Modeling Children's Activity Levels using Wavelet-Based Functional Mixed Models Jeffrey S. Morris University of Texas M.D. Anderson Cancer Center, Houston, TX

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Overview

Introduction

- Planet Health Study/Accelerometer Data
- Mixed Models/Wavelets
- Wavelet-based Functional Mixed Models
- Missing Data Methods for WFMM
- Analysis of Accelerometer Data from Planet Health Study
- Discussion/Conclusions

Planet Health Study

• Planet Health: school-based intervention study to reduce obesity in middle school youth by changing key physical activity and dietary risk factors.

• Design:

- 10 schools paired up, randomized to intervention/control
- 1295 children: nutritional, behavioral, and health-related outcomes measured at baseline (F95) and follow-up (S97)

Challenge: How to measure physical activity levels?

- Questionnaires
- Accelerometers: objectively quantify activity levels

 256 children: monitored using TriTrac-R3D activity monitor for one or two 4-day sessions in S97 8/4/2005

Accelerometer Data

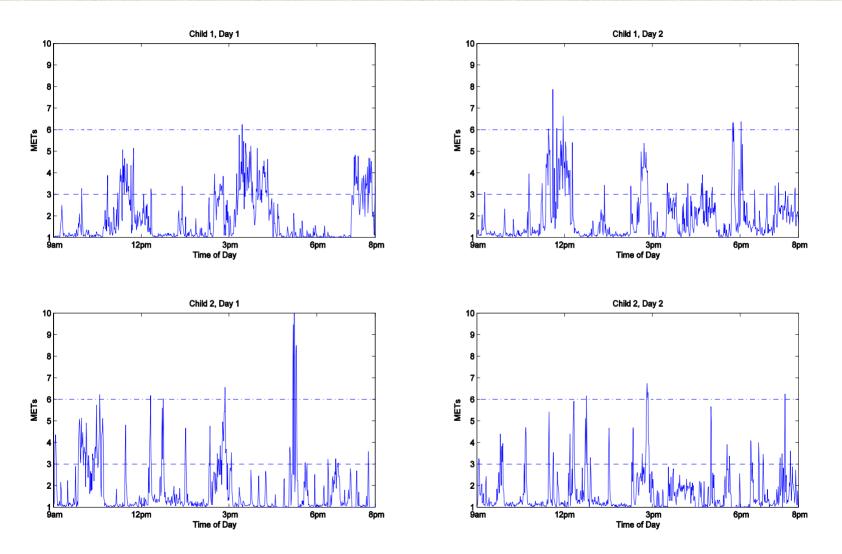
- Accelerometers: small motion sensors that digitally record minute-by-minute activity levels

 Increasingly used in surveillance and intervention studies
- TriTrac-R3D: pocket-sized sensor worn on hip
 - Minute-by-minute record of motion in 3 planes
 - Condensed into single activity level measurement/minute, on either acceleration or "METs" scale (multiple of resting metabolic rate)

• METs for different activities (Rowlands, et al. 2004)

- 1 MET = minimum activity level
- 3-6 MET = moderately intense activity
- >6 MET = vigorous activity

Accelerometer Data



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Analyzing Accelerometer Data

- Major challenge: How to analyze these data?
- Standard analysis strategies involve simple, arbitrary summary measures:
 - Average daily activity levels
 - Proportion of time with activity levels corresponding to sedentary, moderately vigorous, and vigorous activities
 - 30-minute averages
- Limitations of these approaches:
 - Do not make full use of information in functional data
 - Cannot effectively deal with incomplete profiles
- Methods that model the entire functions may be more effective in extracting information from these data

Accelerometer Data

- Here, we focus only on weekday profiles from children at control schools

 550 profiles from 112 children, 1440 meas./profile
 Enormous amount of data; can be displayed using
- Lots of missing data
 - Use profiles >50% complete from 9am-8pm

