Journal of Biomedical Optics 13(2), 1 (March/April 2008)

Highly resolved diffuse optical tomography: a systematic approach using high-pass filtering for value-preserved images

Min-Cheng Pan

Tungnan University Department of Electronic Engineering Taipei County 222, Taiwan

Chien-Hung Chen Liang-Yu Chen Min-Chun Pan

National Central University Graduate Institute of Biomedical Engineering Department of Mechanical Engineering Jhongli City, Taoyuan County 320, Taiwan E-mail: pan_minc@cc.ncu.edu.tw

Yi-Ming Shyr

Taipei Veterans General Hospital Department of Surgery National Yang Ming University Taipei 122, Taiwan

Abstract. We attempt to develop a systematic scheme through adopting high-pass filtering (HPF) to well resolve value-preserved images such as medical images. Our approach is derived from the Poisson maximum a posteriori superresolution algorithm employing the HP filters, where four filters are considered such as two low-pass-filtercombination based filters, wavelet filter, and negative-oriented Laplacian HP filter. The proposed approach is incorporated into the procedure of finite-element-method (FEM)-based image reconstruction for diffuse optical tomography in the direct current domain, posterior to each iteration without altering the original FEM modeling. This approach is justified with various HPF for different cases that breastlike phantoms embedded with two or three inclusions that imitate tumors are employed to examine the resolution performances under certain extreme conditions. The proposed approach to enhancing image resolution is evaluated for all tested cases. A qualitative investigation of reconstruction performance for each case is presented. Following this, we define a set of measures on the quantitative evaluation for a range of resolutions including separation, size, contrast, and location, thereby providing a comparable evaluation to the visual quality. The most satisfactory result is obtained by using the wavelet HP filter, and it successfully justifies our proposed scheme. © 2008 Society of Photo-Optical Instrumentation Engineers. [DOI: 10.1117/1.2907344]

Keywords: high-pass filtering; Poisson maximum *a posteriori*; superresolution algorithm; finite element method; image reconstruction; diffuse optical tomography; quantitative evaluation.

Paper 07288R received Jul. 31, 2007; revised manuscript received Nov. 21, 2007; accepted for publication Nov. 28, 2007.

¹ 1 Introduction

2 Over the last several decades, there has been great enthusiasm 3 in developing medical imaging techniques to assist physicians 4 in detecting and diagnosing tumors and diseases. Today, the 5 efforts drive toward developing imaging systems employing 6 noninvasive, nonradioactive, and relatively low cost instru-7 mentations. Near-infrared (NIR) diffuse optical tomography 8 (DOT) imaging is such an imaging modality that NIR light is 9 used to probe biological tissues and it is promising to continu-10 ously monitor the status of tissues using NIR imaging. There-11 fore, the realization of NIR DOT as a viable clinical imaging 12 modality would be a beneficial advancement in medical diag-13 nosis. Basically, both the absorption and scattering tomogra-14 phic images are evaluated in an NIR imaging system, thereby 15 relating absorption properties to the oxygen saturation of he-16 moglobin content and water content, as well as scattering 17 properties to the scatter size and density or the mitochondrial **18** compartment and blood glucose concentration.^{1–5}

about 19 An NIR spectral window exists from 650 to 1000 nm wherein the absorption is relatively small, 20 which enables transillumination of NIR radiance through bio- 21 logical tissues. With a difficulty arising from strongly scatter- 22 ing effects in human tissues, the contrast and resolution of 23 optical images are severely reduced. Compared with conven- 24 tional x-ray mammography, magnetic resonance imaging 25 (MRI), and ultrasound imaging all with acceptable resolutions 26 $(\sim 10^{0} \text{ mm})$, but low intrinsic contrast $(\sim 10^{-1})$, NIR imaging 27 possesses exceptionally high intrinsic contrast ($\sim 10^{1-2}$), but 28 exhibits inferior spatial resolutions ($\sim 10^1$ mm) as a result of 29 highly scattering nature of biological tissues.^{4,6} Many efforts 30 have been made to improve NIR optical tomographic image **31** resolution through different ways.^{7–34} Hebden and Deply⁷ pro- **32** posed a method using the least-squares fit between the 33 temporal-distribution measures of transmitted light and a 34 model of the diffusion equation to enhance time-resolved im- 35 aging. Moon and Reintjes⁸ applied the Markov-chain tech- 36 nique to enhance optical image resolution. Jiang and Paulsen⁹ 37 and Jiang et al.^{10–12} improved diffuse optical images in the 38direct current (dc) domain using the scheme with total varia- 39 40

Address all correspondence to Min-Chun Pan, Department of Mechanical Engineering, National Central University, No. 300, jhongda Rd.-Jhongli City, 320 Taiwan; Tel: 886-3-4267312; Fax: 886-3-4254501; E-mail: pan_minc@cc.ncu.edu.tw

^{1083-3668/2008/13(2)/1/0/\$25.00 © 2008} SPIE

⁴² tion minimization, dual mesh, and low-pass spatial filtering to 43 achieve a satisfactory result. Recently researchers have 44 adopted hybrid modalities to attain high-resolution NIR to-45 mographic images by the use of *a priori* structural informa-46 tion available from MRI (Refs. 13–18) or ultrasonic 47 imaging.^{19–21} Especially, the structure information acquired 48 from MRI was incorporated with a Laplacian-type regulariza-49 tion integrated in the inversion-computation process.^{22,23} Ad-50 ditionally, the spectral priors acquired from various source 51 wavelengths were combined with the reconstruction process, 52 validating improvement over spatial priors.²⁴

53 For the enhancement of image reconstruction, Kanmani 54 and Vasus²⁵ used a nonlinear approximation of the perturba-55 tion equation through adding the second term involving the 56 Hessian in the Taylor expansion instead of a linear perturba-57 tion model that adopts only the first order derivatives (the 58 Jacobian), which is solved by using conjugate gradient search. 59 Furthermore, Jiang²⁶ reconstructed optical images using the 60 third-order diffusion equation, providing more stable inverse 61 solutions. Pogue et al.²⁷ improved diffuse optical images with 62 spatially variant regularization in the radial orientation, 63 thereby minimizing high-frequency noise and producing con-64 stant image resolution and contrast. Brooks et al.²⁸ and Zhang 65 et al.²⁹ obtained accurate reconstruction images by the joint 66 use of measurement-model agreement, amplitude, and total 67 variation type constraints. Guven et al.^{30,31} proposed an adap-68 tive multigrid algorithm for the enhancement of image reso-69 lutions where two-level meshes were generated to provide **70** high resolution of the region of interest. Stott et al.³² presented 71 a technique to improve optical images through using simulta-72 neous calibration of optode positions that were sensitive to **73** image quality. Furthermore, Ntziachristos et al.³³ and Intes et 74 al.³⁴ employed a fluorescent diagnostic agent, indocyanine 75 green (ICG) to enhance heterogeneity contrast for obtaining 76 better resolutions prior to optical image reconstruction. As to 77 the background information about image processing tech-78 niques for the enhancement of reconstructed optical-property **79** images, especially applied in this study, some **80** monographs $^{35-37}$ and related papers $^{38-40}$ are valuable. Three 81 referred to books are rather appropriate for the beginner, es-82 pecially the third one, and three reference papers are actually 83 the origin of the idea resulting in the proposed algorithm pre-84 sented in this paper. Further, more references were reviewed **85** and introduced.

In this paper, the design of a high-pass filtering (HPF) 86 87 method to enhance optical images is studied. Based on the 88 viewpoint of image processing, generally, visual quality per-89 ception is preferred to actual image values. Moreover, as is 90 known, the effect of an HP filter applied on an image to be 91 processed yields a different image, which cannot be used 92 when true optical property values are required. In this paper, 93 we attempt to develop a systematic scheme through adopting 94 HP filters for value-preserved images such as medical images. 95 Therefore, as simply implementing a specific HP filter to re-96 solve NIR DOT images, it is not suitable that the procedure of 97 the conventional approach takes routine steps like HPF the 98 original image to be weighted and then histogram equaliza-99 tion. As can be understood, this conventional image-**100** processing procedure is performed on optical property images 101 between a reconstruction and a true distribution. For instance, **102** one may be more interested in estimating the true values than obtaining the visual effect. As a result, an approach to system-¹⁰³ atically implementing HPF is demanding. Additionally, recon- 104 structed highly resolved images that preserve true values are 105 extremely expected. In this paper, first we investigate the 106 properties of HP filters that can be classified into two types, 107 i.e., low-pass filter (LPF) combined form and a wavelet-like 108 filter, respectively. To preserve the true value distribution of 109 optical images, the approach proposed and realized is derived 110 from the Poisson maximum a priori (Poisson MAP) super- 111 resolution algorithm for the application of HPF on absorption- 112 and diffusion-coefficient DOT images. Following this, the 113 proposed approach is incorporated into the finite-element- 114 method (FEM)-based image reconstruction in the continuous 115 wave (cw) domain. Simulation results and their corresponding 116 evaluations are demonstrated and investigated by comparing 117 reconstruction with and without filtering. 118

