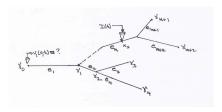
# Inverse Problems for Neuronal Cables on Graphs

Jon Bell and Sergei Avdonin

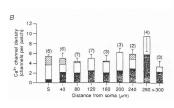
**UMBC** and University of Alaska Fairbanks

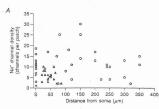
Minisymposium, 9th IPMS, 21-25 May 2018, Malta

## Motivation



Can we use boundary data (voltage & current measurements) to recover various distributed parameters in a dendritic tree model (quantum tree graph)?





# Single Branch, Single Conductance

- 1. JB, G Cracian, 2005: numerical method on
  - single unknown ion density:  $(1 + q(x))u_t + q(x)u = u_{xx}$ , 0 < x < L,  $u_x(0,t) = f(t)$ ,  $u_x(L,t) = 0$
  - unknown spine density in (nonlinear) Baer-Rinzel model:

$$u_t + u = u_{xx} + \rho n(x)(v - u)$$
  

$$v_t + i_{HH}(v, m, n, h) = \rho(u - v)$$

- D Wang, PhD thesis, UMBC, 2008: PDE-constrained optimization methods unknown spine density in Baer-Rinzel model
- 3. S Avdonin, JB, 2013, 2015 Boundary Control Theory on single unknown conductance  $u_t + g(x)u = u_{xx}$



### Dendritic Trees



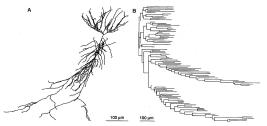
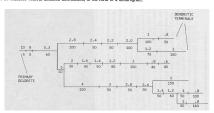


Fig. 3. A sample of representations available with the morphology editor as applied to the CAI cell n184. A: a simple skeleton projection. B: the appropriate lengths and connections of all branches without direction information, in the form of a dendrogram.

#### motoneuron.jpg



## Metric Graphs

Let  $\Gamma = \{E, V\}$  be a finite compact metric graph.

 $E = \{e_j\}_{j=1}^N$  is a set of edges and  $V = \{\nu_j\}_{j=1}^M$  is a set of vertices. A graph is called a metric graph if every edge  $e_j \in E$  is identified with an interval  $(a_{2j-1}, a_{2j})$  of the real line with a positive length  $l_j = |a_{2j-1} - a_{2j}|$ , and a graph is a tree if it has no cycles. The edges are connected at the vertices  $\nu_j$  which can be considered as equivalence classes of the edge end points  $\{a_j\}$ . Let  $\{\gamma_1, \ldots, \gamma_m\} = \partial \Gamma \subset V$  be the boundary vertices.

In this talk, graph = (finite compact) metric tree graph.

# A Simple Tree Example

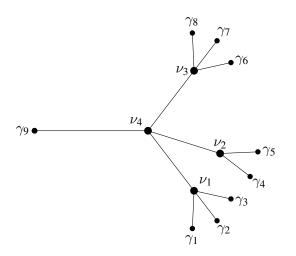


Fig. 1: A metric tree with m = 9 and N = 12

## Quantum Graphs

Quantum graph  $\{\Gamma, H\}$ : differential operator H on metric graph  $\Gamma$ , coupled by specific vertex matching conditions.

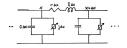
#### Applications include

- Oscillations of the flexible structures made of strings, beams, cables.
- hierarchical materials like ceramic or metallic foams, percolation networks and carbon and graphene nano-tubes.
- Our motivation comes from neurobiology; specifically dendritic trees of CNS neurons.

Control and inverse theories for PDEs on graphs constitute an important part of the rapidly developing area of applied mathematics — analysis on graphs.



## **Neuronal Cable Equation**



Single dendritic branch  $\Rightarrow$  Circuit model  $\Rightarrow$  Cable model Non-dimensionalized neuronal cable equation has the form

$$v_t + \sum_{j=1}^k g_j(x)(v - E_j) = v_{xx}$$

Pick any  $1 \le i \le k$ , and let  $u = v - E_i$ ,  $E_{ji} = E_j - E_i$ , and substitute into the equation:

$$u_t - u_{xx} + q(x)u = h(x)$$

where  $q(x) = \sum_{j=1}^k g_j(x)$ , and  $h(x) = \sum_{j \neq i} E_{ji} g_j(x)$ . The  $E_j s$  are assumed known, but the  $g_j s$ , hence q(x) and h(x), are assumed unknown a priori.

