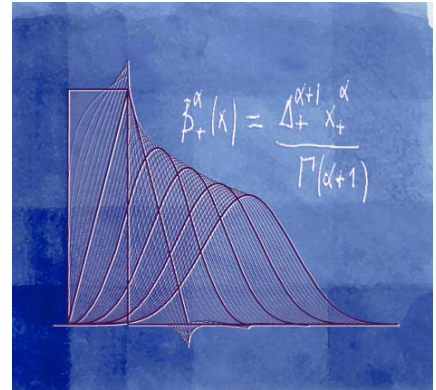


20 years of splines and biomedical imaging: The prehistory

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BIG's 20th Birthday, March 23, 2018, EPFL, Switzerland



1981: Master in EE

1984: Ph.D. in EE



1985-88: Visiting Fellow

National Institutes of health,
Biomedical Engineering and Instrumentation Program

1989-97: Scientist



1997-2000: Associate Professor



2000-present: Full Professor

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1985-89: Electron microscopy

Volume 55 April 1989 713-724 Biophysical Journal 713

Interactions between actin and myosin filaments in skeletal muscle visualized in frozen-hydrated thin sections

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ABSTRACT For the purpose of determining net interactions between actin and myosin filaments in muscle cells, perhaps the single most informative view of the myofibril lattice is its averaged axial projection. We have studied frozen-hydrated transverse thin sections with the goal of obtaining axial projections that are not subject to the limitations of conventional thin sectioning (suspect preservation of native structure) or of equatorial x-ray diffraction analysis (lack of experimental phases). In principle, good preservation of native structure may be achieved with fast freezing, followed by low-dose electron imaging of unstained vitrified cryosections. In practice, how-

ever, cryosections undergo large-scale distortions, including irreversible compression; furthermore, phase contrast imaging results in a nonlinear relationship between the projected density of the specimen and the optical density of the micrograph. To overcome these limitations, we have devised methods of image restoration and generalized correlation averaging, and applied them to cryosections of rabbit psoas fibers in both the relaxed and rigor states. Thus visualized, myosin filaments appear thicker than actin filaments by a much smaller margin than in conventional thin sections, and particularly so for rigor muscle. This may result from a significant

fraction of the myosin S1-cross-bridges averaging out in projection and thus contributing only to the baseline of projected density. Entering rigor incurs a loss of density from an annulus around the myosin filament, with a compensating accumulation of density around the actin filament. This redistribution of mass represents attachment of the fraction of cross-bridges that are visible above background. Myosin filaments in the "nonoverlap" zone appear to broaden on entering rigor, suggesting that, on deprivation of ATP, cross-bridges in situ move outwards even without actin in their immediate proximity.



Ultramicroscopy 30 (1989) 429-434 North-Holland, Amsterdam

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SHORT NOTE

THE SPECTRAL SIGNAL-TO-NOISE RATIO RESOLUTION CRITERION: COMPUTATIONAL EFFICIENCY AND STATISTICAL PRECISION

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Received 14 March 1989

This note describes a practical improvement in the computational efficiency of the spectral signal-to-noise ratio (SSNR) resolution criterion for covariation-averaged images. The total set of N images is randomly partitioned into n_p subsets, each subset is separately averaged, and a reduced form of the SSNR is computed from these average images. In general, larger values of n_p achieve lower statistical uncertainty, while smaller values of n_p are computationally more expedient. It is shown that, for negatively stained data, a judicious compromise is achieved with $10 \leq n_p \leq 20$, regardless of how large N may be.



Jacques Dubochet, Joachim Frank, Richard Henderson

Nobel Prize in Chemistry 2017

"for developing cryo-electron microscopy for the high-resolution structure determination of biomolecules in solution"

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1990-93: Splines and signal processing

IEEE TRANSACTIONS ON PATTERN ANALYSIS AND MACHINE INTELLIGENCE, VOL. 13, NO. 3, MARCH 1991

Fast B-Spline Transforms for Continuous Image Representation and Interpolation

Michael Unser, Akram Aldroubi, and Murray Eden

Abstract—This correspondence describes efficient algorithms for the continuous representation of a discrete signal in terms of B-splines (direct B-spline transform), and for interpolative signal reconstruction (indirect B-spline transform) with an expansion factor m . Expressions for the z -transforms of the sampled B-spline functions are determined and a convolution property of these kernels is established. It is shown that both the direct and indirect spline transforms involve linear operators that are

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A. Aldroubi and M. Eden are with the Biomedical and Instrumentation Program, National Institutes of Health, Bethesda, MD 20892.
IEEE Log Number 9040953.

