E9 285 Biomedical imaging-Inverse problems

Chapter 2.b
Representative reconstruction methods for fluorescence microscopy

Iterative constrained Tikhonov Miller method (ICTM)

[1] H.T.M. van der Voort and K.C. Strasters, "Restoration of confocal images for quantitative image analysis," *J. Microsc.*, vol. 178, no. 2, pp. 165–181, 1995.

[2] G.M.P. van der Kempen, L.J. van Vliet, P.J. Verveer, and H.T.M. van der Voort, "A quantitative comparison of image restoration methods for confocal microscopy," *J. Microsc.*, vol. 185, no. 3, pp. 354–365, 1997.

ICTM

- 1) $\{L_1, L_2\}$: first derivative filters
- 2) Implements steepest descent and conjugate gradient algorithm where the gradient is clipped for negative values at each step.

Recap: Tikhonov filtering

$$x_{opt}(\mathbf{r}) = \underset{x(\mathbf{r})}{\operatorname{arg\,min}} \left[\sum_{\substack{\mathbf{r}' \in [1:N]^D \\ J_e(x)}} ((h*x)(\mathbf{r'}) - D(\mathbf{r'}))^2 + \lambda J_R(x) \right]$$

$$J_R(x) = \sum_{\mathbf{r}'} \left(\sum_{j=1}^{N_F} \left((L_j * x)(\mathbf{r'}) \right)^2 \right)$$

First consider the data term:

$$J_e(x) = \sum_{\mathbf{r}'} ((h * x)(\mathbf{r}'))^2 + \sum_{\mathbf{r}'} D^2(\mathbf{r}') - 2\sum_{\mathbf{r}'} (h * x)(\mathbf{r}')D(\mathbf{r}')$$

Consider the individual terms:

$$\sum_{\mathbf{r}'} ((h * x)(\mathbf{r}'))^2 = \langle h * x, h * x \rangle = \langle x, h^T * h * x \rangle, \text{ where } h^T(\mathbf{r}) = h(-\mathbf{r})$$

$$\sum_{\mathbf{r}'} (h * x)(\mathbf{r}') D(\mathbf{r}') = \langle h * x, D \rangle = \langle x, h^T * D \rangle$$

Hence we get

$$J_e(x) = \langle x, h^T * h * x \rangle - 2\langle x, h^T * D \rangle + \langle D, D \rangle$$

$$J_R(x) = \sum_{\mathbf{r}'} \left(\sum_{j=1}^{N_F} \left((L_j * x)(\mathbf{r}') \right)^2 \right) = \sum_{j=1}^{N_F} \left\langle x, L_j^T * L_j * x \right\rangle$$

Total cost: $J_e(x) + \lambda J_R(x)$

$$J_T(x) = \langle x, M * x \rangle - 2 \langle x, b \rangle + c$$

$$M = \mathbf{h}^{T} * \mathbf{h} + \lambda \sum_{j=1}^{N_{F}} L_{j}^{T} * L_{j}$$
 $b = \mathbf{h}^{T} * D$
 $c : \langle D, D \rangle$

$$J_T(\mathbf{x}) = \mathbf{x}^T \mathbf{Q} \mathbf{x} - 2\mathbf{x}^T \mathbf{b} + c$$

x: scanned vector of the image x

b: scanned vector of the image b

 \mathbf{Q} : convolution matrix of the filter M with respect to the scan order of \mathbf{x}

The gradient:

$$\nabla J_T(\mathbf{x}) = \mathbf{Q}\mathbf{x} - \mathbf{b}$$

$$\nabla J_T(x) = M * x - h^T * D$$

Tikhonov solution:

$$\nabla J_T(x) = M * x - h^T * D = 0$$

$$M(\mathbf{\Omega})x_{opt}(\mathbf{\Omega}) - \tilde{H}(\mathbf{\Omega})D(\mathbf{\Omega}) = 0$$

$$x_{opt}(\mathbf{\Omega}) = \frac{\tilde{H}(\mathbf{\Omega})D(\mathbf{\Omega})}{M(\mathbf{\Omega})}$$

The steepest descent method

Intialization:

Given \mathbf{x}_0

$$\mathbf{d}_0 = -\mathbf{g}_0 = -(\mathbf{Q}\mathbf{x}_0 - \mathbf{b}).$$

Iteration:

$$\mathbf{x}_{k+1} = \mathbf{x}_k + \alpha_k \mathbf{d}_k$$

$$\alpha_k = -\frac{\mathbf{g}_k^{\mathbf{T}} \mathbf{d}_k}{\mathbf{d}_k^{\mathbf{T}} \mathbf{Q} \mathbf{d}_k}$$

$$\mathbf{d}_{k+1} = -\mathbf{g}_{k+1}$$

where
$$\mathbf{g}_k = \mathbf{Q}\mathbf{x}_k - \mathbf{b}$$
.