This paper aims to (1) develop an approach derived from 119 the Poisson MAP superresolution algorithm to systematically 120 implement HP filters on optical property images; (2) justify 121 the proposed approach with various HP filters performed on 122 breast-like optical heterogeneity; (3) demonstrate the reso- 123 lution performance of this approach under certain extreme 124 conditions, significantly improving the reconstruction perfor- 125 mance even in the absence of a priori information or modified 126 reconstruction algorithms; and (4) define a set of measures for 127 the evaluation of computation resolutions on the separation, 128 size, and location of inclusions, and the contrast of inclusions 129 to background. Additionally, further discussions on these 130 measures are also provided. The paper is organized as fol- 131 lows. Section 2 briefly describes processing with HP filters 132 that are used to enhance an image. Following this, a novel 133 approach that starts from the Poisson MAP superresolution 134 algorithm is derived. Section 3 implements four HP filters on 135 several DOT images, and presents both qualitative and quan- 136 titative discussions of the reconstructed images. Finally, in 137 Sec. 4 we draw conclusions and discuss future works. 138

2 Theoretical Analysis of the Proposed Approach

Following from the introduction this section concerns image 141 processing. As is known, a linear image enhancement technique can improve image visual quality but cannot preserve 143 its true values, whereas nonlinear image restoration can obtain 144 an improved and value-preserved image, but is time-145 consuming. Here, we propose an approach derived from the Poisson MAP superresolution algorithm. This approach is in-147 corporated with the procedure of FEM-based image reconstruction to obtain resolution-enhanced images. In this sec-149 tion, conventional image processing is first addressed, then a novel approach to implementing HP filters is derived, and finally our proposed approach is integrated with DOT image reconstruction.

2.1 Conventional Image Processing

In image processing, image enhancement is always used to 155 improve image visual quality. The techniques of contrast enhancement, histogram equalization, and HPF are usually 157 adopted. Contrast enhancement conducts an operation to expand the contrast of features of interest. The procedure of 159 histogram equalization, basically, transforms the histogram 160

139

140



Fig. 1 Two LPF-combined HP filters (a) $\delta - g_2$ and (b) $g_1 - g_2$, and (c) a wavelet-like filter, where δ is the delta function.

161 distribution of an image into an output image with an equal 162 number of pixels at each gray level. This causes a ragged 163 histogram to become flat. HPF is exactly a transfer function 164 with a unit at dc frequency and higher gains associated with 165 larger frequencies. Usually, edge enhancement can be re-166 garded as an alternative to HPF, sharpening the edge but with 167 overshoot.

As described in some image-processing monographs,^{35–37}
the HPF applied to improve image quality follows some routime steps such as the so-called high-frequency emphasis filtering,

$$H_{\rm hef} = a + bH_{\rm hp},\tag{1}$$

 and histogram equalization, where H_{hef} denotes high- frequency emphasis filtering in the frequency domain (FD), H_{hp} is an HP filter, *a* is an offset, and *b* is a weighting number (usually, b > a). Note that the capital to represent a filter is the corresponding frequency function. Therefore, $1-H_{lp}$ is adopted, a complementary filter to low-pass filtering in the FD for an HP filter.

180 To achieve both improved image visual quality and pre-181 serve true value distribution in biological applications, we ar-182 gue in this paper that an in-depth investigation is required to 183 cope with the challenges of emergent biomedical imaging mo-184 dalities. Before proposing our scheme, we first define HP, 185 filters and classify them for the convenience of following dis-186 cussions. There are two types of $h_{\rm hp}$ to be performed, respec-187 tively. One is a differential filter through the combination of 188 two LPFs, and the other is an intrinsic (wavelet-like) HP filter. 189 It is sensible that an HP filter can be described as

$$h_{\rm hp} = h_{\rm lp1} - h_{\rm lp2}, \qquad (2)$$

 where h_{1p1} and h_{1p2} denote LPFs. More precisely, this repre- sents a narrow full width at half maximum (FWHM) LPF with a larger amplitude (A_1) subtracted by a broad FWHM LPF with a smaller amplitude (A_2). Both HP filters must com-ply with two rules of thumb³⁷ as follows:

196
$$H_{hp}(0) = A_1 - A_2$$
 and $(H_{hp})_{max} \le A_1$. (3)

197 The difference between these filters is that the $h_{\rm lpl}$ of the 198 former rolls off faster than that of the latter. In this study,

Gaussian functions with various standard deviations (σ) are ¹⁹⁹ employed for $h_{\rm lp}$. Thus, the HP filter shown in Fig. 1 can be 200 formulated as 201

$$h_{\rm hp}(r) = g_1(r) - g_2(r),$$
 (4) 202

where
$$g_1(r) = [A_1/(2\pi\sigma_1^2)^{1/2}] \exp(-r^2/2\sigma_1^2)$$
 and $g_2(r)$ 203
= $[A_2/(2\pi\sigma_2^2)^{1/2}] \exp(-r^2/2\sigma_2^2)$, respectively, and $\sigma_1 < \sigma_2$. 204

2.1.1 LPF-combined HP filter ($\sigma_1 < \sigma_2$) **205**

This filter is usually determined with a smaller $\sigma_1 (\ll 1)$, as 206 shown in Fig. 1(a); this can also be determined with a larger 207 σ_1 , as shown in Fig. 1(b). If we let σ_1 approach zero, $h_{\rm hp1}$ 208 narrows further to an impulse, then Eq. (4) can be expressed 209 in the frequency domain as $1-H_{\rm lp}$.

A dilated wavelet-like function expressed as

$$\psi_a(r) = \frac{1}{\sqrt{3a^4/\pi}} \left(1 - \frac{r^2}{a^2} \right) \exp\left(-\frac{r^2}{2a^2} \right),$$
 (5) 213

212

where *a* is a dilated factor, as depicted in Fig. 1(c), and can be **214** used as an HP filter. **215**

2.1.3 Negative-oriented Laplacian HP filter 216

Alternatively, a 3×3 negative-orientated Laplacian edge op- 217 erator 218

$$\begin{bmatrix} 0 & -1 & 0 \\ -1 & 4 & -1 \\ 0 & -1 & 0 \end{bmatrix}$$
(6)
219

in a form similar to a wavelet is also considered and em- 220 ployed as an HP filter is this study. 221

As can be seen in Fig. 1, a wavelet-like HP filter has sharp 222 sidelobes rather than a LPF-combined HP filter. 223

172

PROOF COPY 032802JBO

Pan et al.: Highly resolved diffuse optical tomography: a systematic approach...

224 2.2 Novel Approach Implementing HPF for Optical 225 Tomography

 To find the fundamental theoretical basis to explain our pro- posed novel approach, an attempt to derive the value- preserving HPF technique begins with the Poisson MAP su- perresolution algorithm. Mathematically, the algorithm³⁸⁻⁴⁰ is **230** given as

$$\hat{f}_n = \hat{f}_{n-1} \exp\left[\left(\frac{g}{\hat{f}_{n-1} \otimes h} - 1\right) * h\right] = \hat{f}_{n-1}C, \quad n = 1, 2, \dots, N,$$
(7)

232 where

231

233
$$C = \exp\left[\left(\frac{g}{\hat{f}_{n-1} \otimes h} - 1\right) * h\right], \qquad (8)$$

 is regarded as the correction term during the iterative restora- tion progress; \otimes represents a convolution; * represents cor- relation; *h* denotes the point spread function (PSF); *g* is the observed image; and the subscript *n* is the number of iteration. Additionally, \hat{f}_0 defined as *g* is the initial guess of iteration, and \hat{f}_N is the final superresolved image. In terms of the op- eration of Poisson MAP, it is an iterative algorithm, where successive estimate of the restored image is obtained through the multiplication of current estimate by such a quantity close to one that is a function of the interpolated image divided by a convolution of the current estimate with the PSF. Using Taylor series expansion, Eq. (7) can be expanded to be ap-proximate as

$$\hat{f}_{n} \sim \hat{f}_{n-1} \left[1 + \left(\frac{g}{\hat{f}_{n-1} \otimes h} - 1 \right) * h \right] = \hat{f}_{n-1} + \hat{f}_{n-1} \left[\left(\frac{g}{\hat{f}_{n-1} \otimes h} - 1 \right) * h \right] = \hat{f}_{n-1} + \Delta \hat{f}_{n-1},$$
(9)

 where the intermediate obtained image \hat{f}_n can be expressed as adding the previous one \hat{f}_{n-1} with a correction increment $\Delta \hat{f}_{n-1}$. It is nontrivial to further explore the correction incre-**252** ment

$$\Delta \hat{f}_{n-1} = \hat{f}_{n-1} \left[\left(\frac{g}{\hat{f}_{n-1} \otimes h} - 1 \right) * h \right]$$
$$= \hat{f}_{n-1} \left[\left(\frac{g - \hat{f}_{n-1} \otimes h}{\hat{f}_{n-1} \otimes h} \right) * h \right], \tag{10}$$