Heatmap

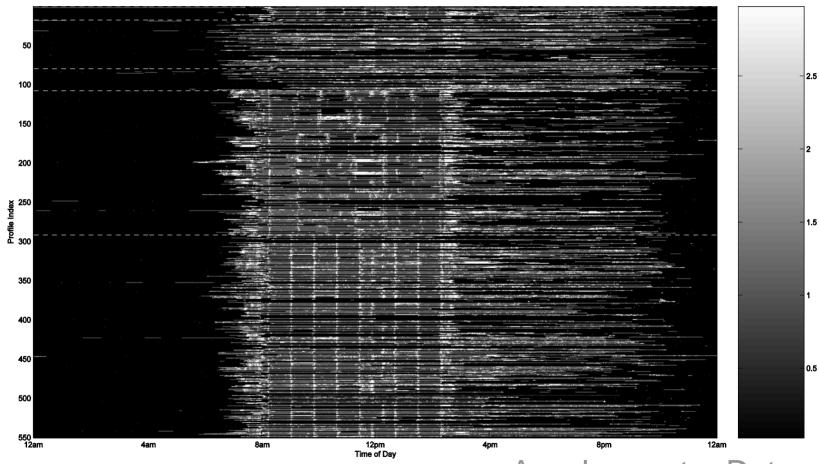
• 292 profiles from 106 children

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Heatmap



Accelerometer Data

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Accelerometer Data

Other measurements:

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- Child-level covariates: school, race, gender, age, weight, height, BMI, % body fat, avg hrs of TV/week
- Day-level covariates: day of week, calendar date
 Goals:
 - 1. Assess how activity levels vary throughout day, across schools, across different days of the week, over time from early to late Spring, and across various child-level covariates.
- 2. Assess relative variability in activity levels from day-to-day and child-to-child, in order to guide future study design.

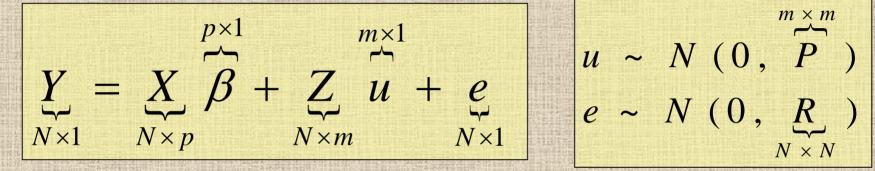
Statistical Challenges

- In order to model these data as functions, we need a method that can:
- 1. Model functions of arbitrary form
- 2. Jointly model *functional effects* of multiple covariates
- 3. Account for correlation between profiles from same child
- 4. Incorporate information from incomplete profiles

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Linear Mixed Models

Linear Mixed Model (Laird and Ware, 1982):



Fixed effects part, Xβ, accommodate a broad class of mean structures, including main effects, interactions, and linear coefficients.
 Random effects part, Zu, provide a convenient mechanism for modeling correlation among the N observations.

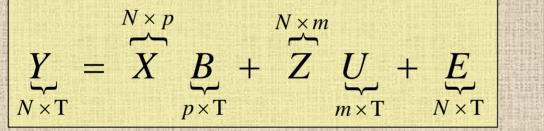
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Functional Mixed Model Morris and Carroll (2004) (Discrete version)

Y= *N*-by-*T* matrix containing the observed curves all sampled on a common equally-spaced grid of length *T*, *t*.



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 $\begin{bmatrix} U \sim MN(P,Q) \\ E \sim MN(R,S) \end{bmatrix}$

• B_{ij} is the effect of covariate *i* at location t_j

- U and E follow the Matrix Normal distribution
 - $U \sim MN(P,Q)$ implies $Cov\{U_{ij}, U_{i'j'}\}=P_{ii'}*Q_{jj'}$
- **P** and **R** are between-curve covariance matrices

• Q and S are within-curve covariance matrices $(T \times T)$

Example: Model Let Y be 292×660 matrix containing 292 accelerometer profiles for each minute from 9am-8pm.

