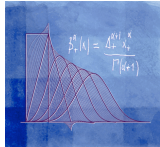


Medical image interpolation: The quest for higher quality

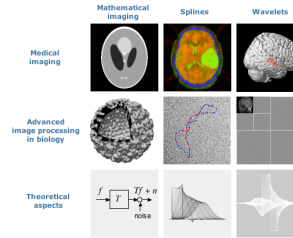
Michael Unser
Biomedical Imaging Group
EPFL, Lausanne
Switzerland



Medical Imaging Conference, Rome, October 16-22, 2004

Biomedical imaging group

Inter-disciplinary research strategy



2

OUTLINE

- Introduction
- Spline interpolation
- Splines and approximation theory
- Application example: image registration
- Interpolation in the presence of noise
- Conclusions

3

Back to the fundamentals: interpolation

Fundamental issue in imaging and signal processing

Linking the *discrete* and the *continuous*

Acquisition

Algorithm design

Mismatch between classical theory and practice

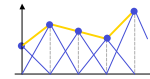
- Classical theory : Shannon's sampling theorem
- Practice: nearest neighbor, linear interpolation

Limitations of Shannon's sampling theory

- Ideal lowpass filters do not exist
- Incompatible with finite support signals
- Gibbs oscillations
- Slow decay of $\text{sinc}(x)$

Basic problem

How do you interpolate a signal ?



4

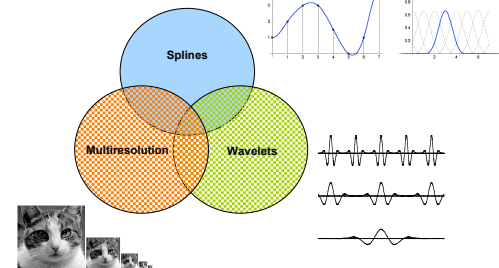
Interpolation and bio-imaging

Image processing task	Image processing	Image processing
Tomographic reconstruction	<ul style="list-style-type: none"> • Filtered backprojection • Fourier reconstruction • Relative techniques • SD + line 	<ul style="list-style-type: none"> • Commercial CT (X-rays) • EM • PET, SPECT • Dynamic CT, SPECT, PET
Sampling grid conversion	<ul style="list-style-type: none"> • Polar-to-cartesian coordinates • Spiral sampling • k-space sampling • Scan conversion 	<ul style="list-style-type: none"> • Ultrasound (endovascular) • Spiral CT, MRI • MRI
Visualization	<ul style="list-style-type: none"> • 2D operations • Zooming, panning, rotation • Resizing, scaling • Stereo imaging • Range stereoscopy 	<ul style="list-style-type: none"> • All • Fundus camera • CCT
3D operations	<ul style="list-style-type: none"> • Reslicing • Max. intensity projection • Simulated X-ray projection • Surface/volume rendering • Iso-surface ray tracing • Gradient-based shading • Stereogram 	<ul style="list-style-type: none"> • CT, MRI, MRA • CT • MRI
Geometrical correction	<ul style="list-style-type: none"> • Multi-angle lenses • Projective mapping • Aspect ratio, fill • Metastatic foci distortions 	<ul style="list-style-type: none"> • Endoscopy • C-Arm fluoroscopy • Dental X-rays • MRI
Registration	<ul style="list-style-type: none"> • Motion compensation • Image subtraction • Masking • Contrast-enhancing • Patient positioning • Retrospective compensation • Multi-modality imaging • Stochastic normalization • Brain warping 	<ul style="list-style-type: none"> • MRI, fundus camera • DSA • Endoscopy, fundus camera, EM microscopy • Surgery, radiotherapy • CT/PET/MR
Feature detection	<ul style="list-style-type: none"> • Contours • Ridge • Differential geometry 	<ul style="list-style-type: none"> • All
Contour extraction	<ul style="list-style-type: none"> • Static and active contours 	<ul style="list-style-type: none"> • MRI, Microscopy (cytology)