253

254

 where $g - \hat{f}_{n-1} \otimes h$ can be reduced to an increment $\Delta \hat{f}_{n-1}$ in a condition of decreasing correction rate. Assuming $\hat{f}_{n-1} \otimes h$ ap- proaches a constant as \hat{f}_{n-1} has a simple distribution. Thus, Eq. (10) is approximated to

$$\Delta \hat{f}_{n-1} \sim \frac{\hat{f}_{n-1}}{\hat{f}_{n-1} \otimes h} (\Delta \hat{f}_{n-1} * h).$$
(11) 259

Through point-by-point multiplying both sides of Eq. (11) 260 with $\Delta \hat{f}_{n-1}$, we obtain 261

$$(\Delta \hat{f}_{n-1})(\Delta \hat{f}_{n-1}) = \frac{\hat{f}_{n-1}}{\hat{f}_{n-1} \otimes h} (\Delta \hat{f}_{n-1} * h) \Delta \hat{f}_{n-1}, \qquad (12)$$
262

and then reorganize Eq. (12) to yield

$$\hat{f}_{n-1} \otimes h = \hat{f}_{n-1} \left[\frac{(\Delta \hat{f}_{n-1} * h) \Delta \hat{f}_{n-1}}{(\Delta \hat{f}_{n-1}) (\Delta \hat{f}_{n-1})} \right].$$
(13) 264

263

276

As defined previously, h is the PSF like an LPF, and a Gauss- 265 ian function is employed in the study. Additionally, the opera- 266 tion of correlation is equivalent to take a convolution due to 267 the symmetry of function h. Therefore, Eq. (13) can be de- 268 rived to the following equation with the HPF definition 269

$$\hat{f}_{n-1} \otimes h_{\rm hp} = \hat{f}_{n-1} \otimes h_{\rm lp1} - \hat{f}_{n-1} \otimes h_{\rm lp2}$$
 270

$$=\hat{f}_{n-1}\left[\frac{(\Delta\hat{f}_{n-1}\otimes h_{\rm hp})\Delta\hat{f}_{n-1}}{(\Delta\hat{f}_{n-1})(\Delta\hat{f}_{n-1})}\right]$$
271

$$= \frac{(\Delta f_{n-1} \otimes h_{\rm hp})\Delta f_{n-1}}{(\Delta \hat{f}_{n-1})(\Delta \hat{f}_{n-1})/\hat{f}_{n-1}}.$$
 (14)

We here consider the quantity obtained through the convolu- 273 tion of an image and HPF as a new correction increment, i.e., 274

$$\Delta \hat{f}_{n-1} \sim \hat{f}_{n-1} \otimes h_{\rm hp}. \tag{15} \ \textbf{275}$$

Thus, Eq. (15) is equivalent to

$$\Delta \hat{f}_{n-1} = \frac{(\Delta \hat{f}_{n-1} \otimes h_{\rm hp}) \Delta \hat{f}_{n-1}}{(\Delta \hat{f}_{n-1}) (\Delta \hat{f}_{n-1}) / \hat{f}_{n-1}}.$$
(16)
277

To further simplify Eq. (16) for numerical evaluation, we as- **278** sume that the denominator is a positive number relative to **279** $\Delta \hat{f}_{n-1}$, and finally get an approximate solution of the correc- **280** tion increment for using HPF, as follows **281**

$$\Delta \hat{f}_{n-1} = \frac{\langle \Delta \hat{f}_{n-1} | h_{\rm hp} | \Delta \hat{f}_{n-1} \rangle}{\langle \Delta \hat{f}_{n-1} | \Delta \hat{f}_{n-1} \rangle / \hat{f}_{n-1}} \sim \frac{\langle \Delta \hat{f}_{n-1} | h_{\rm hp} | \Delta \hat{f}_{n-1} \rangle}{w \| \Delta \hat{f}_{n-1} \|}, \quad (17)$$
282

where the denominator is simplified with the norm of $\Delta \hat{f}_{n-1}$ 283 multiplied by *w*, a weight number 10, used in computation as 284 well as herein the symbols $\langle \cdot |$ and $| \cdot \rangle$ stand for the state and 285 $\langle z | x \text{ and } x | y \rangle$ represent the operations of a point-by-point 286 product (*z* and *x*) and a convolution (*x* and *y*), respectively, 287 resulting in the other state. In considering Eq. (17) used in the 288 computation of NIR DOT imaging, heterogeneities are treated 289 as a perturbation to homogeneous background for a phantom, 290 and incremental values of both absorption and scattering co-291 efficients are estimated from a projection of a high-frequency 292 enhancement to the original increment. 293



Fig. 2 Flowchart of NIR DOT image reconstruction incorporated with the HPF approach.

294 2.3 NIR DOT Image Reconstruction Incorporating 295 with Novel Approach

296 Compared with other medical imaging modalities, NIR imag-297 ing requires the solution of an inverse problem. In NIR DOT298 imaging, the fundamental equation governing the propagation299 of light in biological tissues is the Boltzmann transport equa-300 tion (BTE) to model the optical characteristics of the scatter-301 ing and absorption.

The BTE is an integrodifferential equation, so it is rather 303 difficult to obtain solutions to the BTE under general condi-304 tions. With the use of approximation techniques by assuming 305 the experimental material or tissues have highly scattering 306 properties and that the input radiance is isotropic and modu-307 lated under a 1-GHz frequency, the BTE can be reduced to an 308 easily solvable form of the diffusion approximation. In NIR 309 imaging, mappings of the absorption and/or scattering coeffi-310 cients can be evaluated by using an FEM to invert the diffu-311 sion approximation. The FEM-based image reconstruction in 312 the dc domain is concluded with the following equations. 313 More derivation details can be found in Ref. 41.

As described previously, the physical process can be de-315 duced from a diffusion equation:

316
$$\nabla \cdot D \nabla \Phi(r,\omega) - \left(\mu_a - \frac{i\omega}{c}\right) \Phi(r,\omega) = -S(r,\omega), \quad (18)$$

 where $S(r, \omega)$ and Φ denote the source and the radiance, re- spectively; and μ_a , c, and D are the absorption coefficient, the wave speed in the medium, and the diffusion coefficient, re- spectively. To solve Eq. (18), the boundary condition $-D\nabla\Phi\cdot\hat{n}=\alpha\Phi$ (flux in fact) and the FEM are applied. Since only dc data are considered, ω is set as a null; i.e., the imagi- nary part is to vanish from the subsequent equations. Thus, the following discrete equations in a matrix form,

325

$$\begin{bmatrix}
A_{ij}^{bb} - \alpha B_{ij}^{bb} & A_{ij}^{bl} \\
A_{ij}^{lb} & A_{ij}^{ll}
\end{bmatrix} \quad \left\{ \Phi_{j}^{b} \\
\Phi_{j}^{l} \right\} = \left\{ \begin{array}{c}
C_{i}^{b} \\
C_{i}^{l} \\
C_{i}^{l} \\
\end{array} \right\},$$
326
optical-property matrix radiance matrix source matrix (19)

 can be obtained. Obviously, the forward solution, Φ , can be evaluated through Eq. (19). In terms of the physical process, the radiance matrix is quantitatively and qualitatively depen- dent on the source matrix and the optical-property matrix, respectively, where the optical-property matrix is the inertia of the material in spite of relating to the wavelength. Further- more, the following two equations can be derived for the computation of image reconstruction, i.e.,

$$\begin{bmatrix} A_{bb} - \alpha B_{bb} & A_{bI} \\ A_{Ib} & A_{II} \end{bmatrix} \begin{cases} \frac{\partial \Phi_b}{\partial D_k} \\ \frac{\partial \Phi_I}{\partial D_k} \end{cases} = \begin{bmatrix} -\frac{\partial A_{bb}}{\partial D_k} & -\frac{\partial A_{bI}}{\partial D_k} \\ -\frac{\partial A_{Ib}}{\partial D_k} & -\frac{\partial A_{II}}{\partial D_k} \end{bmatrix} \begin{cases} \Phi_b \\ \Phi_I \end{cases}$$

$$335$$

$$+ \begin{cases} \frac{\partial C_b}{\partial D_k} \\ \frac{\partial C_I}{\partial D_k} \end{cases}, \qquad (20)$$