Y = XB + ZU + E

X = 292 × 14 matrix of covariates

 School effects (5), gender, % body fat, BMI, day-of-week (4), daylight savings time, avg tv hrs/wk

• $B = 14 \times 660$ matrix of fixed effects functions

 $-B_{ij}$ is effect of covariate *i* at time t_j

Z = 292 × 106 matrix indicating child for each profile
 U = 106 × 660 matrix of random effect functions (1/child)

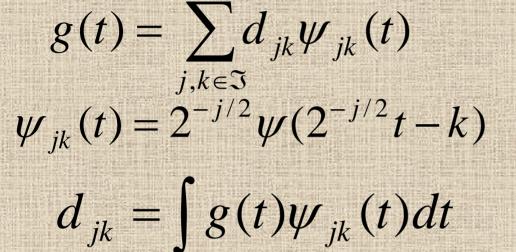
• E = 292 × 660 matrix of residual errors

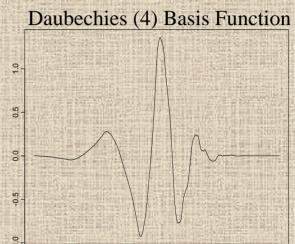
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Functional Mixed Models

- **Key feature of FMM:** Does not require specification of parametric form for curves
- Most existing literature for nonparametrically modeling functional data is based on kernels or splines (Guo 2002).
- Kernels/fixed-knot splines may not work well for spatially heterogeneous data
- Wavelet Regression: nonparametric regression technique that better preserves local features present in the curves.

Introduction to Wavelets Wavelets: families of orthonormal basis functions





- Discrete Wavelet Transform (DWT): fast algorithm {O(T)} for obtaining T empirical wavelet coefficients for curves sampled on equally-spaced grid of length T.
- Linear Representation: d = y W'- W' = T-by-T orthogonal projection matrix
- Inverse DWT (IDWT): y = d8/4/2005 Harvard SPH November 20

Wavelet Regression

Wavelet Regression:

- Row vector y: response on equally-spaced grid t (length T)

$$\mathbf{y} = g(\mathbf{t}) + \varepsilon \qquad \varepsilon \sim N(0, \sigma^2 I_T)$$

Project data into wavelet space using DWT.
 d=y W' where W' is the orthogonal DWT matrix

Wavelet Regression

• Wavelet Regression:

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- Row vector y: response on equally-spaced grid t (length T)

 $\mathbf{d} = \mathbf{\theta} + \boldsymbol{\varepsilon}^* \qquad \boldsymbol{\varepsilon}^* \sim N(0, \sigma^2 I_T)$

1. **Project data into wavelet space using DWT.**

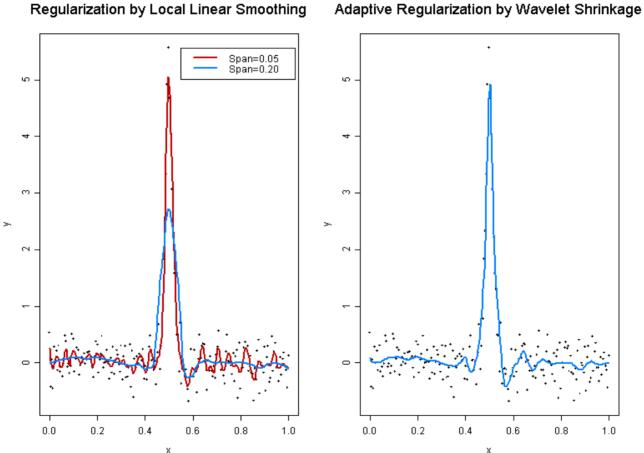
d=y W' where W' is the orthogonal DWT matrix

Estimate θ by thresholding (or placing mixture prior on θ)
 Project back to data space using IDWT

$$\hat{g}(t) = \hat{\theta} W$$

Yields *adaptive regularized* nonparametric estimate of g(t).

Adaptive Regularization



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Wavelet-based FMM:

- Morris, et al. (2003) introduced new Bayesian method generalizing wavelet regression to hierarchical functional framework
 - Bayesian inference on mean curve and random effect functions for nested functional data
- Morris and Carroll (2004) extended this work to functional mixed model
 - Arbitrary functional fixed and random effects
 - Wavelet-based functional mixed models

Wavelet-based FMM:

General Approach

1. Project observed functions Y into wavelet space. 2. Fit FMM in wavelet space. (Use MCMC to get posterior samples) **3. Project** wavelet-space estimates (posterior samples) back to data space.