Intialization:
$$x_0(\mathbf{r}) = F^{-1} \left[\frac{\tilde{H}(\mathbf{\Omega})D(\mathbf{\Omega})}{M(\mathbf{\Omega})} \right]$$

$$d_0 = -g_0 = -\nabla J_T(x) =$$

$$-\left(M * x_0 - h^T * D\right)$$

Iteration:

$$x_{k+1} = x_k + \alpha_k d_k$$

$$\alpha_{k} = -\frac{\langle g_{k}, d_{k} \rangle}{\langle d_{k}, M * d_{k} \rangle}$$

$$d_{k+1} = -g_{k+1}$$

where

$$g_k = -\left(M * x_0 - h^T * D\right)$$

ICTM

Intialization:
$$x_0(\mathbf{r}) = \text{ClipNeg}\left[F^{-1}\left[\frac{\tilde{H}(\mathbf{\Omega})D(\mathbf{\Omega})}{M(\mathbf{\Omega})}\right]\right]$$

$$d_0 = -g_0 = -\nabla J_T(x) =$$

$$\mathbf{ClipNeg}\Big(-\Big(M*x_0 - h^T*D\Big)\Big)$$

Iteration:

$$x_{k+1} = x_k + \alpha_k d_k$$

$$\alpha_{k} = -\frac{\langle g_{k}, d_{k} \rangle}{\langle d_{k}, M * d_{k} \rangle}$$

$$d_{k+1} = \mathbf{ClipNeg}(-g_{k+1})$$

where

$$g_k = -\left(M * x_0 - h^T * D\right)$$

The conjugate gradient method:

Given an initial guess for \mathbf{x}_0 for the minimum of the quadratic form $f(\mathbf{x}) = (1/2)\mathbf{x}^T\mathbf{Q}\mathbf{x} - \mathbf{b}^T\mathbf{x}$, let $\mathbf{d}_0 = -\mathbf{g}_0 = -(\mathbf{Q}\mathbf{x}_0 - \mathbf{b})$.

Then the iteration is given by

$$\mathbf{x}_{k+1} = \mathbf{x}_k + \alpha_k \mathbf{d}_k, \quad \alpha_k = -\frac{\mathbf{g}_k^{\mathsf{T}} \mathbf{d}_k}{\mathbf{d}_k^{\mathsf{T}} \mathbf{Q} \mathbf{d}_k}$$

$$\mathbf{d}_{k+1} = -\mathbf{g}_{k+1} + \boldsymbol{\beta}_k \mathbf{d}_k, \quad \boldsymbol{\beta}_k = \frac{\mathbf{g}_{k+1}^{\mathbf{T}} \mathbf{Q} \mathbf{d}_k}{\mathbf{d}_k^{\mathbf{T}} \mathbf{Q} \mathbf{d}_k},$$

where $\mathbf{g}_k = \mathbf{Q}\mathbf{x}_k - \mathbf{b}$.

ICTM-CG

Intialization:
$$x_0(\mathbf{r}) = \text{ClipNeg}\left(F^{-1}\left[\frac{\tilde{H}(\mathbf{\Omega})D(\mathbf{\Omega})}{M(\mathbf{\Omega})}\right]\right)$$

$$d_0 = -g_0 = -\nabla J_T(x) =$$

$$\mathbf{ClipNeg}\Big(-\Big(M*x_0 - h^T*D\Big)\Big)$$

Iteration:

$$x_{k+1} = x_k + \alpha_k d_k, \quad \alpha_k = -\frac{\langle g_k, d_k \rangle}{\langle d_k, M * d_k \rangle}$$

$$\tilde{d}_{k+1} = \mathbf{ClipNeg} \left(-g_{k+1} \right)$$

$$d_{k+1} = \tilde{d}_{k+1} + \beta_k d_k, \beta_k = \frac{\left\langle \tilde{d}_{k+1}, M * d_k \right\rangle}{\left\langle d_k, M * d_k \right\rangle}$$

where

$$g_k = -\left(M * x_k - h^T * D\right)$$

Assignment 3:

Write a program to compute $\nabla J_T(x) = M * x - h^T * D$.