337

and

$$\begin{bmatrix} A_{bb} - \alpha B_{bb} & A_{bI} \\ A_{Ib} & A_{II} \end{bmatrix} \begin{cases} \frac{\partial \Phi_b}{\partial \mu_l} \\ \frac{\partial \Phi_I}{\partial \mu_l} \end{cases} = \begin{bmatrix} -\frac{\partial A_{bb}}{\partial \mu_l} & -\frac{\partial A_{bI}}{\partial \mu_l} \\ -\frac{\partial A_{Ib}}{\partial \mu_l} & -\frac{\partial A_{II}}{\partial \mu_l} \end{bmatrix} \begin{pmatrix} \Phi_b \\ \Phi_I \end{pmatrix}$$

$$338$$

$$+ \begin{cases} \frac{\partial C_b}{\partial \mu_l} \\ \frac{\partial C_I}{\partial \mu_l} \end{cases} . (21)$$

where the superscripts *I* and *b* denote interior and boundary 340 nodes, and D_k for $k=1,2,\ldots,K$ and μ_l for $l=1,2,\ldots,L$ are 341 the reconstruction parameters for the optical property profile. 342 For the inverse problem to update absorption/scattering coef- 343 ficients, the partial differentiation of boundary radiance to the 344 parameters of interest, $\partial \Phi_b / \partial \mu_l$ or $\partial \Phi_b / \partial D_k$, must be ob- 345 tained from Eqs. (20) and (21). The Newton-Raphson tech- 346 nique regularized by a Levenberg-Marquardt algorithm and 347 with the Tikhonov regularization parameter is adopted to it- 348 eratively update the diffusion and absorption coefficients, i.e., 349

$$(\mathbf{J}^T \mathbf{J} + \lambda I) \Delta \chi = \mathbf{J}^T (\Phi^o - \Phi^c) = \mathbf{J}^T \Delta \Phi, \qquad (22) \ \mathbf{350}$$

where Jacobian matrix **J** denotes $\mathbf{J}(\partial \Phi_b / \partial D_k, \partial \Phi_b / \partial \mu_l)$, $\Delta \chi$ 351 means $\Delta \chi (\Delta D_k, \Delta \mu_l)$, and λ is a Tikhonov regularization parameter of the Jacobian matrix. As described, this inversion 353 generally requires the construction of the Jacobian matrix; 354 actually, the Jacobian represents a highly underdetermined 355 system of equations. Although it is possible to obtain a leastsquares solution to underdetermined systems of equations, the 357 resulting images are usually inaccurate relating to inferior resolution. The procedure of FEM-based image reconstruction in 359 the dc domain is illustrated in Fig. 2. As indicated, the proposed approach is merely implemented once, subsequently 361 posterior to each iteration without altering the original FEM 362



Fig. 3 Schematic diagram for the dimensions of four different test cases in simulation. (a) to (d) are cases 1 to 4, respectively, where *R* is radius in millimeters.

³⁶³ modeling. The following section illustrates the comparison
364 and effectiveness of the incorporated resolution-enhanced
365 schemes.

366 3 Results and Discussion

367 The phantoms employed to justify our proposed technique 368 incorporate two or three inclusions with various sizes, loca-**369** tions, and separations, illustrated in Fig. 3, where R denotes 370 the radius in millimeters. In this paper, four HPFs and four 371 phantom cases were performed. The numerical simulations of 372 multiinclusion phantoms provide further information concern-373 ing the spatial resolution (separation, size, and location) and 374 the contrast resolution beyond that of the single-inclusion **375** case. Of the phantom, the background absorption (μ_a) and **376** reduced scattering (μ'_s) values are about 0.0025 and $377 0.25 \text{ mm}^{-1}$, respectively, while the maximum absorption and **378** reduced scattering for the inclusion are 0.025 and 2.5 mm^{-1} , 379 if we assume the contrast ratio of the inclusion to background 380 is 10:1, because high contrast results in much more overlap-**381** ping effects than low contrast, although a contrast of 2 to 10 382 was used throughout other published works.

383 As depicted in Fig. 3, cases 1 and 2 and cases 3 and 4, 384 respectively, have three and two inclusions separated by a 385 similar distance but of different sizes. As the separation reso-386 lution of inclusions is examined, several (two or three) em-387 bedded inclusions are necessary, and different inclusion sizes 388 are considered as well. To test the limitation of each HPF **389** employed here, the phantoms of cases 1 and 4 with larger **390** inclusions and closer to the phantom center were designed, **391** compared with case 2 and case 3 designs. For the convenience 392 of discussion, we denote M0 to M4 as the reconstructions **393** with the schemes using nonfiltering, $\delta - g2(\sigma_2 = 1.5)$, g1 **394** $-g2(\sigma_1=0.75, \sigma_2=1.5)$, wavelet (a=0.5), and Laplacian 395 HPF in their 2-D form, respectively. Currently, absorption-396 coefficient images are presented for our cw image reconstruc-**397** tion algorithm.

 In FEM-based image reconstruction, the homogeneous background ($\mu_a = 0.0025 \text{ mm}^{-1}$, $\mu'_s = 0.25 \text{ mm}^{-1}$) was adopted as an initial guess. For both the forward and inverse processes, 256 elements and 257 nodes were used, associated with a desktop PC with a 3.6-GHz CPU and 4 Gbytes of RAM, respectively. Thirty iteration assignments were em- ployed for each case as the normalized increasing rate, i.e., mean value of $|(\Phi_{n+1}-\Phi_n)/\Phi_n|^2$, reaches less than 10^{-2} , where each iteration takes about 2 min. Meanwhile, the absorption- and diffusion-coefficient images were updated concurrently in spite of the fact that reconstruction began ⁴⁰⁸ from a homogeneous condition and only the acquired dc data ⁴⁰⁹ were employed. ⁴¹⁰

First, a qualitative investigation of the reconstruction per- 411 formance of each case is presented in Sec. 3.1. Following this 412 in Sec. 3.2, we describe quantitative performance measures 413 for various HPFs for a range of resolutions including separa- 414 tion, size, contrast, and location. Finally, we further investi- 415 gate and discuss the significance of the proposed measures in 416 Sec. 3.3. 417

3.1 Examples Illustration 418

419

434

3.1.1 Case 1

Figure 4 shows a set of reconstructed absorption-coefficient 420 images [Figs. 4(a)-4(e)] and quantitative information [Figs. 421 4(f)-4(j)] for the images along with their corresponding cir- 422 cular transaction profiles. Comparing the reconstructed ab- 423 sorption images, it is obvious that all of the reconstructions 424 show roughly correct images of the inclusions and all of the 425 reconstruction techniques can highly resolve images and sepa- 426 rate inclusions, except the result using M0. However, the M1 427 to M4 schemes underestimate the computed absorption coef- 428 ficients of the inclusions. For further inspection, M4 generated 429 highly ringing artifacts between inclusions. At this phase, it is 430 not easy to speculate about the causes of such artifacts that 431 might be referred to as 'false' inclusions and concluded as a 432 wrong judgment.

3.1.2 Case 2

It can be found that compared with only the image reconstruc- 435 tion employed, considerable improvement is observed in the 436 reconstructed images, as illustrated in Figs. 5(a)-5(e) when 437 the HPF approach is invoked. Evidently, M0 retains highly 438 blurred inclusions, while the other reconstruction schemes can 439 better differentiate inclusions, and the M4 scheme overesti-440 mated the absorption coefficients. Again, ringing artifacts are 441 produced surrounding inclusions and their optical properties 442 are lower than background levels, as depicted in Figs. 443 5(f)-5(j).

From the results for cases 1 and 2 note that schemes with 445 filtering can discriminate even small size inclusions, whereas 446 scheme M0 cannot meet even the basic requirements of image 447 reconstruction, especially for small inclusions (case 2). 448

Pan et al.: Highly resolved diffuse optical tomography: a systematic approach...



Fig. 4 Case 1, 2-D reconstructed absorption images (a) without HPF (M0) and (b) to (e) with M1, M2, M3, M4 filtering, respectively; (f) to (j) 1-D sectional profiles corresponding to (a) to (e), where the solid lines are the designed and the dotted lines represent the reconstructed schemes.

449 3.1.3 Case 3

 Compared with previous two cases, this case was designed as a phantom with three smaller inclusions. Several improved images were obtained by using appropriate filtering, as shown in Figs. 6(b)-6(e). Likewise, M2 resulted in a worse-resolved image than the others with HP filtering. Negative artifacts occurred in each reconstructed image, as depicted in Figs. 6(g)-6(j). It is well noted that M4 overestimated the inclusion amplitudes, which yields a higher inclusion-to-background contrast.

459 3.1.4 Case 4

 In this highly challenging case, a phantom with two closest- separation inclusions was designed. As shown in Figs. 7(a)-7(e), all reconstructed images underestimated inclusions, and offered relatively poor resolution for two separate inclu- sions. This is rather competitive for these employed filters. Based on a quantitative comparison, as depicted in Figs. 7(i) and 7(j), the M3 and M4 schemes demonstrate better resolution discrimination to separate longer and closer inclusions **467** in comparison with case 3. **468**

From the results of cases 3 and 4 for a phantom with 469 inclusions of both small size and close separation, it can be 470 concluded that the wavelet-like HP filtering (M3) demon- 471 strates the best spatial-resolution capability to the inclusions. 472

This evidently shows that the enhancement of reconstruc- 473 tion through the incorporation of our proposed HPF approach 474 can effectively improve computed images. As already illus- 475 trated, the wavelet-like HP filtering schemes (M3 and M4) 476 further yield better results than the LPF-combined HP filtering 477 schemes (M1 and M2). In the aspects of sensitivity and sta- 478 bility of evaluation, the M3 scheme yielded results closest to 479 the true absorption property compared to the other schemes. 480 However, scheme M4 visually characterizes the inclusion-to- 481 background contrast best. 482



Fig. 5 Reconstructed case 2 images, with (a) to (j) as described for Fig. 4.