Wavelet-based FMM:

General Approach

1. Project observed functions Y into wavelet space.

Fit FMM in wavelet space

 (Use MCMC to get posterior samples)

 Project wavelet-space estimates

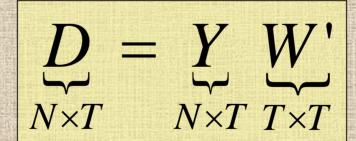
 (posterior samples) back to data space.

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Wavelet-based FMM

1. Project observed functions Y to wavelet space

• Apply DWT to rows of Y to get wavelet coefficients corresponding to each observed function



Projects the observed curves into the space spanned by the wavelet bases.

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Wavelet-based FMM: <u>General Approach</u>

1. Project observed functions Y into wavelet space.

2. Fit FMM in wavelet space (Use MCMC to get posterior samples)

3. Project wavelet-space estimates (posterior samples) back to data space.

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= X B + Z U + E $N \times p$ $N \times T$

 $U \sim MN(P,Q)$ $E \sim MN(R,S)$

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 $\overrightarrow{\mathbf{W'}} = \overrightarrow{X} \underbrace{B}_{p \times T} + \overrightarrow{Z} \underbrace{U}_{m \times T} + \underbrace{E}_{N \times T}$ $N \times T$

 $U \sim MN(P,Q)$ $E \sim MN(R, S)$

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 $\vec{T} = \vec{X} \quad \vec{B} \quad \vec{W'} + \vec{Z} \quad \vec{U} \quad \vec{W'} + \vec{E}$ $T \times T$ $T \times T$ $N \times T$ $p \times T$ $m \times T$

 $U \sim MN(P,Q)$ $E \sim MN(R, S)$

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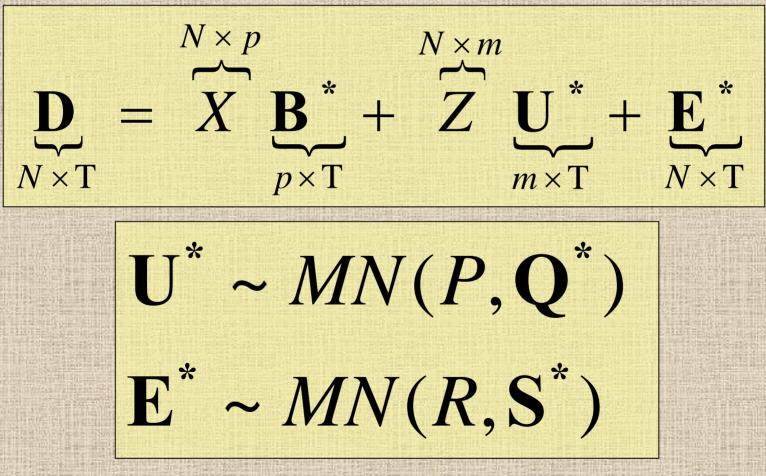
$\underbrace{Y}_{N\times T} \underbrace{\stackrel{N\times p}{\mathbf{W'}}}_{p\times T} = \underbrace{X}_{p\times T} \underbrace{B}_{p\times T} \underbrace{\stackrel{T\times T}{\mathbf{W'}}}_{p\times T} + \underbrace{Z}_{m\times T} \underbrace{U}_{m\times T} \underbrace{\stackrel{T\times T}{\mathbf{W'}}}_{N\times T} + \underbrace{E}_{N\times T} \underbrace{\stackrel{T\times T}{\mathbf{W'}}}_{N\times T}$

$UW' \sim MN(P, WQW')$ $EW' \sim MN(R, WSW')$

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Prior Assumptions

Mixture prior on B^{*}_{ijk}:

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$$B_{ijk}^* = \gamma_{ijk}^* N(0, \tau_{ij}) + (1 - \gamma_{ijk}^*) \delta_0$$

 $\gamma_{ijk}^* = \text{Bernoulli}(\pi_{ij})$

- Nonlinearly shrinks B_{ijk}^* towards 0, leading to adaptively regularized estimates of $B_i(t)$.
- τ_{ij} & π_{ij} are regularization parameters