5

Splines: a unifying framework

Linking the discrete and the continuous



6

Splines: Bad press phenomenon

- Classical review article on interpolation, IEEE TMI, 1983
Comparison of four interpolators:
"The cubic B-spline provides the most smoothing."
- Classical book on *Digital Image Processing*, 1991 (2nd ed)
About high order B-splines:
"[out-of-band] interpolation error reduces significantly for higher order interpolation functions, but at the expense of resolution error [i.e., distortion]"
- Recent book on *Volume Rendering*, 1998
"The results of scaling the original image using [cubic] B-spline interpolation are shown in Figure 5.20. You can see the blurring effects"

7

SPLINE INTERPOLATION

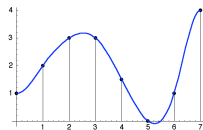
- Splines: definition
- B-spline basis functions
- B-spline interpolation
- Spline interpolators
- Geometric transformation of images

8

Splines: definition

Definition: A polynomial spline of degree n , $s(x)$, with knots $\dots < x_k < x_{k+1} < \dots$ is a function with the following two properties:

- Piecewise polynomial:
 $s(x)$ is a polynomial of degree n in each interval $[x_k, x_{k+1})$;
- Higher-order continuity:
 $s(x), s'(x), \dots, s^{(n-1)}(x)$ are continuous at the knots x_k .



- Effective degrees of freedom per segment:

$$\begin{matrix} n+1 & - & n & = & 1 \\ \text{(polynomial coefficients)} & & \text{(constraints)} & & \end{matrix}$$

- Cardinal splines** = unit spacing and infinite number of knots

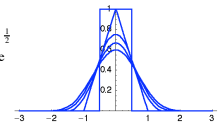
⇒ The right framework for signal processing

9

B-spline basis functions

- B-spline of degree n

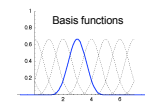
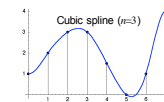
$$\beta^n(x) = \underbrace{\beta^0 * \beta^0 * \dots * \beta^0}_{(n+1) \text{ times}}(x), \quad \beta^0(x) = \begin{cases} 1 & -\frac{1}{2} \leq x < \frac{1}{2} \\ 0 & \text{otherwise} \end{cases}$$



- Theorem** (Schoenberg, 1946)

Any cardinal spline can be represented as a linear combination of shifted B-splines:

$$s(x) = \sum_k c(k) \beta^n(x-k)$$

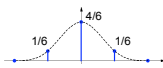


10

B-spline interpolation

- Discrete B-spline kernels

$$b_k^n(k) = \beta^n(x) \Big|_{x=k} \longleftrightarrow B^n(z) = \sum_{k=-\lfloor n/2 \rfloor}^{\lfloor n/2 \rfloor} \beta^n(k) z^{-k}$$



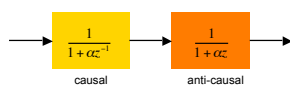
- B-spline interpolation: filtering solution

$$f(x) \Big|_{x=k} = \sum_l c(l) \beta^n(x-l) \Big|_{x=k} = (b_k^n * c)(k) \Rightarrow c(k) = ((b_k^n)^{-1} * f)(k)$$

- Efficient recursive solution

$$(b_k^n)^{-1}(k) \longleftrightarrow \frac{6}{z+4+z^{-1}} = \frac{-6\alpha}{(1-\alpha z)(1-\alpha z^{-1})} \quad \text{(symmetric exponential)}$$

⇒ Cascade of first order recursive filters



11

Generic C-code (splines of any degree n)

- Main recursion

```
void ConvertToInterpolationCoefficients (
    double c[], long DataLength, double z[], long NbPoles, double Tolerance)
{
    double Lambda = 1.0; long n, k;
    if (DataLength == 1) return;
    for (k = 0L; k < NbPoles; k++) Lambda = Lambda * (1.0 - z[k]) * (1.0 - 1.0 / z[k]);
    for (n = 0L; n < DataLength; n++) c[n] = Lambda;
    for (k = 0L; k < NbPoles; k++) {
        c[0] = InitialCausalCoefficient(c, DataLength, z[k], Tolerance);
        for (n = 1L; n < DataLength; n++) c[n] += z[k] * c[n-1];
        c[DataLength-1] = (c[0] / c[0] * z[k] - 1.0)
            * z[k] * c[DataLength-2L] + c[DataLength-1L];
        for (n = DataLength-2L; 0 <= n; n--) c[n] = z[k] * (c[n+1L] * c[n]);
    }
}
```