Pan et al.: Highly resolved diffuse optical tomography: a systematic approach...



Fig. 6 Reconstructed case 3 images, with (a) to (j) as described for Fig. 4.

483 3.2 Performance Investigation

 In terms of the optical properties within the inclusion and the background, note that the image reconstruction not only pur- sues qualitative correctness but also obtains favorably quanti- tative information about the optical properties of either the inclusions or the background. The parameters of interest, such as size, contrast, and location variations associated with image quantification measures are most frequently investigated and discussed.

492 Several measures^{42–45} have been used to evaluate the per-493 formance of the NIR imaging algorithms or systems. Song et 494 al.⁴² used the contrast-to-noise ratio (CNR), which is defined 495 as the difference between the region of interest (ROI) and the 496 background region values of the optical properties divided by 497 the average variation in the background, where one inclusion 498 was considered. Furthermore, informative works (Pogue et 499 al.⁴⁴) provided an overview of the three major methods uti-500 lized for image analysis in the imaging science and medical 501 physics communities, which lie in the areas of the spatial resolution, the contrast detail (CD) analysis, and human perception of images. Briefly, the first one relates to the modula-503 tion transfer function (MTF) profile; the second one, to the 504 CD curve (contrast versus size) obtained by human observa-505 tion; and the last concerns the receiver operating characteristic 506 (ROC) curve and location receiver operating characteristic 507 curve (LROC) obtained by the human observer detection of 508 abnormalities. In our cases, however, several inclusions in the 509 background of a phantom were considered and further investigations of the contrast, size, separation, and location were 511 conducted so that it is essential that these four terms are respectively defined and discussed. 513

To provide a quantitative assessment for these recon- 514 structed images through using various HPF approaches, we 515 designate and formulate four measures over the ROI for the 516 evaluation of these filtering schemes, and these measures are 517 normalized to be in-between a null and unit with the ratio of 518 the reconstructed to the original images. To interpret these 519 measures in detail, we describe them using Fig. 8, where 520



Fig. 7 Reconstructed case 4 images, with (a) to (j) as described for Fig. 4.

and



Fig. 8 Diagram for the explanation of defined measures.

521 1D_ROI is chosen as the line segment between the two out-522 most nodes of the inclusions, and 2D ROI is the possibly 523 smallest region around and/or covering inclusions. Moreover, 524 although it is usually a difficult task to define a separation 525 resolution, here we regard an inclusion and a separation as a 526 bump and a cave, respectively.

527 3.2.1 *Contrast resolution* (*R*^{1D,2D}_{cont.})

528 The measure $R_{\text{cont.}}^{1\text{D,2D}}$ is defined to evaluate the resolution on 529 the contrast of optical property values with the inclusions to 530 the background region especially between inclusions.

531
$$R_{\text{cont.}}^{1\text{D},2\text{D}} = \frac{(\overline{\max}^{\Delta \text{incl.}}/\overline{\min}^{\Delta \text{incl.}})_{\text{reconstruction}}}{(\overline{\max}^{\Delta \text{incl.}}/\overline{\min}^{\Delta \text{incl.}})_{\text{original}}},$$
(23)

532 and

533

$$R_{\text{cont.}}^{1D,2D} = 2 - R_{\text{cont.}}^{1D,2D}, \text{ if } 1 < R_{\text{cont.}}^{1D,2D} < 2,$$
 (24)

534 where $\overline{\max}$ and $\overline{\min}$ denote the average of maxima and 535 minima over all the selected regions as the superscripts **536** (Δ incl. and $\Delta_{\overline{\text{incl}}}$). Also incl. and incl correspondingly repre-537 sent inclusions and complementary inclusions as well as **538** Δ incl. and $\Delta_{\overline{\text{incl}}}$ are chosen with several nodes around central 539 area of incl. and incl.

540 3.2.2 Separation resolution $(R_{sep}^{1D,2D})$ **541** The measure $R_{sep}^{1D,2D}$ is designed to evaluate the resolution on **542** the separation between inclusions.

543
$$R_{\text{sep}}^{\text{1D,2D}} = \left\{ \left[1 - \frac{(\text{MSE}^{\overline{\text{incl.}}})_{\text{Recon.2Ori.}}}{(\text{MSE}^{\overline{\text{incl.}}})_{\text{Ori.2Baseline}}} \right] R_{\text{cont.}}^{\text{1D,2D}} \right\}^{1/2}$$

544
$$\equiv (Ro_{\text{sep}}^{\text{1D,2D}} \times R_{\text{cont.}}^{\text{1D,2D}})^{1/2}, \qquad (25)$$

545 where MSE is the mean square error over all the selected 546 region as the superscript (incl) and Baseline is used with **547** 0.025 mm⁻¹; Ori.2Baseline is, here, 0.0025 to 0.025 mm⁻¹, 548 and Recon.20ri. is the reconstructed value in the region incl **549** to 0.0025 mm^{-1} .

550 3.2.3 Size resolution $(R_{size}^{1D,2D})$

551 The measure $R_{\text{size}}^{1\text{D,2D}}$ is designed to evaluate the resolution on **552** the size over all inclusions.

$$R_{\text{size}}^{1\text{D},2\text{D}} = \left\{ \left[1 - \frac{(\text{MSE}^{\text{incl.}})_{\text{Recon.2Ori.}}}{(\text{MSE}^{\text{incl.}})_{\text{Ori.2baseline}}} \right] R_{\text{cont.}}^{1\text{D},2\text{D}} \right\}^{1/2}$$

$$= (R_0 e^{1\text{D},2\text{D}} \times R^{1\text{D},2\text{D}})^{1/2}$$
(26) ----

$$\equiv (Ro_{\text{size}}^{1D,2D} \times R_{\text{cont.}}^{1D,2D})^{1/2}, \qquad (26) \ 554$$

where MSE is over the selected region incl. and baseline is 555 used with 0.0025 mm⁻¹. Note that the FWHM usually oper- 556 ated manually and subjectively is not adopted for the evalua- 557 tion of inclusion size. Here, attempt to automatically estimate 558 this resolution with the idea of the capacity rate for the term 559 of interest. 560

3.2.4 Location resolution
$$(R_{locat}^{1D,2D})$$
 561

The measure $R_{\text{locat}}^{1D,2D}$ is defined to evaluate the resolution on **562** the location over all inclusions. **563**

$$R_{\text{locat}}^{\text{1D,2D}} = \left[1 - \frac{(\overline{\text{CM}^{\text{incl.}}})_{\text{Reconstruction}}}{(\overline{\text{CM}^{\text{incl.}}})_{\text{Original}}}R_{\text{cont.}}^{\text{1D,2D}}\right]^{1/2}$$
564

$$\equiv (Ro_{\text{locat}}^{1D,2D} \times R_{\text{cont.}}^{1D,2D})^{1/2}, \qquad (27) \ 565$$

566

$$Ro_{\text{locat}}^{1D,2D} = 2 - Ro_{\text{locat}}^{1D,2D}, \text{ if } 1 < Ro_{\text{locat}}^{1D,2D} < 2,$$
 (28) 567

where \overline{CM} is the average of the center of mass over all the 568

selected region as the superscript (incl.). 569 Note that the resolutions, $R_{sep}^{1D,2D}$, $R_{size}^{1D,2D}$, and $R_{local}^{1D,2D}$ in- 570 clude a multiplication operation by $R_{cont.}^{1D,2D}$ to avoid the low- 571 contrast reconstruction with high-consistent inclusions or 572 complements. Furthermore, it is expected that the resolution is 573 higher because the R value approaching more closely to a 574 unit.