IEEE TRANSACTIONS ON SIGNAL PROCESSING, VOL. 41, NO. 2, FEBRUARY 1993

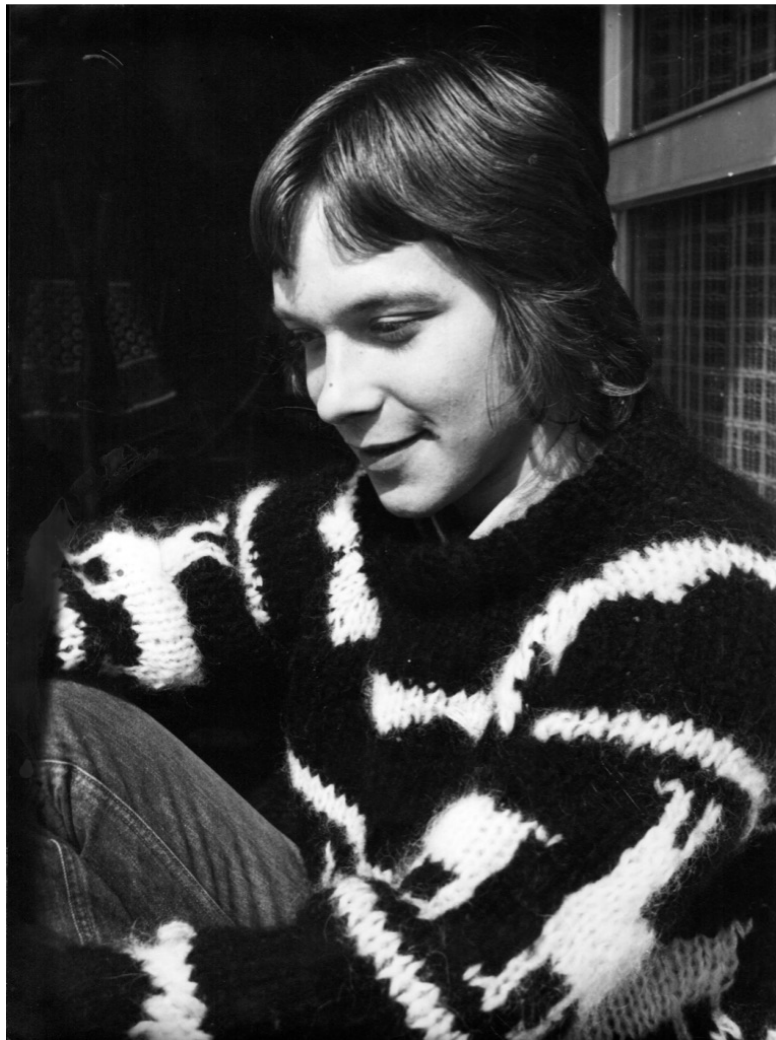
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B-Spline Signal Processing: Part I—Theory

Michael Unser, Member, IEEE, Akram Aldroubi, and Murray Eden, Life Fellow, IEEE

Abstract—This paper describes a set of efficient filtering techniques for the processing and representation of signals in terms of continuous B-spline basis functions. We first consider the problem of determining the spline coefficients for an exact signal interpolation (direct B-spline transform). The reverse operation is the signal reconstruction from its spline coefficients with an optional zooming factor m (indirect B-spline transform). We derive general expressions for the z -transforms and the equivalent continuous impulse responses of B-spline interpolators of order n . We present simple techniques for signal differentiation and filtering in the transformed domain. We then derive recursive filters that efficiently solve the problems of smoothing spline and least squares approximations. The smoothing spline technique approximates a signal with a complete set of coefficients subject to certain regularization or smoothness constraints. The least squares approach, on the other hand, uses a reduced number of B-spline coefficients with equally spaced nodes; this technique is in many ways analogous to the application of antialiasing low-pass filter prior to decimation in order to represent a signal correctly with a reduced number of samples.