Assignment 4:

Write a program for ICTM

Assignment 5:

Write a program for ICTM-CG

Quadratic data fitting with L1 regularization

S. Lefkimmiatis, A. Bourquard, M. Unser, "Hessian-Based Regularization for 3-D Microscopy Image Restoration", Proceedings of the Ninth IEEE International Symposium on Biomedical Imaging: From Nano to Macro (ISBI'12), Barcelona, Spain, May 2-5, 2012, pp. 1731-1734.

The minimization problem

$$x_{opt}(\mathbf{r}) = \underset{x(\mathbf{r})}{\operatorname{arg\,min}} \left[\sum_{\mathbf{r}' \in [1:N]^{D}} ((h * x)(\mathbf{r}') - D(\mathbf{r}'))^{2} + \lambda J_{R}(x) \right]$$

$$J_R(x) = \sum_{\mathbf{r}'} \sqrt{\sum_{j=1}^{N_F} \left((L_j * x)(\mathbf{r}') \right)^2}$$

The regularization functional

Consider the following:

$$\overline{R}_{\overline{x}}^{2}(\mathbf{r}) = \left(\frac{\partial^{2}}{\partial r_{2}^{2}} \overline{x}(\mathbf{r})\right)^{2} + \left(\frac{\partial^{2}}{\partial r_{1}^{2}} \overline{x}(\mathbf{r})\right)^{2} + \left(\frac{\partial^{2}}{\partial r_{3}^{2}} \overline{x}(\mathbf{r})\right)^{2}$$

$$\left(\frac{\partial^{2}}{\partial r_{2}^{2}} \right)^{2} \left(\frac{\partial^{2}}{\partial r_{3}^{2}} \right)^{2} \left(\frac{\partial^{2}}{\partial r_{3}^{2}} \right)^{2}$$

$$2\left(\frac{\partial^2}{\partial r_1 \partial r_3} \overline{x}(\mathbf{r})\right)^2 + 2\left(\frac{\partial^2}{\partial r_1 \partial r_2} \overline{x}(\mathbf{r})\right)^2 + 2\left(\frac{\partial^2}{\partial r_2 \partial r_3} \overline{x}(\mathbf{r})\right)^2,$$

where $\bar{x}(\mathbf{r})$ is a continuous domain function.

If \overline{x}_1 and \overline{x}_2 are images such that $\overline{x}_1(\mathbf{r}) = \overline{x}_2(\Theta \mathbf{r})$, where Θ is a 3 × 3 rotation matrix, then we have

$$\overline{R}_{\overline{x}_1}(\mathbf{r}) = \overline{R}_{\overline{x}_2}(\Theta \mathbf{r})$$

Now consider

$$R_x^2(\mathbf{r}) = \sum_{j=1}^{N_F} \left((L_j * x)(\mathbf{r'}) \right)^2.$$

Question: what are the filters that make R_x be an approximation $\overline{R}_{\overline{x}}(\mathbf{r})$

$$\mathbf{r} = (l, m, n)$$
 (compare it with $\mathbf{r} = (r_1, r_2, r_3)$ for continuous domain)

$$\delta = \frac{xy \text{ step size}}{z \text{ step size}}$$

$$L_1(\mathbf{r}) * x(\mathbf{r}) = x(l-1,m,n) - 2x(l,m,n) + x(l+1,m,n)$$

$$L_2(\mathbf{r}) * x(\mathbf{r}) = x(l, m-1, n) - 2x(l, m, n) + x(l, m+1, n)$$

$$L_3(\mathbf{r}) * x(\mathbf{r}) = \delta^2 (x(l,m,n-1) - 2x(l,m,n) + x(l,m,n+1))$$

$$L_4(\mathbf{r}) * x(\mathbf{r}) = \sqrt{2} \left[x(l,m,n) - x(l-1,m,n) - x(l,m-1,n) + x(l-1,m-1,n) \right]$$

$$L_5(\mathbf{r}) * x(\mathbf{r}) = \sqrt{2}\delta \left[x(l,m,n) - x(l,m-1,n) - x(l,m,n-1) + x(l,m-1,n-1) \right]$$

$$L_6(\mathbf{r}) * x(\mathbf{r}) = \sqrt{2}\delta [x(l,m,n) - x(l-1,m,n) - x(l,m,n-1) + x(l-1,m,n-1)],$$