Based on the preceding definitions, the evaluated resolu- 576 tions of 1-D profiles and 2-D images with multiinclusions are 577 listed in Tables 1-4 for each phantom case, respectively. The 578 quantities for $R_{\text{sep}}^{\text{1D,2D}}$, $R_{\text{size}}^{\text{1D,2D}}$, and $R_{\text{local}}^{\text{1D,2D}}$ are small because 579 the defined measures are quite strict. For overall cases, it is 580 found that location resolution is above 0.95 prior to $Ro_{locat}^{1D,2D}$ 581 multiplied by $R_{\text{cont.}}^{1D,2D}$, and less difference exists between these 582 two; whereas the contrast, separation, and size resolution have 583 comparable differences. Figures 9-12 illustrate comparisons 584 of the separation, size, and contrast resolutions among various 585 HP filters to clarify our observation. Overall, the resolutions 586 obtained in cases 1 to 3 are better than those in case 4, as 587 expected. Basically, our approach demonstrates the effective- 588 ness of separation and size resolution rather than contrast res- 589 olution. A discussion of each individual case follows. 590

Figure 9(a) shows the resolution performance of schemes M3, 592 M1, M2, M4, and M0, respectively. The results show that the 593 M4 scheme yielded false inclusions. For the 2-D condition, 594 the revealed performance is similar to that in the 1-D condi- 595 tion except for the M0 and M4 schemes. Obviously, this 596 evaluation is consistent with that for Fig. 4 based on the visual 597 perception. 598

3.2.6 Case 2

Figure 10(a) shows the performance of schemes M4, M1, M3, 600 M2, and M0, respectively. The performance in the 2-D con- 601

| | | | 1-D | | 2-D | | | | | |
|----|----------------|----------------|----------|----------------|----------------|----------------|----------|------------------|------------------|--|
| | Sep. 0 Sep. | Size 0 Size | Contrast | Loc. 0 Loc. | Sep. 0 Sep. | Size 0 Size | Contrast | Locx. 0 Locx. | Locy. 0 Locy. | |
| MO | 0.46 | 0.93 | 0.12 | 1.00 | 0.92 | 0.94 | 0.58 | 1.00 | 0.98 | |
| | 0.23 | 0.33 | | 0.35 | 0.73 | 0.74 | | 0.76 | 0.76 | |
| M1 | 0.69 | 0.66 | 0.31 | 0.95 | 0.99 | 0.66 | 0.68 | 0.96 | 0.99 | |
| | 0.46 | 0.45 | | 0.54 | 0.82 | 0.67 | | 0.81 | 0.82 | |
| M2 | 0.70 | 0.63 | 0.30 | 0.98 | 0.99 | 0.65 | 0.54 | 1.00 | 1.00 | |
| | 0.46 | 0.44 | | 0.55 | 0.73 | 0.59 | | 0.73 | 0.73 | |
| M3 | 0.84 | 0.85 | 0.36 | 1.00 | 0.99 | 0.78 | 0.87 | 1.00 | 0.99 | |
| | 0.55 | 0.55 | | 0.60 | 0.93 | 0.82 | | 0.93 | 0.93 | |
| M4 | 0.74 | 0.71 | 0.15 | 1.00 | 0.97 | 0.65 | 0.80 | 1.00 | 0.98 | |
| | 0.33 | 0.32 | | 0.38 | 0.88 | 0.72 | | 0.89 | 0.88 | |

Table 1 Case 1 separation, size, contrast, and location resolutions for various filtering on 1-D and 2-D conditions.

602 dition is similar to that in the 1-D condition except for the M4 603 scheme. Unfortunately, a negative value occurs in the 2-D 604 condition [Fig. 10(b)], which means that M4 highly overesti-605 mated the inclusion size. highly overestimated effect of M4 attenuates the measure values. Basically, other filtering schemes obtain the measure distribution as expected. 611

Generally speaking, the M1 and M3 schemes perform bet- 612 ter than the M2 and M4 schemes on either of the defined 613 measures or the visual perception for cases 2 and 3. 614

606 3.2.7 Case 3

607 Obviously, the performance of case 3 is similar to that of case **608** 2, shown as Fig. 11. For the same reasons as in case 2, the

Table 2 Case 2 separation, size, contrast, and location resolutions for various filtering on 1-D and 2-D conditions.

| | 1-D | | | | 2-D | | | | | |
|----|----------------|----------------|----------|----------------|----------------|----------------|----------|------------------|------------------|--|
| | Sep. 0 Sep. | Size O Size | Contrast | Loc. 0 Loc. | Sep. 0 Sep. | Size 0 Size | Contrast | Locx. 0 Locx. | Locy. 0 Locy. | |
| MO | 0.84 | 0.85 | 0.10 | 1.00 | 0.95 | 0.87 | 0.24 | 1.00 | 0.98 | |
| | 0.29 | 0.29 | | 0.32 | 0.48 | 0.46 | | 0.49 | 0.49 | |
| M1 | 0.94 | 0.95 | 0.33 | 0.98 | 0.95 | 0.89 | 0.95 | 0.96 | 0.99 | |
| | 0.56 | 0.56 | | 0.57 | 0.95 | 0.92 | | 0.95 | 0.97 | |
| M2 | 0.86 | 0.91 | 0.17 | 0.99 | 0.97 | 0.90 | 0.83 | 0.99 | 0.96 | |
| | 0.38 | 0.39 | | 0.41 | 0.90 | 0.86 | | 0.91 | 0.89 | |
| M3 | 0.88 | 0.94 | 0.17 | 1.00 | 0.97 | 0.95 | 0.70 | 1.00 | 0.99 | |
| | 0.39 | 0.41 | | 0.42 | 0.83 | 0.82 | | 0.84 | 0.84 | |
| M4 | 0.87 | 0.87 | 0.40 | 1.00 | 0.74 | -0.07 | 0.50 | 1.00 | 0.98 | |
| | 0.59 | 0.59 | | 0.63 | 0.61 | -0.19 | | 0.71 | 0.70 | |

Table 3 Case 3 separation, size, contrast, and location resolutions for various filtering on 1-D and 2-D conditions.

| | | 1 | -D | | 2-D | | | | | |
|----|----------------|----------------|----------|----------------|----------------|----------------|----------|------------------|------------------|--|
| | Sep. 0 Sep. | Size O Size | Contrast | Loc. 0 Loc. | Sep. 0 Sep. | Size 0 Size | Contrast | Locx. 0 Locx. | Locy. 0 Locy. | |
| MO | 0.85 | 0.74 | 0.10 | 1.00 | 0.98 | 0.75 | 0.18 | 1.00 | 1.00 | |
| | 0.29 | 0.27 | | 0.32 | 0.42 | 0.36 | | 0.42 | 0.42 | |
| M1 | 0.84 | 0.95 | 0.20 | 0.99 | 0.92 | 0.97 | 0.40 | 0.98 | 0.98 | |
| | 0.41 | 0.43 | | 0.44 | 0.61 | 0.62 | | 0.62 | 0.62 | |
| M2 | 0.57 | 0.86 | 0.12 | 0.98 | 0.91 | 0.94 | 0.33 | 0.96 | 0.99 | |
| | 0.26 | 0.32 | | 0.34 | 0.55 | 0.56 | | 0.57 | 0.57 | |
| M3 | 0.91 | 0.94 | 0.16 | 1.00 | 0.95 | 0.92 | 0.38 | 1.00 | 0.99 | |
| | 0.39 | 0.39 | | 0.41 | 0.60 | 0.59 | | 0.62 | 0.61 | |
| M4 | 0.74 | 0.80 | 0.42 | 1.00 | 0.82 | 0.69 | 0.42 | 0.99 | 0.98 | |
| | 0.56 | 0.58 | | 0.65 | 0.59 | 0.54 | | 0.65 | 0.65 | |

615 3.2.8 Case 4

616 In this case, Fig. 7 shows that only wavelet-like HP filtering is 617 able to resolve images well. As expected, Fig. 12 shows a 618 better performance of schemes M4 and M3 than that of 619 schemes M1, M2, and M0 on both the 1-D and the 2-D mea-620 sures.

621 In summary, it can be seen that the evaluations depicted in
622 Figs. 9–12 using our defined measures are quite consistent
623 with those evaluations based on visual perception on Figs.
624 4–7.

Case 1 is the only example that is resolved to some extent ⁶²⁵ without having to use HPF (M0). However, scheme M0 made ⁶²⁶ some measure evaluation better than others since the corre- ⁶²⁷ sponding M0 reconstructions have a nearly uniform distribu- ⁶²⁸ tion. In spite of this, the measures we defined remain effective ⁶²⁹ for most of the 1-D and 2-D cases. ⁶³⁰

3.3 Evaluation on Defined Measures

In an aspect of quantitative discussions on resolution, we em- 632 ploy these four measures to explain the effectiveness of each 633

631

Table 4 Case 4 separation, size, contrast, and location resolutions for various filtering on 1-D and 2-D conditions.

| | 1-D | | | | 2-D | | | | | |
|----|----------------|----------------|----------|----------------|----------------|----------------|----------|------------------|------------------|--|
| | Sep. 0 Sep. | Size O Size | Contrast | Loc. 0 Loc. | Sep. 0 Sep. | Size 0 Size | Contrast | Locx. 0 Locx. | Locy. 0 Locy. | |
| MO | 0.85 | 0.81 | 0.09 | 1.00 | 0.97 | 0.82 | 0.19 | 0.99 | 1.00 | |
| | 0.28 | 0.27 | | 0.30 | 0.43 | 0.40 | | 0.44 | 0.44 | |
| M1 | 0.79 | 0.83 | 0.09 | 0.98 | 0.97 | 0.84 | 0.40 | 0.97 | 0.96 | |
| | 0.27 | 0.27 | | 0.30 | 0.62 | 0.58 | | 0.62 | 0.62 | |
| M2 | 0.63 | 0.81 | 0.08 | 0.98 | 0.94 | 0.85 | 0.31 | 0.96 | 0.99 | |
| | 0.22 | 0.25 | | 0.28 | 0.54 | 0.51 | | 0.54 | 0.55 | |
| M3 | 0.83 | 0.89 | 0.11 | 1.00 | 0.97 | 0.87 | 0.41 | 1.00 | 1.00 | |
| | 0.31 | 0.32 | | 0.34 | 0.63 | 0.60 | | 0.64 | 0.64 | |
| M4 | 0.83 | 0.91 | 0.12 | 0.99 | 0.97 | 0.89 | 0.44 | 1.00 | 0.96 | |
| | 0.32 | 0.34 | | 0.35 | 0.65 | 0.63 | | 0.66 | 0.65 | |

Pan et al.: Highly resolved diffuse optical tomography: a systematic approach...