Edge detection is a good example in which the use of a continuous signal representation is particularly appropriate. Most algorithms are based on the evaluation of spatial gradients or Laplacians [5]. Early techniques relied on finite differences to estimate these quantities [6], [7]; however, these simple operators used on noisy images perform poorly. More recent approaches often depend on the concept of fitting a continuous surface locally to the data [5], [8], [9]. Haralick used local least squares polynomial fits to determine the zero crossing of the directional second derivatives [9]. Poggio *et al.* proposed a smoothing cubic spline technique to improve the estimation of the intensity gradient in the presence of noise [10], [11]. These authors showed the approach to be more or less equivalent to smoothing the image with a Gaussian low-pass filter in a preprocessing step. In fact, an initial smoothing operation is implicit to all least squares techniques and is used in almost any modern edge detection



1991-93: Splines and wavelets

864 IEEE TRANSACTIONS ON INFORMATION THEORY, VOL. 38, NO. 2, MARCH 1992

On the Asymptotic Convergence of *B*-Spline Wavelets to Gabor Functions

Michael Unser, Member, IEEE, Akram Aldroubi, and Murray Eden, Life Fellow, IEEE

Abstract—A family of nonorthogonal polynomial spline wavelet transforms is considered. These transforms are fully reversible and can be implemented efficiently. The corresponding wavelet functions have a compact support. It is proven that these *B*-spline wavelets converge to Gabor functions (modulated Gaussians) pointwise and in all L_p norms with $1 \leq p < +\infty$ as the order of the spline (n) tends to infinity. In fact, the approximation error for the cubic *B*-spline wavelet ($n = 3$) is already less than 3%; this function is also near optimal in terms of its time/frequency localization in the sense that its variance product is within 2% of the limit specified by the uncertainty principle.

Index Terms—Wavelet transform, Gabor transform, uncertainty principle, polynomial spline, *B*-splines, time-frequency localization.

The corresponding basis functions are obtained by translation (index k) and dilation (index l) of a single prototype: the wavelet function ψ . One of the interesting properties of the wavelet transform is that it is relatively easy to construct a function $\tilde{\psi}$ that satisfies the biorthogonality condition

$$\langle \tilde{\psi}(2^{-l}x - k), \psi(2^{-j}x - l) \rangle = \begin{cases} 2^l & \text{for } (i = j) \text{ and } (k = l) \\ 0, & \text{otherwise.} \end{cases} \quad (1.4)$$

This function can be used to obtain the expansion coefficients by simple inner product

$$d_{i,j}(k) = \langle g(x), 2^{-j/2} \tilde{\psi}(2^{-j}x - k) \rangle. \quad (1.5)$$


Signal Processing 30 (1993) 141–162 Elsevier 141

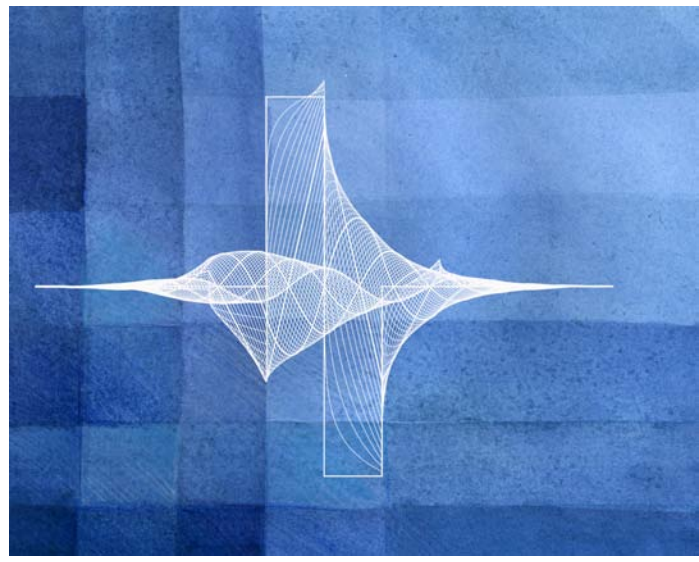
A family of polynomial spline wavelet transforms

Michael Unser, Akram Aldroubi and Murray Eden
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Received 25 July 1991
 Revised 24 January 1992 and 25 May 1992

Abstract. This paper presents an extension of the family of orthogonal Battle/Lemarié spline wavelet transforms with emphasis on filter bank implementation. Spline wavelets that are not necessarily orthogonal within the same resolution level, are constructed by linear combination of polynomial spline wavelets of compact support, the natural counterpart of classical *B*-spline functions. Mallat's fast wavelet transform algorithm is extended to deal with these non-orthogonal basis functions. The impulse and frequency responses of the corresponding analysis and synthesis filters are derived explicitly for polynomial splines of any order n (n odd). The link with the general framework of biorthogonal wavelet transforms is also made explicit. The special cases of orthogonal, *B*-spline, cardinal and dual wavelets are considered in greater detail. The *B*-spline (respectively dual) representation is associated with simple FIR binomial synthesis (respectively analysis) filters and recursive analysis (respectively synthesis) filters. The cardinal representation provides a sampled representation of the underlying continuous functions (interpolation property). The distinction between cardinal and orthogonal representation vanishes as the order of the spline is increased; both wavelets tend asymptotically to the bandlimited sinc-wavelet. The distinctive features of these various representations are discussed and illustrated with a texture analysis example.