 Can be estimated from the data using empirical Bayes
 Extend Clyde&George (1999) to functional mixed model

Model Fitting

 Let Ω be a vector containing ALL covariance parameters (i.e. for P, Q*, R, and S*).
 MCMC Steps

 Sample from f(B*/D,Ω): Mixture of normals and point masses at 0 for each i,j,k.
 Sample from f(Ω/D,B*): Metropolis-Hastings steps for each j,k
 If desired, sample from f(U*/D,B*,Ω): Multivariate normals

Wavelet-based FMM: General Approach

1. Project observed functions Y into wavelet space. 2. Fit FMM in wavelet space (Use MCMC to get posterior samples) **3. Project** wavelet-space estimates (posterior samples) back to data space.

Wavelet-based FMM

- 3. **Project** wavelet-space estimates (posterior samples) back to data space.
- Apply IDWT to posterior samples of *B** to get posterior samples of *B*

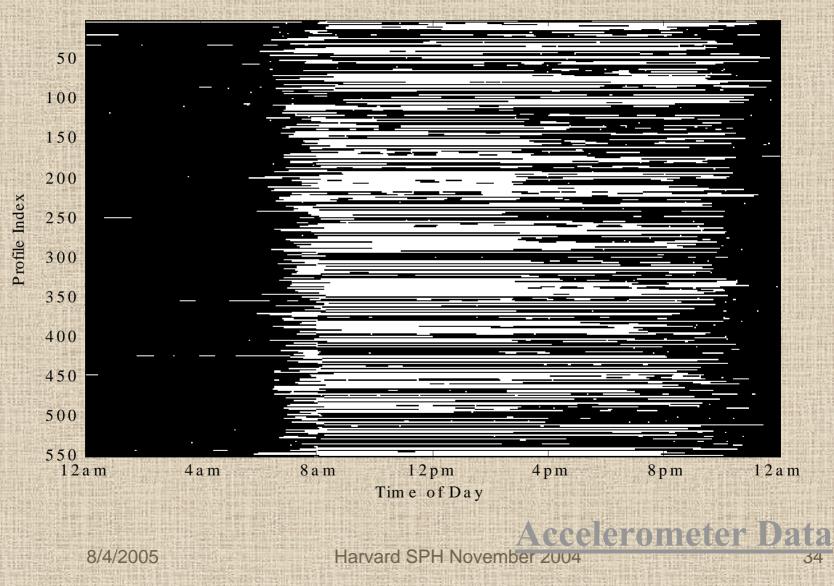
- **B=B*W**

- Also for U, P, Q, R, and S
- Can be used for Bayesian inference/prediction
 - Posterior probabilities
 - Bayes Factors
 - Posterior predictive probabilities

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Incomplete Profiles • Lots of missing data (Missing Data) **Example of** incomplete profile • WFMM can only be applied to *complete profiles* (with no missing regions) - 95 of the 292 profiles complete from 9am-8pm How do we incorporate information from other 197 incomplete profiles?