Fig. 9 Case 1: (a) 1-D measures and (b) 2-D measures, where solid, +, \star , Δ , and \bigcirc lines represent using schemes of M0 to M4, respectively.

634 proposed filtering. Particularly, accurate demonstrations for 635 the 1-D condition are almost fully matched with the evalua-636 tion in quality. To an extent, most are also promising for the 637 2-D condition. In other words, the evaluation implies that our 638 defined measures are quite acceptable. For further inspection, 639 these measures can be seen based on individual inclusion or 640 separation as well. In this subsection, more discussion of the 641 defined measures is given. First, the contrast resolution can be 642 also defined as

$$R_{\text{cont.}}^{1\text{D,2D}} = \frac{(\text{mean}^{\text{incl.}}/\text{mean}^{\text{incl.}})_{\text{Reconstruction}}}{(\text{mean}^{\text{incl.}}/\text{mean}^{\text{incl.}})_{\text{Original}}},$$
(29)

644 where mean is to find the average value of the selected re-645 gions as the superscripts (incl. and incl). A definition in this 646 manner, however, is not suitable for our cases 1 to 3. For 647 further investigation, Eq. (23) can be concluded as



Fig. 10 Case 2 with (a), (b) and line key as described for Fig. 9.



Fig. 11 Case 3 with (a), (b) and line key as described for Fig. 9.

$$R_{\text{cont.}}^{1\text{D},2\text{D}} \begin{cases} >c & \text{normal situation} \\ =c & \text{no contrast} \\
(30)
(30)$$

where c is equal to $1/(\mu^{\text{incl.}}/\mu^{\text{incl.}})$ (0.1 is used here) and the 649 abnormal situation, here, means the optical value of the inclu- 650 sion is smaller than that of the separation region. Likewise, 651 the separation and size resolution can be defined as 652

$$Ro_{\text{sep;size}}^{1\text{D,2D}} = \frac{(\text{MSE}^{\text{incl.;incl.}})_{\text{Recon.2B(b)aseline}}}{(\text{MSE}^{\overline{\text{incl.;incl.}}})_{\text{Ori.2B(b)aseline}}}.$$
 (31)

It is found that Eq. (31) eventually regards a reconstructed **654** "inclusion" as a reverse cave with values ranging between 0 **655** and 1, and Eq. (31) always gives positive values. When con- **656** sidering Eqs. (25) and (26), it can be proven that both **657** $Ro_{\text{sep.:size}}^{1D,2D}$ are always smaller than a unit and, moreover, are **658**



Fig. 12 Case 4 with (a), (b) and line key as described for Fig. 9.

659 negative values to denote a high underestimation or overesti-660 mation. Note that our 2-D ROI can be determined automati-661 cally using a computer program but not with manual selec-662 tion, whereas the FWHM is not adopted in this paper because 663 it is selected manually. Finally, the location resolution is regu-664 lated by Eq. (28). Prior to adjustment, the positive or negative 665 errors to the unit can be explained as denoting the multiinclu-666 sion position in a reverse direction.

667 4 Concluding Remarks

668 We proposed and implemented a resolution-enhancing tech-669 nique using HPF incorporated with the FEM-based inverse 670 computation to obtain highly resolved NIR diffuse optical im-671 ages in a systematical manner. As mentioned previously, our 672 approach, derived from the Poisson MAP, was justified by 673 various HPFs for different designated phantoms. Qualitative 674 visual perception and quantitative evaluations of the recon-675 structions also validate the proposed approaches.

676 Obviously, the wavelet-like HP filtering is superior to the 677 LPF-combined HPF, as shown in Figs. 4–7. In summary, the 678 approach to use the wavelet-like HP filtering, M3, is recom-679 mended in terms of its resolving ability and computational 680 stability. It is observed that the M4 scheme demonstrates a 681 high resolution result as well, but reveals worse stability than 682 the M3 scheme. Additionally, a small inclusion-to-background 683 diameter ratio, 2:20, is detectable and distinguished.

Due to the variation in the choice of σ_1 and σ_2 associated 684 685 with each filter, various filters result in different reconstruc-686 tion results. In this paper, we did not attempt to conduct a 687 wide comparison and an extensive study over a range of HP 688 filters and phantom cases, but rather chose to begin with two 689 categories of filters and a set of more-or-less extreme cases. 690 Although the resolutions of absorption images enhanced with 691 our proposed techniques were presented, this approach re-692 mains effective to improve the scattering images for the 693 frequency-domain DOT imaging system as well. In future 694 work, a thorough investigation of HP filters used in the pro-695 posed approach will be conducted to find one or several ap-696 propriate filters. Moreover, the resolution limit should be 697 specified with a set of designed cases. A further study is also 698 required to identify exact causes of the negative-value arti-**699** facts shown in Figs. 5(f)-5(j) and 6(f)-6(j) which are depen-700 dent either on the filters or on the cases themselves. Owing to 701 the lack of a sound method for quantitative evaluation, it is 702 believed that even to objectively define a measure correspond-703 ing to visual perception is quite complicated. Alternatively, 704 four reasonable measure definitions were considered and de-705 fined to provide an initial basis for quantitative evaluations, **706** from which further explorations of an individual inclusion or 707 separation ROI can begin. Briefly, our proposed measures 708 mainly provide approximate information, and areas remain for 709 further investigation and improvement.

710 Acknowledgments

711 The authors would like to acknowledge the funding support
712 from the grants by the Veteran General Hospital/University
713 System of Taiwan Joint Research Program (VGHUST95-P4714 13, VGHUST96-P4-17) and the National Science Council
715 (NSC 95-2221-E-236-002) in Taiwan.

References

 G. A. Millikan, "The oximeter, an instrument for measuring continuously the oxygen saturation of arterial blood in man," *Rev. Sci. In-* 718 *strum.* 13, 434–444 (1942). 719

- J. S. Maier, S. A. Walker, S. Fantini, M. A. Franceschini, and E. 720 Gratton, "Possible correlation between blood glucose concentration 721 and the reduced scattering coefficient of tissues in the near infrared," 722 *Opt. Lett.* 19, 2062–2064 (1994). 723
- M. Kohl, M. Cope, M. Essenpreis, and D. Böcker, "Influence of 724 glucose concentration on light scattering in tissue-simulating phantoms," *Opt. Lett.* 19, 2170–2172 (1994). 726
- B. W. Pogue, S. D. Jiang, H. Dehghani, C. Kogel, S. Soho, S. Srini-727 vasan, X. M. Song, T. D. Tosteson, S. P. Poplack, and K. D. Paulsen, 728 "Characterization of hemoglobin, water, and NIR scattering in breast 729 tissue: analysis of intersubject variability and menstrual cycle 730 changes," *J. Biomed. Opt.* 9(3), 541–552 (2004). 731
- B. Brooksby, S. D. Jiang, H. Dehghani, B. W. Pogue, K. D. Paulsen, 732 J. Weaver, C. Kogel, and S. P. Poplack, "Combining near-infrared 733 tomography and magnetic resonance imaging to study *in vivo* breast 734 tissue: implementation of a Laplacian-type regularization to incorporate magnetic resonance structure," *J. Biomed. Opt.* 10, 051504 736 (2005). 737
- P. J. Cassidy and G. K. Radda, "Molecular imaging perspectives," J. 738 R. Soc., Interface 2, 133–144 (2005). 739
- J. C. Hebden and D. T. Deply, "Enhanced time-resolved imaging with 740 a diffusion model of photon transport," *Opt. Lett.* 19(5), 311–313 741 (1994). 742
- J. A. Moon and J. Reintjes, "Image resolution by use of multiply 743 scattered light," *Opt. Lett.* 19(8), 521–523 (1994). 744
- H. Jiang and K. D. Paulsen, "A finite element based higher-order 745 diffusion approximation of light propagation in tissues," *Proc. SPIE* 746 2389, 608–614 (1995). 747
- H. Jiang, K. D. Paulsen, and U. L. Österberg, "Optical image reconstruction using DC data: simulations and experiments," *Phys. Med.* 749 *Biol.* 41, 1483–1498 (1996). 750
- H. Jiang, K. D. Paulsen, U. L. Österberg, and M. S. Patterson, 751 "Frequency-domain optical image reconstruction in turbid media: an 752 experimental study of single-target detectability," *Appl. Opt.* 36, 753 52–63 (1997). 754
- H. Jiang, K. D. Paulsen, U. L. Österberg, and M. S. Patterson, "Improved continuous light diffusion imaging in single- and multi-target tissue-like phantoms," *Phys. Med. Biol.* 43, 675–693 (1998).
- B. W. Pogue and K. D. Paulsen, "High-resolution near-infrared tomographic imaging simulations of the rat cranium by use of a priori magnetic resonance imaging structural information," *Opt. Lett.* 23, 760 1716–1718 (1998).
- B. W. Pogue, T. McBride, C. Nwaigwe, U. L. Österberg, J. F. Dunn, 762 and K. D. Paulsen, "Near-infrared diffuse tomography with a priori 763 MRI structural information: testing a hybrid image reconstruction 764 methodology with functional imaging of the rat cranium," *Proc. SPIE* 765 3597, 484–492 (1999). 766
- V. Ntziachristos, "Concurrent diffuse optical tomography, spectros- 767 copy and magnetic resonance imaging of breast cancer," PhD Diss., 768 University of Pennsylvania (2000). 769
- H. Dehghani, B. W. Pogue, and K. D. Paulsen, "Development of 770 hybrid NIR/MRI imaging system algorithm: use of a-priori information for tumor detection in the female breast," in *Proc. IEEE Int.* 772 *Symp. on Biomedical Imaging*, pp. 657–660 (2002). 773
- H. Xu, H. Dehghani, B. W. Pogue, K. D. Paulsen, and J. F. Dunn, 774 "Hybrid MR/near infrared imaging of the murine brain: optimization 775 of optical fiber arrangement and use of *a-priori* knowledge," in *Proc.* 776 *IEEE Int. Symp. on Biomedical Imaging*, pp. 74–77 (2002). 777
- B. A. Brooksby, H. Dehghani, B. W. Pogue, and K. D. Paulsen, 778 "Near-infrared (NIR) tomography breast image reconstruction with a 779 priori structural information from MRI: algorithm development for 780 reconstructing heterogeneities," *IEEE J. Sel. Top. Quantum Electron.* 781 9, 199–209 (2003). 782
- Q. Zhu, T. Durduran, V. Ntziachristos, M. Holboke, and A. G. Yodh, 783
 "Imager that combines near-infrared diffusive light and ultrasound," 784
 Opt. Lett. 24, 1050–1052 (1999). 785
- P. Guo, D. Piao, Q. Zhu, and J. Fikiet, "A combined 2-D ultrasound 786 and NIR imaging system," in *Proc. IEEE 26th Annu. Northeast* 787 *Bioengineering Conf.*, pp. 77–78 (2000). 788
- M. J. Holboke, B. J. Tromberg, X. Li, N. Shah, J. Fishkin, D. Kidney, 789
 J. Butler, B. Chance, and A. G. Yodh, "Three-dimensional diffuse 790

