof – II

Global variational formulation semi-discretized in time

Find $m{u}^n\coloneqq (u^n,u^n_{\mathrm{i}},u^n_1,\ldots,u^n_L,U^n)\in\mathcal{H}_0$, and $J^n_s\in\mathbb{R}^{N_s}$ with

$$\begin{split} \mathcal{H} &\coloneqq H^1\left(\Omega_{\mathsf{M}}\right) \times H^1\left(\Omega\right) \times \Pi_{\ell=1}^L L^2\left(\Gamma_\ell\right)^{N_\ell} \times \mathbb{R}^{N_{\mathsf{n}}},\\ \mathcal{H}_0 &\coloneqq \{\boldsymbol{u} \in \mathcal{H} \text{ s.t. } \boxed{\gamma_0 \boldsymbol{U} = 0, \gamma_\ell \boldsymbol{U} = \gamma_1 \boldsymbol{u}_\ell, \gamma \boldsymbol{u} = \gamma_2 \boldsymbol{u}_\ell}, \ell = 1, \dots, L\}, \end{split}$$

verifying, for all $\pmb{\phi}\coloneqq(\phi,\phi_{\mathrm{i}},\phi_{1},\ldots,\phi_{L},\Phi)\in\mathcal{H}_{0}$,

$$\begin{split} a\left(\frac{\boldsymbol{u}^{n}-\boldsymbol{u}^{n-1}}{\Delta t},\boldsymbol{\phi}\right) + b\left(\boldsymbol{u}^{n},\boldsymbol{\phi}\right) + \left(f\left(\left[\boldsymbol{u}^{n}\right]\right),\left[\boldsymbol{\phi}\right]\right)_{\Omega_{\mathsf{M}}} + A_{\mathsf{s}}J_{\mathsf{s}}^{n}\cdot\boldsymbol{\Phi} = \boldsymbol{0},\\ A_{\mathsf{s}}^{\mathsf{T}}\boldsymbol{U}^{n} = V_{\mathsf{s}}^{n} \end{split}$$

$$\begin{split} a\left(\delta \boldsymbol{u},\boldsymbol{\phi}\right) &= \left(c_{\mathrm{m}}\left[\delta \boldsymbol{u}\right],\left[\boldsymbol{\phi}\right]\right)_{\Omega_{\mathsf{M}}} + \sum_{\ell=1}^{L} \left(C_{\ell}\boldsymbol{A}_{\ell,\mathrm{c}}^{\mathsf{T}}\delta\boldsymbol{u}_{\ell},\boldsymbol{A}_{\ell,\mathrm{c}}^{\mathsf{T}}\boldsymbol{\phi}_{\ell}\right)_{\Gamma_{\ell}} + C\boldsymbol{A}_{\mathrm{c}}^{\mathsf{T}}\delta\boldsymbol{U}\cdot\boldsymbol{A}_{\mathrm{c}}^{\mathsf{T}}\boldsymbol{\Phi},\\ b\left(\boldsymbol{u},\boldsymbol{\phi}\right) &= \left(\sigma_{\mathrm{i}}\nabla\boldsymbol{u}_{\mathrm{i}},\nabla\phi_{\mathrm{i}}\right)_{\Omega_{\mathsf{M}}} + \left(\sigma\nabla\boldsymbol{u},\nabla\phi\right)_{\Omega} + \sum_{\ell=1}^{L} \left(G_{\ell}\boldsymbol{A}_{\ell,\mathrm{g}}^{\mathsf{T}}\boldsymbol{u}_{\ell},\boldsymbol{A}_{\ell,\mathrm{g}}^{\mathsf{T}}\boldsymbol{\phi}_{\ell}\right)_{\Gamma_{\ell}} + G\boldsymbol{A}_{\mathrm{g}}^{\mathsf{T}}\boldsymbol{U}\cdot\boldsymbol{A}_{\mathrm{g}}^{\mathsf{T}}\boldsymbol{\Phi}. \end{split}$$



Sketch of the proof - III

Existence

Hypothesis

— Consequence A_s has full column rank $\ker(A_s) = \{0\}$ and $\operatorname{Im}(A_s^{\mathsf{T}}) = \ker(A_s)^{\perp} = \mathbb{R}^{N_s}, \Rightarrow \forall V_s^n \in \mathbb{R}^{N_s}$ there exists $U_0^n \in \mathbb{R}^{N_n}$ such as $A_s^{\mathsf{T}} U_0^n = V_s^n$ and $\|\boldsymbol{u}_0^n\|_{\mathcal{U}} \leq cst \|V_s^n\|_{\mathbb{R}^{N_n}}$.

 Variational formulation in the constraint space Find $\boldsymbol{u}_{\star}^{n} \in \mathcal{H}_{0}^{\mathrm{ker}}$, with $\boldsymbol{u}^{n} = \boldsymbol{u}_{\star}^{n} + \boldsymbol{u}_{0}^{n}$ and $\left| \mathcal{H}_{0}^{\mathrm{ker}} \coloneqq \left\{ \boldsymbol{u} \in \mathcal{H}_{0} \mid \boldsymbol{U} \in \mathrm{ker}\left(\boldsymbol{A}_{\mathrm{s}}^{\mathsf{T}}\right) \right\} \right|$, verifying for all $\phi_{\star} \in \mathcal{H}_{0}^{\mathrm{ker}},$ $a\left(\frac{\boldsymbol{u}_{\star}^{n}+\boldsymbol{u}_{0}^{n}-\boldsymbol{u}^{n-1}}{\Lambda \boldsymbol{t}},\boldsymbol{\phi}_{\star}\right)-b\left(\boldsymbol{u}_{\star}^{n}+\boldsymbol{u}_{0}^{n},\boldsymbol{\phi}_{\star}\right)+\left(f\left(\left[\boldsymbol{u}^{n}\right]\right),\left[\boldsymbol{\phi}\right]\right)_{\Omega_{\mathsf{M}}}=0$

Lax-Milgram and Banach fixed-point theorems: existence of $\boldsymbol{u}_{\perp}^{n}$ in $\mathcal{H}_{\circ}^{\mathrm{ker}}$.

- Resolution of I_s^n equation in \mathbb{R}^{N_s}
 - $\forall \{e_k\}$ basis of \mathbb{R}^{N_s} , $\exists \{\phi_k\} \subset \mathcal{H}_0$ s.t. $e_k = A_s^{\mathsf{T}} \Phi_k$, and J_s^n solves the linear systems $J_s^n \cdot e_k = w_k$,
 - w_k are independent of the choice of $\{\phi_k\}$.



We have a sequence (\boldsymbol{u}_0^n) bounded by data, and $(\boldsymbol{u}_{\star}^n)$ in $\mathcal{H}_0^{\text{ker}}$ with a priori estimates.

— A priori estimates on

$$a(\mathbf{u}^{n},\mathbf{u}^{n}) \qquad \sum_{n} \Delta t b(\mathbf{u}^{n},\mathbf{u}^{n}) \qquad \sum_{n} \Delta t a\left(\frac{\mathbf{u}^{n}-\mathbf{u}^{n-1}}{\Delta t},\frac{\mathbf{u}^{n}-\mathbf{u}^{n-1}}{\Delta t}\right)$$
for compactness (as *f* is nonlinear)

Compactness: discrete Aubin-Simon (Gallouët-Latché theorem)

 \Rightarrow up to subsequence $oldsymbol{u}^n
ightarrow oldsymbol{u}$ (strong convergence)

— Passing to the limit

 \Rightarrow *u* solution of the continuous problem.





Figure: Pacemaker + 2 R//C contacts + geometry inspired of a standard VEGA lead