## IBVP on a Graph

Consider a system described by neuronal cable theory on a graph (adding p(t), with  $p(0) \neq 0$ ):

$$\partial_t u_j - \partial_x^2 u_j + q_j(x)u_j = p(t)h_j(x)$$
 on  $e_j \times (0,T) \ \forall e_j \in E$  (1)

or shortly,  $\partial_t u - \partial_x^2 u + q(x)u = p(t)h(x)$  on  $E \times (0,T)$ 

**KN**: 
$$\sum_{e_j \sim \nu} \partial u_j(\nu, t) = 0$$
 at each vertex  $\nu \in V \setminus \partial \Gamma$ ,  $u(\cdot, t)$  is continuous at each vertex, for  $t \in [0, T]$  (2)

$$\partial u = f$$
 on  $\partial \Gamma \times [0, T]$ ,  $u|_{t=0} = 0$  on  $E$ . (3)

In (2) (and below)  $\partial u_j(\nu)$  denotes the derivative of u at the vertex  $\nu$  taken along the edge  $e_j$  in the direction outwards the vertex. Also,  $e_j \sim \nu$  means edge  $e_j$  is incident to vertex  $\nu$ , and the sum is taken over all edges incident to  $\nu$ .

#### **IBVP** and Inverse Problem

Let  $\mathcal{H} = L^2(\Gamma)$  and  $\mathcal{F}^T = L^2([0,T];\mathbb{R}^m)$ .

**Theorem 1.** If  $f, p \in \mathcal{F}^T$ ,  $h \in \mathcal{H}$ , then for any  $t \in [0, T]$ ,  $u^f(\cdot, t) \in \mathcal{H}$  and  $u^f \in C([0, T]; \mathcal{H})$ .

For the inverse problem purposes we assume that  $p \in H^1(0,T), p(0) \neq 0$ .

The response operator,  $R^T: \mathcal{F}^T \to \mathcal{F}^T$ , is defined as

$$(R^T f)(t) = u^f(\cdot, t)|_{\partial \Gamma}, \ t \in [0, T].$$

**Theorem 2.** Operator  $R^T$  known for any T > 0 uniquely determines the graph topology, the lengths of the edges, the potentials  $q_j$  and sources  $h_j$ , j = 1, ..., N.



# Separating Problem into Two Auxiliary Problems

Solution of the problem (1)–(3) can be presented in the form u = y + z:

$$\partial_t y - \partial_x^2 y + q(x)y = 0 \text{ on } E \times (0, T)$$
 (4)

$$\partial y = f \quad \text{on} \quad \partial \Gamma \times [0, T]$$
 (5)

$$\partial_t z - \partial_x^2 z + q(x)z = p(t)h(x) \text{ on } E \times (0,T)$$
 (6)

$$\partial z = 0 \quad \text{on} \quad \partial \Gamma \times [0, T]$$
 (7)

with the KN matching conditions and zero initial conditions. We see that  $z=u^0$  so,  $y^f|_{\partial\Gamma}=u^f|_{\partial\Gamma}-u^0|_{\partial\Gamma}=R^Tf-R^T0$ .