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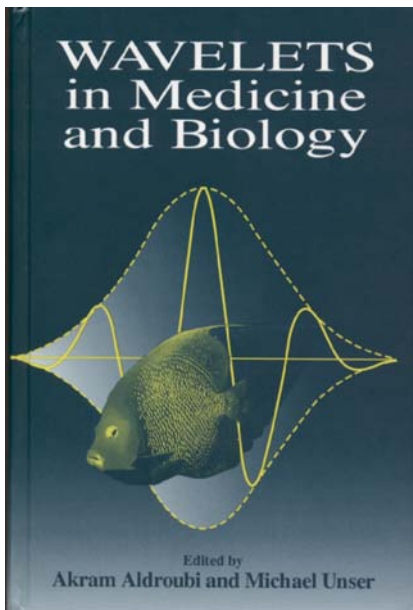


Lumini 2009



San Diego 2003

1991-97: Wavelets in medicine and biology



CRC Press, 1996

PROCEEDINGS OF THE IEEE, VOL. 84, NO. 4, APRIL, 1996

A Review of Wavelets in Biomedical Applications

MICHAEL UNSER, SENIOR MEMBER, IEEE, AND AKRAM ALDROUBI

Invited Paper

In this paper, we present an overview of the various uses of the wavelet transform (WT) in medicine and biology. We start by describing the wavelet properties that are the most important for biomedical applications. In particular, we provide an interpretation of the continuous wavelet transform (CWT) as a prewhitening multiscale matched filter. We also briefly indicate the analogy between the WT and some of the biological processing that occurs in the early components of the auditory and visual system. We then review the uses of the WT for the analysis of 1-D physiological signals obtained by phonocardiography, electrocardiography (ECG), and electroencephalography (EEG), including evoked response potentials. Next, we provide a survey of recent wavelet developments in medical imaging. These include biomedical image processing algorithms (e.g., noise reduction, image enhancement, and detection of microcalcifications in mammograms), image reconstruction and acquisition schemes (tomography, and magnetic resonance imaging (MRI)), and multiresolution methods for the registration and statistical analysis of functional images of the brain (positron emission tomography (PET) and functional MRI (fMRI)). In each case, we provide the reader with some general background information and a brief explanation of how the methods work.

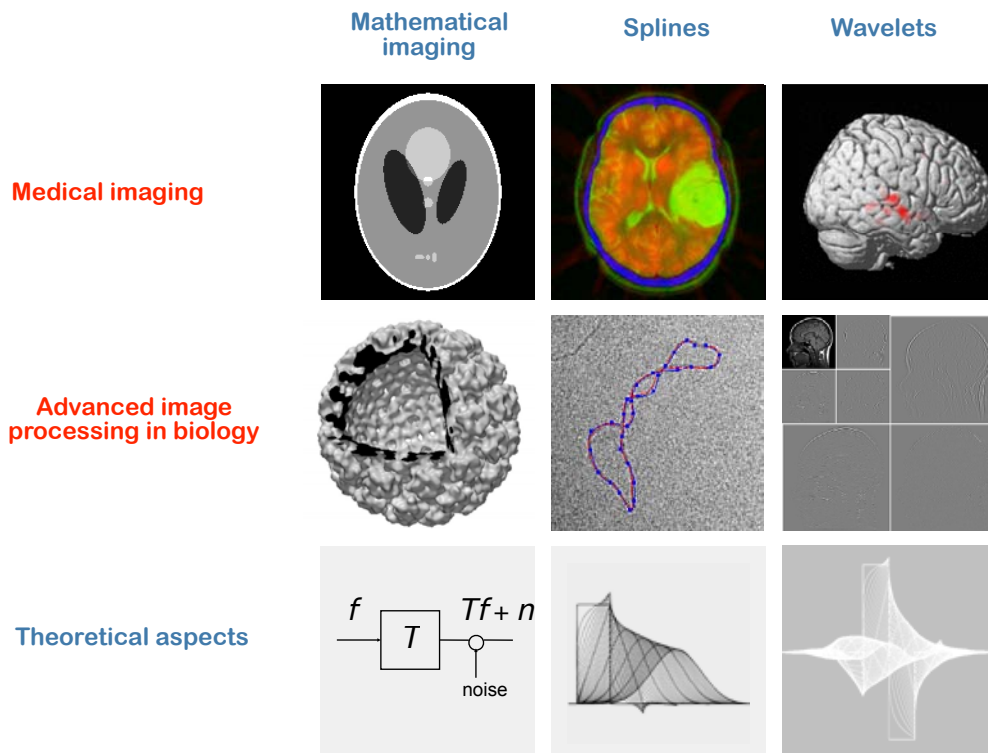
tomography (PET) and magnetic resonance imaging (MRI). The main difficulty in dealing with biomedical objects is the extreme variability of the signals and the necessity to operate on a case by case basis. Often, one does not know *a priori* what is the pertinent information and/or at which scale it is located. For example, it is frequently the deviation of some signal feature from the normal that is the most relevant information for diagnosis. As a result, the problems tend to be less well defined than those in engineering and the emphasis is more on designing robust methods that work in most circumstances, rather than procedures that are optimal under very specific assumptions. Another important aspect of biomedical signals is that the information of interest is often a combination of features that are well localized temporally or spatially (e.g., spikes and transients in electroencephalograph (EEG) signals and microcalcifications in mammograms) and others that are more diffuse (e.g., small oscillations, bursts, and texture). This requires the use of analysis methods sufficiently versatile to handle events that can be at opposite extremes in terms of their time-frequency localization.