Missing Data



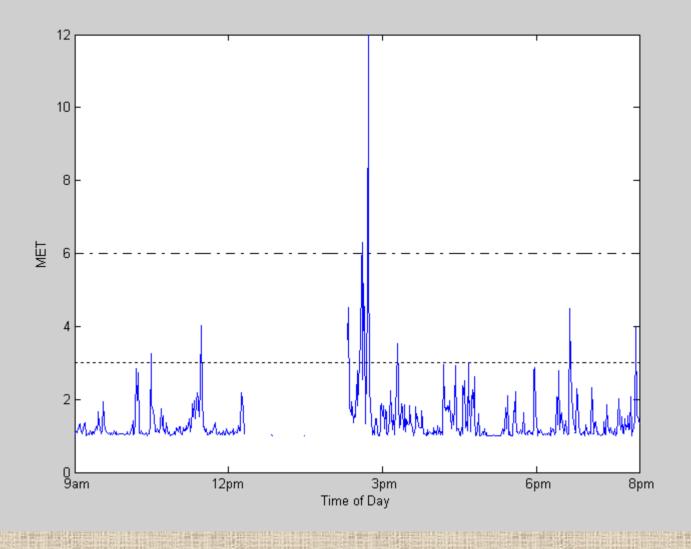
Approach: Incomplete Profiles

- 1. First fit model to *complete profiles*, get estimates and standard errors for model parameters.
- 2. Use these to estimate *predictive distributions* for the the incomplete profiles (fig)
 - Borrow information about what the curves in these regions look like.
 - Account for child-specific and day-specific covariates.
- 3. Regress missing data on the observed data to obtain *imputation distribution* for missing regions (fig)
 - Borrow information from nearby times in incomplete profiles.
 - Makes predictions for missing regions "connected" with observed.
- 4. Supplement WFMM with step to stochastically impute values for missing data.

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• Inference appropriately accounts for uncertainty in imputation

Incomplete Profile



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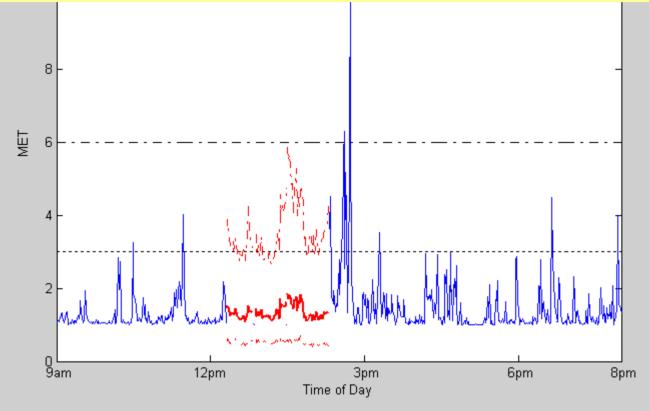
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Predictive Distribution

$\mu_{i}(t) = E\{Y_{i}(t) | Y^{C}\} = \int Y_{i}(t) f\{Y_{i}(t) | X, Z, \Theta\} f(\Theta | Y^{C}) d\Theta$ $\Sigma_{i}(t_{1}, t_{2}) = COV\{Y_{i}(t_{1}), Y_{i}(t_{2}) | Y^{C}\}$