- Initialization

```
double InitialCausalCoefficient (
    double c[], long DataLength, double z, double Tolerance)
{
    double Sum, zn, z2n; for long n, Horizon;
    Horizon = (long)(ceil(log(Tolerance) / log(fabs(z))));
    if (DataLength < Horizon) Horizon = DataLength;
    zn = z; Sum = c[0];
    for (n = 1L; n < Horizon; n++) (Sum += zn * c[n]; zn *= z);
    return(Sum);
}
```

12

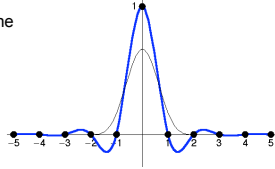
Spline interpolation

- Equivalent forms of spline representation

$$s(x) = \sum_{k \in \mathbb{Z}} c(k) \beta^n(x-k) = \sum_{k \in \mathbb{Z}} (s(k) * (b^n)^{-1}(k)) \beta^n(x-k) = \sum_{k \in \mathbb{Z}} s(k) \varphi_{\text{int}}^n(x-k)$$

- Cardinal (or fundamental) spline

$$\varphi_{\text{int}}^n(x) = \sum_{k \in \mathbb{Z}} (b^n)^{-1}(k) \beta^n(x-k)$$



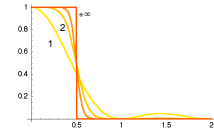
⇒ Finite cost implementation of an infinite impulse response interpolator !

13

Limiting behavior

- Spline interpolator

Impulse response $\varphi_{\text{int}}^n(x) \xrightarrow{\text{Fourier}} H^n(\omega) = \left(\frac{\sin(\omega/2)}{\omega/2} \right)^{n+1} \frac{1}{B_1^n(e^{j\omega})}$



- Asymptotic property

The cardinal spline interpolators converge to the sinc-interpolator (ideal filter) as $n \rightarrow +\infty$:

$$\lim_{n \rightarrow \infty} \varphi_{\text{int}}^n(x) = \text{sinc}(x), \quad \lim_{n \rightarrow \infty} H^n(\omega) = \text{rect}(\omega/2\pi) \quad (\text{in all } L_p\text{-norms})$$

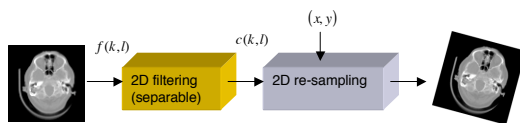
⇒ Includes Shannon's theory as a particular case !

14

Geometric transformation of images

- 2D separable model

$$f(x,y) = \sum_{k=-K_x}^{K_x-1} \sum_{l=-K_y}^{K_y-1} c(k,l) \beta^n(x-k) \beta^n(y-l)$$

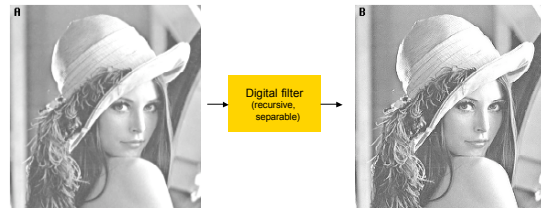


- Applications

zooming, rotation, re-sizing, re-formatting, warping

15

Cubic spline coefficients in 2D



Pixel values $f(k,l)$

B-spline coefficients $c(k,l)$

16

Interpolation benchmark

Cumulative rotation experiment: the best algorithm wins !



Bilinear

Windowed-sinc

Cubic spline

17

SPLINES AND APPROXIMATION THEORY

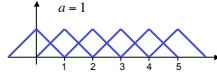
- Order of approximation
- Quantitative L_2 approximation

18

Order of approximation

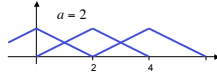
- General "shift-invariant" space at scale a

$$V_a(\varphi) = \left\{ s_a(x) = \sum_{k \in \mathbb{Z}} c(k) \varphi\left(\frac{x-k}{a}\right) : c \in \ell_2 \right\}$$



- Projection operator

$$\forall f \in L_2, P_a f = \arg \min_{s \in V_a} \|f - s\|_{L_2} \in V_a$$



- Order of approximation

DEFINITION

A scaling/generating function φ has order of approximation L iff

$$\forall f \in W_2^L, \|f - P_a f\|_{L_2} \leq C \cdot a^L \cdot \|f^{(L)}\|_{L_2}$$