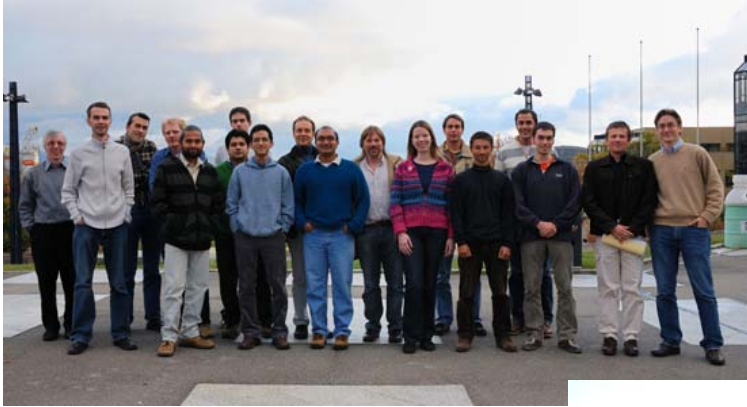
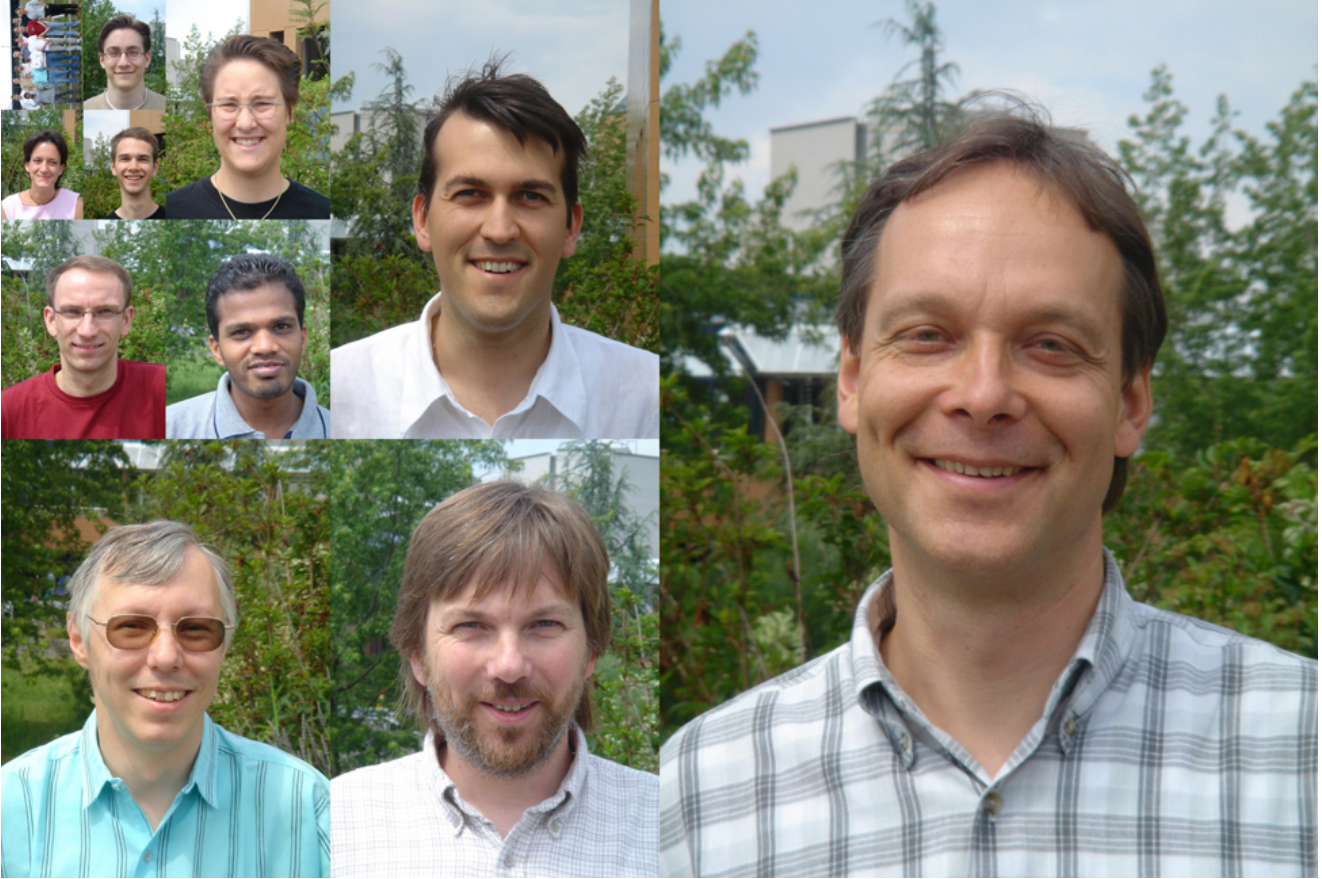
I. INTRODUCTION