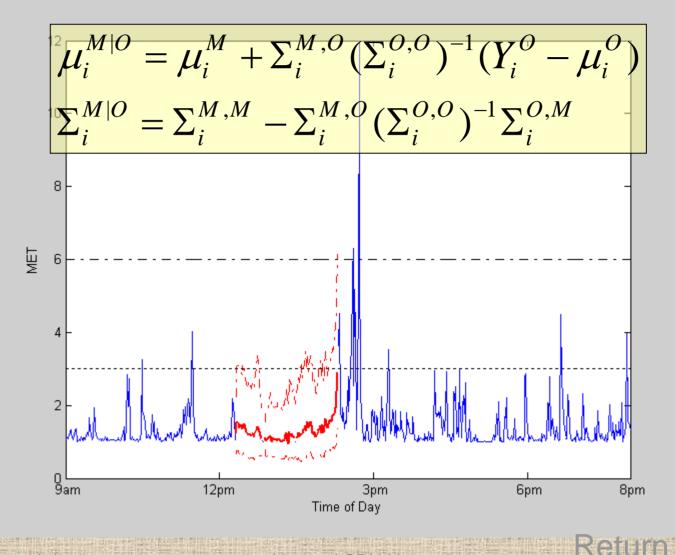


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Keti

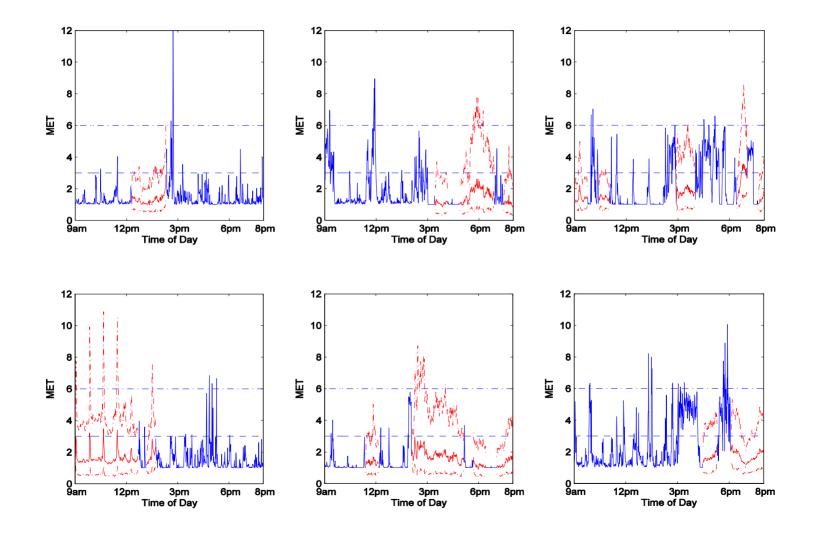
Imputation distribution



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Incomplete Profiles



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Missing Data in the WFMM

- **Problem:** Imputation distribution in data space, modeling done in wavelet space
- Solution: Project Imputation Distributions into wavelet space

 $M_{i}(t) = \begin{cases} Y_{i}(t) \text{ if } t \text{ observed} \\ \mu_{i}^{M|O}(t) \text{ otherwise} \end{cases}$

 $V_i(t_1,t_2) = \begin{cases} 0 \text{ if either } t_1 \text{ or } t_2 \text{ obs.} \\ \Sigma_i^{M|O}(t_1,t_2) \text{ otherwise} \end{cases}$

 $M_i^* = M_i W'$ $V_i^* = W V_i W'$

 Add step to MCMC whereby "missing" wavelet coefficients D_{ijk}~N(M*_{ijk}, V*_{ijk}) 8/4/2005
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Model

Let **Y** be 292×660 matrix containing 292 accelerometer profiles for each minute from 9am-8pm.

Y = XB + ZU + E

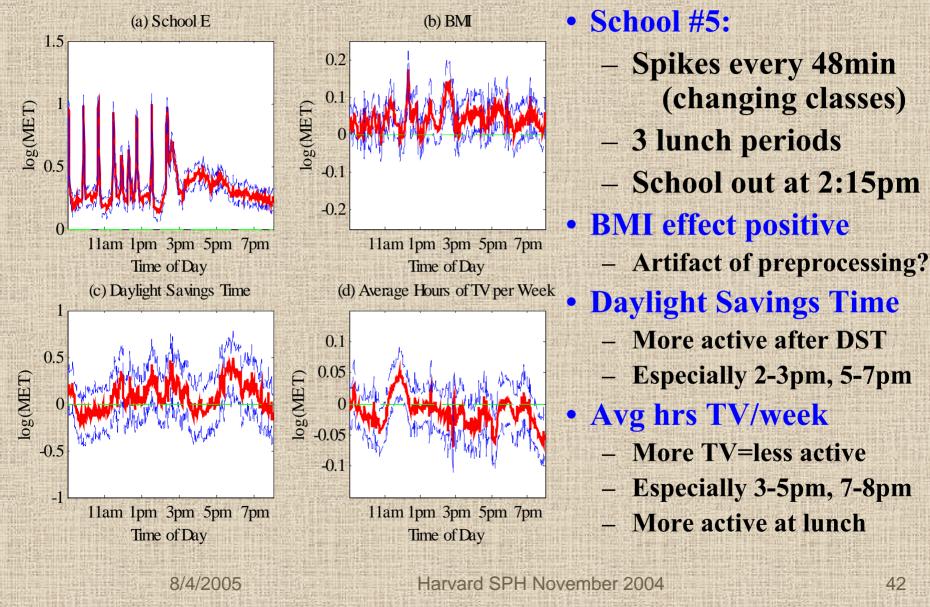
• X = 292 × 14 matrix of covariates

- School effects (5), gender, % body fat, BMI, day-ofweek (4), daylight savings time, avg tv hrs/wk
- $B = 14 \times 660$ matrix of fixed effects functions
 - $-B_{ij}$ is effect of covariate *i* at time t_j
- Z = 292 × 106 matrix indicating child for each profile
 U = 106 × 660 matrix of random effect functions (1/child)
- E = 292 × 660 matrix of residual errors