➔ B-splines of degree n have order of approximation $L=n+1$

19

Spline reconstruction of a CAT-scan

Piecewise constant
 $L = 1$



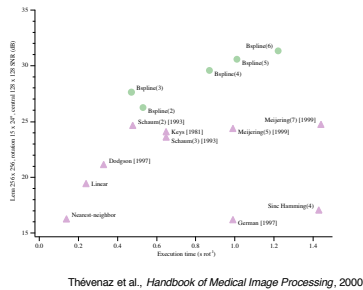
Cubic spline
 $L = 4$



20

High-quality image interpolation

- B-splines & O-MOMS: best cost-performance tradeoff



Thévenaz et al., *Handbook of Medical Image Processing*, 2000

21

APPLICATION EXAMPLE

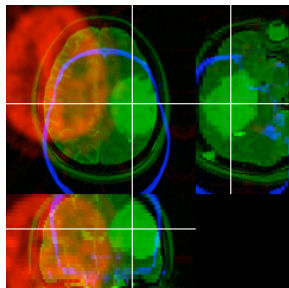
- Image Registration
 - Rigid body
 - Elastic

22

Splines: multi-modal image registration

Specificities of the approach

- Criterion: mutual-information (Colignon et al., 1996)
- Cubic spline model
 - high quality
 - sub-pixel accuracy
- Multiresolution strategy
- Marquardt-Levenberg like optimizer
 - Speed
 - Robustness



Thévenaz and Unser, *IEEE Trans. Imag Proc.*, 2000

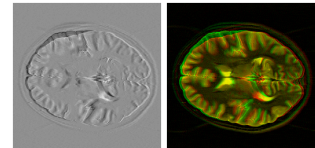
23

Splines: registration by elastic deformation

- Problem formulation

- Optimization criterion:

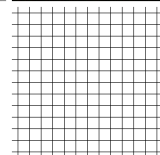
$$\min_{T \in \mathcal{T}} \|f_1(x) - f_2(T(x))\|$$



Number: 0
Image: 256x256
Pix/Node: 32x32
E: 749.685

- Principles of the method

- Splines
 - continuous image representation
 - deformation model with adjustable size
- Multiresolution; coarse-to-fine
 - image pyramids
 - deformation model
- Non-linear optimization



Kybic et al., *IEEE Trans. Medical Imaging*, 2000

24

INTERPOLATION OF NOISY DATA

- Tikhonov regularization
- Smoothing splines
- MMSE (or Wiener) solution

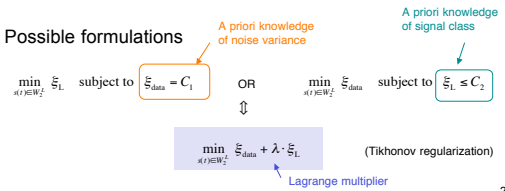
25

Fitting noisy data: Tikhonov regularization

Context

- Input data $\{f[k]\}_{k \in \mathbb{Z}}$ corrupted by noise
- Model : continuous-time function $s(t)$
- Data term : $\xi_{\text{data}} = \sum_{k \in \mathbb{Z}} |f[k] - s(k)|^2$
- Spline energy : $\xi_L = \|D^L s\|_{L_2}^2$ (measures lack of smoothness)

Possible formulations



26

Smoothing splines

Theorem [Schoenberg, 1964]

For a given sequence $f[k] \in \ell_2$ and regularization parameter $\lambda \geq 0$, the minimizer of

$$s_\lambda(t) = \arg \min_{s \in W_2^L} \left\{ \sum_{k \in \mathbb{Z}} (f[k] - s(k))^2 + \lambda \cdot \|D^L s\|_{L_2}^2 \right\}$$

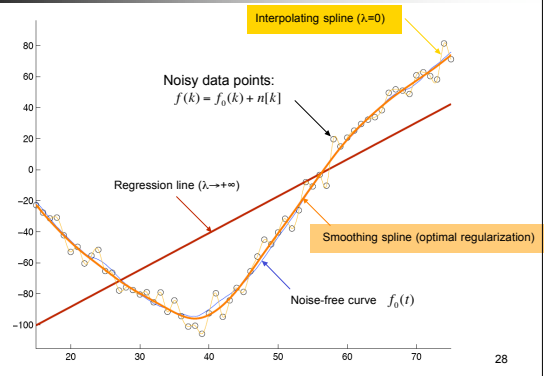
is unique and is a cardinal polynomial spline of degree $n = 2L - 1$.