Comparison:

$$\overline{R}_{\overline{x}}^{2}(\mathbf{r}) = \left(\frac{\partial^{2}}{\partial r_{2}^{2}} \overline{x}(\mathbf{r})\right)^{2} + \left(\frac{\partial^{2}}{\partial r_{1}^{2}} \overline{x}(\mathbf{r})\right)^{2} + \left(\frac{\partial^{2}}{\partial r_{3}^{2}} \overline{x}(\mathbf{r})\right)^{2}
2\left(\frac{\partial^{2}}{\partial r_{1}\partial r_{3}} \overline{x}(\mathbf{r})\right)^{2} + 2\left(\frac{\partial^{2}}{\partial r_{1}\partial r_{2}} \overline{x}(\mathbf{r})\right)^{2} + 2\left(\frac{\partial^{2}}{\partial r_{2}\partial r_{3}} \overline{x}(\mathbf{r})\right)^{2},$$

The majorization-minimization method

The mojorization relation: for any non-negative funtion, g, we have

$$\sqrt{g(y)} \le 0.5\sqrt{g(z)} + 0.5\frac{g(y)}{\sqrt{g(z)}}$$
 for every y and z

satisfying $g(y) \ge 0$ and $g(z) \ge 0$.

Given two images x and x', substitute $g(\bullet) = (\bullet)^2$ and $y=R_x(\mathbf{r}), z=R_{x'}(\mathbf{r})$:

$$R_{x}(\mathbf{r}) \leq 0.5R_{x'}(\mathbf{r}) + 0.5\frac{R_{x}^{2}(\mathbf{r})}{R_{x'}(\mathbf{r})}$$

$$\Rightarrow \sum_{\mathbf{r}} R_{x}(\mathbf{r}) \leq 0.5 \sum_{\mathbf{r}} R_{x'}(\mathbf{r}) + 0.5 \sum_{\mathbf{r}} \frac{R_{x}^{2}(\mathbf{r})}{R_{x'}(\mathbf{r})}$$

$$Q'_{R}(x,x')$$

$$\sum_{\mathbf{r}} R_{x}(\mathbf{r}) \leq 0.5 \sum_{\mathbf{r}} R_{x'}(\mathbf{r}) + 0.5 \sum_{\mathbf{r}} \frac{R_{x}^{2}(\mathbf{r})}{R_{x'}(\mathbf{r})}$$

$$Q'_{R}(x,x')$$

 $Q'_R(x,x')$ is quadratic in x.

Given $x^{(t)}$, define

$$x^{(t+1)} = \underset{\mathcal{X}}{\operatorname{arg\,min}} J_e(x) + \lambda Q_R'(x, x^{(t)}).$$

Then

$$\begin{split} &J_{e}(x^{(t+1)}) + \lambda J_{R}(x^{(t+1)}) \leq \\ &J_{e}(x^{(t+1)}) + \lambda Q'(x^{(t+1)}, x^{(t)}) \leq J_{e}(x^{(t)}) + \lambda Q'(x^{(t)}, x^{(t)}) \\ &= J_{e}(x^{(t)}) + \lambda J_{R}(x^{(t)}) \end{split}$$

$$Q_R'(x, x^{(t)}) = 0.5 \sum_{\mathbf{r}} R_{x^{(t)}}(\mathbf{r}) + 0.5 \sum_{\mathbf{r}} \frac{R_x^2(\mathbf{r})}{R_{x^{(t)}}(\mathbf{r})}$$

$$Q_R'(x, x^{(t)}) = 0.5 \sum_{\mathbf{r}} R_{x^{(t)}}(\mathbf{r}) + 0.5 \sum_{\mathbf{r}} \frac{R_x^2(\mathbf{r})}{R_{x^{(t)}}(\mathbf{r})}$$