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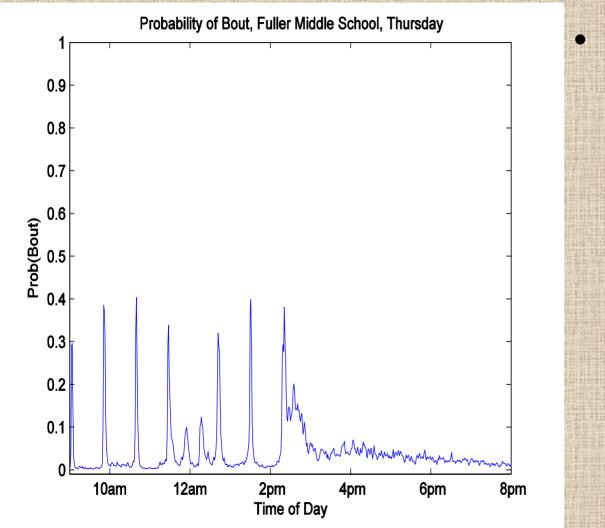
Some Results



Results: Covariance Analysis

- Variability: 91% day-to-day, 9% child-to-child
 - Important to have many days per child
- Study variability as function of t
 - Child-to-child variability: school day > after school
 - Day-to-day variability: after school > school day
 - Relative day-to-day variability after school: 95%-99%
- Equivalent designs:
 - 108 children, 4 days/child
 - 72 children, 8 days/child
 - 54 children, 16 days/child
- Less children, more days, save \$\$\$?

Results: Bouts



Can compute posterior predictive probabilities of bouts for children

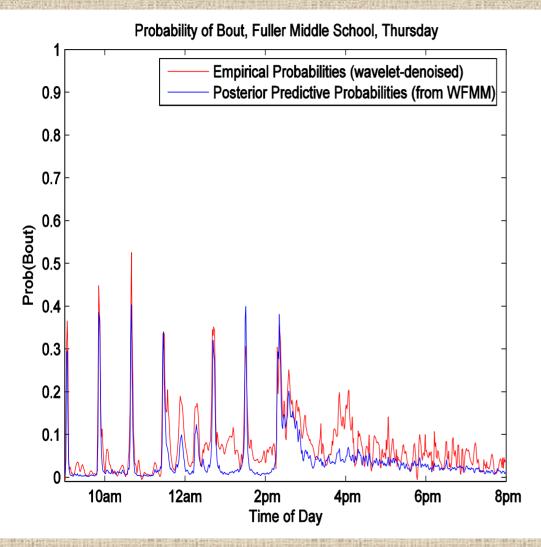
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Results: Bouts

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Can compute posterior predictive probabilities of bouts for children Model-based predictive probabilites not far from empiricallyestimated probabilities May want heavier tails

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Discussion

- WFMM: unified modeling approach for FDA
 - Extract information from samples of accelerometer profiles
- Method based on mixed models; is FLEXIBLE
 - Accommodates a wide range of experimental designs
 - Addresses large number of research questions
- Posterior samples allow Bayesian inference and prediction
 - Posterior credible intervals; pointwise or joint
 - Predictive distributions for future sampled curves
 - Predictive probabilities for group membership of new curves
 - Bayesian functional inference can be done via Bayes Factors
- Since a unified modeling approach is used, all sources of variability in the model propagated throughout inference.

Discussion

- Since functions adaptively regularized using wavelet shrinkage, the method is appropriate for spatially heterogeneous functional data, like accelerometer data.
- Approach is Bayesian. The only informative priors to elicit are regularization parameters, which can be estimated from data using empirical Bayes.
- Method complex, but code straightforward to implement.
- Method generalizes to higher dimensional functions, e.g. image data, space/time (fixed domain) data.
- We used wavelet bases, but approach can be generalized to other orthogonal basis functions.
- Unified modeling approach makes it possible to develop rigorous methods for incorporating of observed functions with missing data
 - Makes it possible to relax equally-spaced grid assumption
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