Statistical challenges and opportunities for reliable CNS interfaces

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A different perspective: optimal decoding of retinal spike train data

Preparation: macaque retina in vitro (Litke et al '04)

— extracellularly-recorded responses of populations of ganglion cells





Decoding the spatiotemporally-filtered stimulus via fast Bayesian methods



(Ahmadian et al '11); note: motor decoders provide much lower bandwidth.

Outline

- The state of the art: state-space models
- Fast estimation of nonlinear, nonstationary encoding models
- Robust subspace identification
- New methods for optimal Bayesian decoding: fast approximate Kalman-based methods and sequential Markov chain Monte Carlo
- Non-myopic optimal experimental design
- Using all the available information: fast approximate methods for hierarchical regularized models
- Modeling and exploiting co-adaptation.

State-space models

Some notation:

- $x_t =$ kinematic state
- $r_t =$ neural response
- encoding model: $p(r_t|x_t)$
- kinematics model: $p(x_t|x_{t-1})$
- optimal Bayesian decoder: $p(x_t|r_{1:t})$

This approach is flexible, computationally efficient (because the decoder is computed recursively at each time step t), and currently provides the best available performance.

Challenge: estimating the encoder $p(r_t|x_t)$

— Simple, near-linear models have sufficed so far in 2d planar hand tracking studies; no longer true for > 20-DOF movements (see, e.g., Vargas-Irwin et al '10)

 Nonstationarity is a key, unavoidable issue: the subject adapts to the controller as the controller adapts to the subject.
Re-estimating parameters every once in a while is a suboptimal solution.

— Speed is essential, since we have to estimate an encoding model for each observed neural channel.

— Reliability requirements translate into a need to avoid local minima in parameter searches: convex optimization is key tool

New methods for tractably estimating nonlinear models

Simplest example: higher-order terms (e.g., quadratic; Li '09):

$$E(r_t|x_t) = b + k^T x_t + x_t^T A x_t.$$

Problem: number of parameters explodes as we add more terms. However, we can exploit new convex optimization ideas from the machine learning literature (low-rank tensor completion) to avoid overfitting and obtain quite robust and accurate estimates:











New methods for fast optimal Bayesian nonstationarity tracking

A flexible framework: generalized additive models, $E(r_t) = F\left[\sum_{i=1}^d a_i(t)g_i(x_t)\right]$.

As $a_i(t)$ changes, so do the response properties.

Goal: track the a_i 's given limited, noisy data.

Fast, robust methods for tracking nonstationarities (optimal Bayesian inference requires just O(dT) time; Paninski et al, '11)



Another challenge: latent variable models

$$E(r_t) = F(x_t, z_t)$$

 z_t = latent state variable whose dynamics are restricted to a low-dimensional subspace. Captures the fact that firing rates are modulated by many variables that we don't observe directly (i.e., not just x_t).

Similar models proposed by Yu, Sahani, Shenoy et al and Wu, Paninski et al: significant improvements over models with no z_t term

Problem: estimation of these models is more challenging, since z_t is never observed.

Subspace identification

Previous work applied iterative methods (expectation-maximization) for estimating the latent model parameters.

These methods are slow and non-robust: many iterations, prone to local optima.

Idea: borrow methods from control literature (e.g., Liu and Vandenberghe '09). Use matrix completion again to identify the subspace using limited, noisy data.

Convex problem: no local optima.

Robust decoding

Once model is identified, how to decode?

If $p(x_t|r_{1:t})$ is unimodal, then Kalman-based methods suffice. we can exploit our new fast O(d) Kalman methods here as well. If $p(x_t|x_{t-1})$ or $p(r_t|x_t)$ is strongly multimodal, Monte Carlo methods are necessary.

"Particle filtering" is the most common approach; in principle, these methods are very general. However, in practice they are often slow and very non-robust.

Speed issues are solvable: method is embarrassingly parallel at each timestep t, and could be implemented on a GPU.

Robustness issues are more fundamental: standard methods put particles in the wrong place in many cases

Sequential MCMC methods are much more robust than the particle filter



0.1 times the weights



100 times the weights:



(Vidne and Paninski '11)

Optimal experimental design

Idea: choose test movements to best constrain the model parameters (simple example: Cunningham et al '08).

Previous approaches: choose one test movement at a time, in a greedy (myopic) way (Mackay, '92).

New approach: exploit connection to classical problem in information theory to compute the globally optimal sequence: no greedy local optimization required.



Hierarchical models

We record from many units simultaneously, but typically estimate encoding models one unit at a time: this is suboptimal.



(Field et al, Nature '10; Sadeghi et al, in preparation)

Modeling and exploiting co-adaptation

A number of intriguing results, but no good quantitative models (to my knowledge).

State-space ideas provide a possible starting point.

Simplest case: linear filtering: $\hat{x}_t = \theta_t^T r_t$

Subspace model: $E(r_t) = Ku_t$: subject can only influence r_t within a subspace K of dim = DOF.

Co-adaptation: experimentalist optimizes θ_t to optimize accuracy; subject tries to infer θ_t based on recent history, in order to best choose the control signal u_t .

Modeling and exploiting co-adaptation

Natural state-space model: subject tracks mapping $\hat{\theta}_t$ (with uncertainty) via Bayesian updates. Similar models in motor psychophysics literature on adaptation of sensorimotor maps.

Goal: exploit co-adaptation instead of fighting it:

- natural hierarchical model for tracking encoding models
- improved decoder: incorporate subject's uncertainty about θ

— connection to optimal design: the subject will try control signals u_t that balance exploration (to optimize information about θ) and minimization of error. Qualitatively consistent with nonstationarities during tracking (e.g., results from Carmena, Schwartz labs).

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