Example : L=2

- Spline energy : $\|D^2 s\|_{L_2}^2 = \int_{-\infty}^{+\infty} |\ddot{s}(t)|^2 dt$ (bending energy)
- Optimal solution: cubic spline ($n=2 \times 2 - 1 = 3$)
- Extreme cases:
 - $\lambda \rightarrow +\infty$: best fitting line (linear regression)
 - $\lambda \rightarrow 0$: cubic spline interpolant (minimum curvature solution)

27

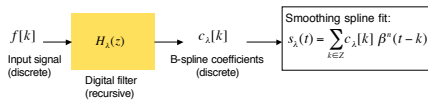
Cubic smoothing spline: example



28

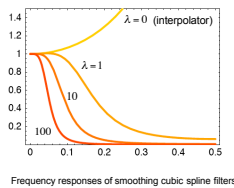
Smoothing splines: implementation

Digital filter-based implementation



Smoothing spline filter

$$H_\lambda(z) = \frac{1}{\sum_{k \in \mathbb{Z}} \beta^2(k) z^{-k} + \lambda \cdot (-z + 2 - z^{-1})^L}$$



Generalization: L'L smoothing splines

$$L = D^L + a_1 D^{L-1} + \dots + a_L I : \text{differential operator of order } L \quad \longleftrightarrow \quad \hat{L}(\omega) = \prod_{k=1}^L (j\omega - \alpha_k)$$

Theorem [U-Blu, 2004]

For a given sequence $f[k] \in \ell_2$ and regularization parameter $\lambda \geq 0$, the minimizer of

$$s_\lambda(t) = \arg \min_{s \in W_2^L} \left\{ \sum_{k \in \mathbb{Z}} (f[k] - s(k))^2 + \lambda \cdot \|L s\|_{L_2}^2 \right\}$$

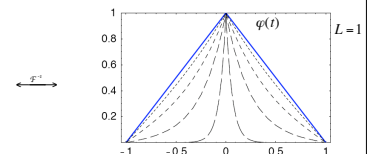
is the unique cardinal L'L-spline : $s_\lambda(t) = \sum_{k \in \mathbb{Z}} (h_\lambda * f)[k] \varphi(t-k)$,

where $\varphi(t)$ is a corresponding exponential B-spline of order $2L$ and

where h_λ is a suitable smoothing spline filter.

Exponential B-spline

$$\hat{\varphi}(\omega) = \frac{\prod_{k=1}^L |1 - e^{j\omega - \alpha_k}|^2}{\prod_{k=1}^L |j\omega - \alpha_k|^2}$$



Stochastic signal models

Wide sense stationary processes

Realization of the stochastic process: $x(t)$

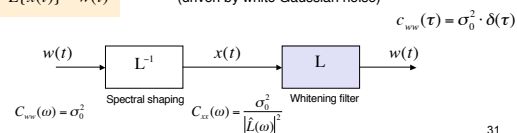
Zero mean: $E\{x(t)\} = 0$

Autocorrelation function: $E\{x(t)x(t+\tau)\} = c_{xx}(\tau) \in L_2$

Spectral density function: $C_{xx}(\omega) = \mathcal{F}\{c_{xx}(\tau)\} \in L_2$

Stochastic differential equation

$L\{x(t)\} = w(t)$ (driven by white Gaussian noise)



31

MMSE (or Wiener) solution

Statistical hypotheses

Discrete measurements (signal + noise): $y[k] = x(k) + n[k]$

Signal autocorrelation: $c_{xx}(\tau)$ such that $L\{L\{c_{xx}(\tau)\}\} = \sigma_0^2 \cdot \delta(\tau)$

Discrete white noise with variance $\sigma^2 \Rightarrow c_{nn}[k] = \sigma^2 \cdot \delta[k]$

MMSE continuous-time signal estimation

Theorem [U.-Blu, 2004]

Under the above assumptions, the linear Minimum Mean Square Error estimator of $x(t)$ at time $t = t_0$ given the measurements $\{y[k]\}_{k \in \mathbb{Z}}$ is $s_x(t_0)$ with $\lambda = \sigma^2 / \sigma_0^2$, where $s_x(t)$ is LL cardinal smoothing spline fit of $\{y[k]\}_{k \in \mathbb{Z}}$, as specified previously.