- optical mammography with ultrasound localization in a human subject," *J. Biomed. Opt.* 5, 237–247 (2000).
- 793 22. B. Brooksby, S. Jiang, H. Dehghani, B. W. Pogue, K. D. Paulsen, J.
 794 Weaver, C. Kogel, and S. P. Poplack, "Combining near-infrared to-
- mography and magnetic resonance imaging to study *in vivo* breast
- tissue: implementation of a Laplacian-type regularization to incorporate magnetic resonance structure," *J. Biomed. Opt.* 10, 051504 (2005).
- 799 23. B. Brooksby, B. W. Pogue, S. Jiang, H. Dehghani, S. Srinivasan, C.
 Kogel, T. D. Tosteson, J. Weaver, S. P. Poplack, and K. D. Paulsen,
 "Imaging breast adipose and fibroglandular tissue molecular signatures by using hybrid MRI-guided near-infrared spectral tomogra-

803 phy," *Proc. Natl. Acad. Sci. U.S.A.* 103(23), 8828–8833 (2006).
 804 24. B. Brooksby, S. Srinivasan, S. Jiang, H. Dehghani, B. W. Pogue, a

- 804 24. B. Brooksby, S. Srinivasan, S. Jiang, H. Dehghani, B. W. Pogue, and
 805 K. D. Paulsen, "Spectral priors improve near-infrared diffuse tomography more than spatial priors," *Opt. Lett.* 30(15), 1968–1970 (2005).
- raphy more than spatial priors," *Opt. Lett.* 30(15), 1968–1970 (2005).
 807 25. B. Kanmani and R. M. Vasu, "Diffuse optical tomography through solving a system of quadratic equations without re-estimating the derivatives: the Frozen-Newton method," in *Proc. IEEE Int. Work-shop on Biomedical Circuits and Systems*, pp. S2.2-17–20 (2004).
- **811** 26. H. Jiang, "Optical image reconstruction based on the third-order dif-
- 812 fusion equations," *Opt. Express* 4, 241–246 (1999).
 813 27. B. W. Pogue, T. O. McBride, J. Prewitt, U. L. Österberg, and K. D. Paulsen, "Spatially variant regularization improves diffuse optical to815 mercenetary" *Acut.* 20(2) 2050-2061 (1000)
- 815 mography," *Appl. Opt.* 38, 2950–2961 (1999).
 816 28. D. H. Brooks, R. J. Gaudette, E. L. Miller, C. A. DiMarzio, D. Boas,
- and M. Kilmer, "An admissible solution approach for diffuse optical
- and M. Klimer, An admissible solution approach for diffuse optical tomography," in *Proc. IEEE 34th Asilomar Conf. on Signals, Systems and Computers*, pp. 333–337 (2000).
- 820 29. Y. H. Zhang, D. H. Brooks, and D. Boas, "A multi-resolution admissible solution approach for diffuse optical tomography," in *Proc.*
- 822 IEEE Int. Symp. on Biomedical Imaging, pp. 1005–1008 (2002).
 823 30. M. Guven, B. Yazici, X. Intes, and B. Chance, "An adaptive multi-
- grid algorithm for region of interest diffuse optical tomography," in
 Proc. IEEE Int. Conf. Image Processing, Vol. II, pp. 823–826 (2003).
- 826 31. M. Guven, B. Yzazici, X. Intes, and B. Chance, "An adaptive v-grid algorithm for diffuse optical tomography," in *Proc. IEEE 29th Annu.*
- 828 Northeast Bioengineering Conf., pp. 95–96 (2003).
- 829 32. J. J. Stott, J. P. Culver, S. R. Arridge, and D. A. Boas, "Optode positional calibration in diffuse optical tomography," *Appl. Opt.* 42,

3154-3162 (2003).

- V. Ntziachristos, A. G. Yodh, M. Schnall, and B. Chance, "Concurrent MRI and diffuse optical tomography of breast after indocyanine green enhancement," *Proc. Natl. Acad. Sci. U.S.A.* 97, 2767–2772 834 (2000).
- X. Intes, J. Ripoll, Y. Chen, S. Nioka, A. G. Yodh, and B. Chance, "In 836 vivo continuous-wave optical breast imaging enhanced with indocyanine green," *Med. Phys.* 30, 1039–1047 (2003).
- 35. W. K. Pratt, Digital Image Processing, Wiley, New York (1991). 839
- R. J. Schalkoff, Digital Image Processing and Computer Vision, 840 Wiley, New York (1989).
- R. C. Gonzalez, R. E. Woods, and S. L. Eddins, *Digital Image Pro-* 842 cessing, Prentice Hall, Upper Saddle River, NJ (2004).
 843
- B. R. Hunt and P. J. Sementilli, "Description of a Poisson imagery 844 super-resolution algorithm," *Astron. Data Anal. Software System I.*, 845 *A.S.P. Conf. Ser.* 25, 196–199 (1992).
- M.-C. Pan, "Improving a single down-sampled image using 847 probability-filtering-based interpolation and improved Poisson maximum a posteriori super-resolution," *EURASIP J. Appl. Signal Pro-* 849 *cess.* 2006, 97492 (2006).
- M.-C. Pan, "A novel blind super-resolution algorithm for restoring 851 Gaussian blurred images," *Int. J. Imaging Syst. Technol.* 12(6), 239– 852
 853
- K. D. Paulsen and H. Jiang, "Spatially varying optical property reconstruction using a finite element diffusion equation approximation," *Med. Phys.* 22, 691–701 (1995).
- X. Song, B. W. Pogue, S. Jiang, M. M. Doyley, H. Dehghani, T. D. 857 Tosteson, and K. D. Paulsen, "Automated region detection based on the contrast-to-noise ratio in near-infrared tomography," *Appl. Opt.* 859 43(5), 1053–1062 (2004).
- S. C. Davis, B. W. Pogue, H. Dehghani, and K. D. Paulsen, 861 "Contrast-detail analysis characterizes diffuse optical fluorescence tomography image reconstruction," *J. Biomed. Opt.* 10(5), 050501-1–3 863 (2005).
- B. W. Pogue, S. C. Davis, X. Song, B. A. Brooksby, H. Dehghani, 865 and K. D. Paulsen, "Image analysis methods for diffuse optical tomography," *J. Biomed. Opt.* 11(3), 033001-1–16 (2006).
- C. A. Kelsey, R. D. Moseley, Jr., J. F. Garcia, F. A. Mettler, Jr., T. W. 868 Parker, and J. H. Juhl, "ROC and contrast detail image evaluation tests compared," *Radiology* 154(3), 629–631 (1985).