The iteration:

$$x^{(t+1)} = \underset{x}{\operatorname{arg\,min}} \left[\sum_{\mathbf{r} \in [1:N]^{D}} ((h * x)(\mathbf{r}) - D(\mathbf{r}))^{2} + 0.5 \lambda Q_{R}(x, x^{(t)}) \right]$$

$$Q_R(x, x^{(t)}) = \sum_{\mathbf{r}} \left[\frac{\sum_{j=1}^{N_F} \left((L_j * x)(\mathbf{r}) \right)^2}{\sqrt{\sum_{j=1}^{N_F} \left((L_j * x^{(t)})(\mathbf{r}) \right)^2}} \right]$$

$$Q_R(x, x^{(t)}) = \sum_{\mathbf{r}} \left[\frac{\sum_{j=1}^{N_F} \left((L_j * x)(\mathbf{r}) \right)^2}{\sqrt{\sum_{j=1}^{N_F} \left((L_j * x^{(t)})(\mathbf{r}) \right)^2}} \right]$$

Define
$$W_{x^{(t)}}(\mathbf{r}) = \frac{1}{\sqrt{\sum_{j=1}^{N_F} ((L_j * x^{(t)})(\mathbf{r}))^2}}$$
.

Then
$$Q_R(x, x^{(t)}) = \sum_{j=1}^{N_F} \langle L_j * x, W_{x^{(t)}}(L_j * x) \rangle$$

= $\sum_{j=1}^{N_F} \langle x, L_j^T * (W_{x^{(t)}}(L_j * x)) \rangle$

Relating to the standard quadratic form

$$J_{x^{(t)}}(x) = \langle x, A_{x^{(t)}} x \rangle - 2 \langle x, b \rangle + c$$

$$A_{t}x = h^{T} * h * x + \lambda \sum_{j=1}^{N_{F}} L_{j}^{T} * (W_{x^{(t)}}(L_{j} * x))$$

$$b = h^{T} * D$$

$$c = \langle D, D \rangle$$

$$J_{\mathbf{x}^{(t)}}(\mathbf{x}) = \mathbf{x}^T \mathbf{Q}_t \mathbf{x} - 2\mathbf{b}^T \mathbf{x} + c$$

 \mathbf{x} : scanned vector of x

b: scanned vector of b

 \mathbf{Q}_t : matrix equivalent of A_t with respect to the scan order

 $D_{x^{(t)}}$: Diagonal approximation of $A_{x^{(t)}}$.

Define

$$\tilde{A}_{x^{(t)}}x = D^{-1/2}\left(h^{T} * h * \left(D_{x^{(t)}}^{-1/2}x\right)\right) + \lambda \sum_{j=1}^{N_{F}} D^{-1/2}\left(L_{j}^{T} * \left(W_{x^{(t)}}\left(L_{j} * \left(D_{x^{(t)}}^{-1/2}x\right)\right)\right)\right)$$

Minimization of $J_{x^{(t)}}(x)$

Intialization: $x_0 = x^{(t)}$

Compute $\tilde{A}_{x^{(t)}}$ from $A_{x^{(t)}}$

$$d_0 = -g_0 = -\nabla J_{x^{(t)}}(x_0) = -\left(\tilde{A}_{x^{(t)}}x_0 - h^T * D\right)$$

Iteration:

$$x_{k+1} = x_k + \alpha_k d_k, \quad \alpha_k = -\frac{\langle g_k, d_k \rangle}{\langle d_k, \tilde{A}_{x^{(t)}} d_k \rangle}$$

$$d_{k+1} = -g_{k+1} + \beta_k d_k, \quad \beta_k = \frac{\left\langle d_{k+1}, \tilde{A}_{x^{(t)}} d_k \right\rangle}{\left\langle d_k, \tilde{A}_{x^{(t)}} d_k \right\rangle}$$

where
$$g_k = -\left(\tilde{A}_{x^{(t)}} x_k - h^T * D\right)$$

What is $D_{x^{(t)}}$?

 $\mathbf{Q}_{x^{(t)}}$: matrix equivalent of $A_{x^{(t)}}$ with respect to the scan order

Let $\mathbf{d}_{x^{(t)}}$ be the diagonal of $\mathbf{Q}_{x^{(t)}}$.

Then $D_{x^{(t)}}$ obtained from reassembling $\mathbf{d}_{x^{(t)}}$.

Direct definition:

$$D_{x^{(t)}}(\mathbf{r'}) = B_{x^{(t)}}(\mathbf{r'}), \text{ where } B_{x^{(t)}}(\mathbf{r}) = A_{x^{(t)}}\delta(\mathbf{r} - \mathbf{r'})$$

Assignment 6:

Derive a formula to compute $D_{x^{(t)}}(\mathbf{r})$ from $W_{x^{(t)}}(\mathbf{r})$, h, and derivative filters L_1, \dots, L_6 .