Remark: optimal over all estimators if one adds the assumption of Gaussianity

32

On the optimality of splines

Splines and continuous-time Tikhonov regularization

- Spline interpolators are optimal: they have minimum « spline energy » (e.g., curvature) among all possible interpolants
- Smoothing splines are optimal: they provide the best regularized fit of the input data, among all possible functions

Splines are optimal statistical estimators

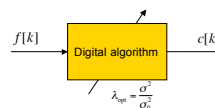
- Smoothing spline = MMSE estimator for fractal-like ($1/\omega^L$) processes
- Optimal regularization factor: $\lambda \propto \sigma^2$
- Can yield optimal estimators of derivatives, etc.
- Estimator can be fine tuned to the spectral characteristics of input signal \Rightarrow generalized splines

(work in progress)

33

Bottom line for practical applications

- Selection of the “optimal” spline space (e.g., cubic splines)
- B-spline coefficients determination via an appropriate filtering algorithm (interpolation or smoothing spline)



- Interpolation step (remains the same in all cases)

$$s(x) = \sum_{k \in \mathbb{Z}} c[k] \varphi(x - k)$$

34

CONCLUSION

Distinctive features of splines

- Simple to manipulate
- Smooth and well-behaved
- Excellent approximation properties
- Multiresolution properties (Wavelets !)
- Optimality properties (variational, statistical, ...)

Splines and medical imaging

- A story of avoidance and, more recently, love....
- Best cost/performance tradeoff
- Many applications

Unifying signal processing formulation

- Tools: digital filters, convolution operators
- Efficient recursive filtering solution
- Exact calculus (differential operators, etc.)
- Flexibility: piecewise constant to bandlimited

35

Splines: the end of the tunnel

Recent survey article on interpolation, *IEEE TMI*, 2000

Comparison of 31 interpolation algorithms:
“It [the cubic B-spline interpolator] produces one of the best results in terms of similarity to the original images, and of the top methods, it runs fastest.”

Addendum on spline interpolation, *IEEE TMI*, 2001

“Therefore, high degree B-splines are preferable interpolators for numerous applications in medical imaging, particularly if high precision is required.”

Recent evaluation of interpolation, *Med. Image Anal.*, 2001

Comparison of 126 interpolation algorithms:
“The results show that spline interpolation is to be preferred over all other methods, both for its accuracy and its relatively low cost.”

High-quality spline interpolation algorithms were included in the 2003 release of SPM (version 2b), a freely-available software package that is used worldwide for the statistical analysis of fMRI data.

36

Acknowledgments

Many thanks to

- Dr. Thierry Blu
- Prof. Akram Aldroubi
- Prof. Murray Eden
- Dr. Philippe Thévenaz
- Annette Unser, Artist

+ many other researchers,
and graduate students



- Software and demos at: <http://bigwww.epfl.ch>

37

Key references

- **Spline basics**
M. Unser, "Splines: a perfect fit for signal processing," *IEEE Signal Processing Magazine*, vol. 16, no. 6, pp. 22-38, November 1999.
- **Splines and approximation theory**
T. Blu, M. Unser, "Quantitative Fourier analysis of approximation techniques: Part I—Interpolators and projectors," *IEEE Trans. Signal Processing*, vol. 47, no. 10, pp. 2783-2795, October 1999.
- **Comparison of interpolators**
P. Thévenaz, T. Blu, M. Unser, "Interpolation revisited," *IEEE Trans. Medical Imaging*, vol. 19, no. 7, pp. 739-758, July 2000.
- **Interpolation of noisy data**
M. Unser, T. Blu, "Generalized smoothing splines and the optimal discretization of the Wiener filter," *IEEE Trans. Signal Processing*, in press.
- Preprints and software can be downloaded at:
<http://bigwww.epfl.ch>

38