# Solving Problem 1: Main Ideas

$$\mathcal{L}\phi \doteq -\phi'' + q(x)\phi = \lambda\phi \text{ on } \Gamma \setminus V, \text{ KN at } V \setminus \partial\Gamma, \phi'(\gamma_j, \lambda) = f_j,$$
  $f = col(f_1, \ldots, f_m), \rightarrow \exists ! \phi = \phi^f$ 

- 1. BC method allows spectral data SD= $\{\lambda_n,\phi_n^f|_{\partial\Gamma}\}_{n\in\mathbb{N}}$  to be recovered from  $R^T$
- 2. SD determines TW matrix function M:  $\phi^f_{|_{\partial\Gamma}} = M(\lambda)f$
- 3. TW matrix function M used in reduction method Essence: recalculate M from original graph to smaller graph by 'pruning' the boundary edges. By iterative process we reduce original IP (Prob. 1) to IP for **companion wave problem** on single edge.

$$w_{tt} - w_{xx} + q(x)w = 0 \text{ on } e = (0, \ell), \text{ zero i.c.s,} \ w_x(0, t) = f(t), w_x(\ell, t) = 0, \ f \in \mathcal{H}^T \doteq \{f \in L^2(0, \ell) \mid supp f \subset [0, T]\}$$



#### Problem 1 continued

$$\begin{split} F(t) &= \int_0^t f(s) ds \to \\ w^f(x,t) &= -F(t-x) + \int_x^t \kappa(x,s) F(t-s) ds, \, x < t, \, w^f(x,t) = 0, x \ge t \\ \kappa(x,t) &= \text{sol. to a certain Goursat prob. on } 0 < x < t < T \le \ell, \\ r(t) &= \kappa(0,t) \\ \psi'' - q(x)\psi &= 0, \, \psi(0) = 1, \psi'(0) = 0, \quad q \in L^1(0,\ell) \\ \exists f = f^T \in L^2(0,T) \text{ s.t. } w_t^{f^T}(x,T) = \psi(x), \, x \le T, = 0, x > T \\ f^T(t) - \frac{1}{2} \int_0^T \{r(|t-s|) + r(2T-t-s)\} f(s) ds = -1 \quad t \in [0,T] \\ w_t^{f^T}(T-,T) &= -f^T(0+) = \psi(T) \in C^2 \Rightarrow q(T) = \frac{\psi''(T)}{s^{b}(T)} \end{split}$$

See Avdonin and Bell, J. Inv. Prob. and Imaging, 9(3)(2015)



# Solving Problem 2

$$z(x,t) = \sum_{n=1}^{\infty} \varphi_n(x) h_n \int_0^t p(t-\tau) e^{-\lambda_n \tau} d\tau ,$$

where  $h_n=\langle h,\varphi_n\rangle_{\mathcal{H}}$ . Recalling  $p(0)\neq 0$ ,  $\mu(t)\doteq u^0(\cdot,t)|_{\partial\Gamma}$ 

$$\mu(t) = \int_0^t p(t-\tau)W(\tau) d\tau, \quad W(\tau) \doteq \sum_{n=1}^\infty h_n \varphi_n(0) e^{-\lambda_n \tau},$$

$$\mu'(t) = p(0) W(t) + \int_0^t p'(t-\tau)W(\tau) d\tau.$$

The last equation allows finding W(t) as the solution to the VIE.



#### Problem 2 continued

**Theorem 3** (controllability): implies that the family  $\{\varphi_n(0)e^{-\lambda_n t}\}$  is minimal in  $\mathcal{F}^T=L^2([0,T];\mathbb{R}^m)$  (for any T>0) with biorthogonal family  $\{\theta_n(t)\}$ .

Thus, with  $W(\cdot)$  and SD determined, we then have the  $h_ns$  and h:

$$h_n = \langle W, \theta_n \rangle_{\mathcal{F}^T}, \quad h(x) = \sum_{n=1}^{\infty} h_n \varphi_n(x).$$

Recovery of original distributed parameters requires solving Problem 1 once, solving Problem 2 k-1 times.

See Avdonin, Bell, and Nurtazina, Math. Meth. Appl. Sci. 40(2017)



#### Other Problems

#### Scaled (linear) models:

- One unknown ion species:  $(1 + q(x))u_t + q(x)u = u_{xx}$  $(C_m = C_0 + C_1N(x), g = g_1N(x))$
- Unknown radius, known conductance:  $u_t + q(x)u = \frac{1}{a(x)}(a(x)^2u_x)_x$

First on single interval, then on a graph.



## Next: Graphs with Cycles

cyclomatic number (first Betti number) = |E|-|V|+1(>0) Much more challenging extending control theory than for tree graphs