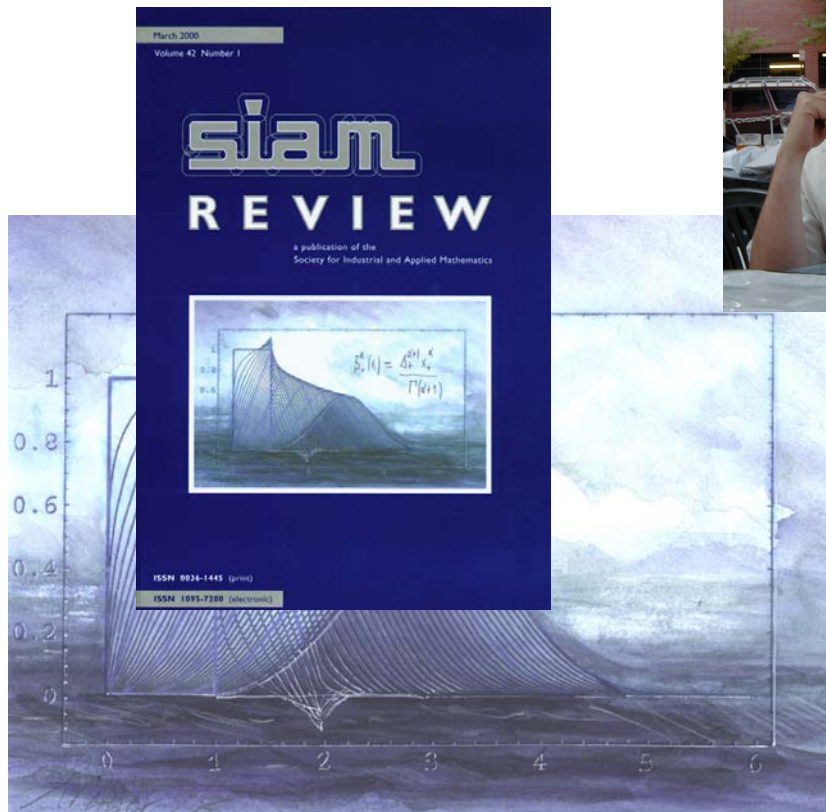


BIG's research agenda





One BIG Memory: going fractional



IEEE Signal Processing Magazine



Splines

A Perfect Fit for a scientific career

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