

Sample of Level 3 Editing (Biomedical & Imaging)

Assessment of Reduced Encoding Diffusion Spectrum Imaging

Implemented with a Bi-Gaussian Model Using Phantoms and

Manganese-Enhanced Optic Tracts

Abstract

Diffusion spectrum imaging (DSI) can map complex fiber microstructures in tissues by characterizing their 3-D water diffusion spectra. However, a long acquisition time is required for adequate q-space sampling to completely reconstruct the 3-D diffusion probability density function (PDF). Furthermore, to achieve a high q- or b-value encoding for sufficient spatial resolution, the diffusion gradient duration and diffusion times are usually ~~enlarged-lengthened~~ on a clinical scanner, ~~which~~resulting in a long echo time and low signal-to-noise ratio (SNR) of diffusion images. To bypass long acquisition times and ~~strong-strict~~ gradient requirements, the reduced-encoding DSI with a bi-Gaussian diffusion model (RE-DSI) is presented in this study. The bi-Gaussian extrapolation kernel, ~~which is~~ based on the assumption of a bi-Gaussian diffusion signal curves across biological tissues, is performed to fulfill a high q-value ~~request~~

requirement on the reduced-encoding scheme. ~~Both~~The intersecting capillary~~ies~~ phantom model~~s~~ and the manganese-enhanced rat model~~s~~ ~~were~~ served as standards for accuracy assessment in RE-DSI, ~~the~~ The errors of RE-DSI in defining fiber orientation were quantified and the results were found to be close to the noise limit. Evidences from a human study demonstrated that RE-DSI significantly decreased the acquisition times, required to ~~meanwhile~~ resolve d complex neural fibers. The presented acquisition method facilitates the application~~s~~ of DSI analysis on a clinical magnetic resonance imaging (MRI) system.

Keywords: diffusion spectrum imaging; phantom model; manganese-enhanced rat model

Comment [WL1]: CHECK:
“...served as standards” for what? I think you might mean they “served as standards for accuracy assessment in RE-DSI”. This is used later in the article.

Introduction

Diffusion MRI has become an essential tool for contrast imaging ~~mechanism of the~~ for central nervouse system. This, has led to and ~~made a~~ significant improvement in clinical diagnosis. Further ~~progress advancement~~ to the technique has been made with the ~~design~~ introduction of diffusion tensor imaging (DTI) [1, 2]. The ~~technique makes further progress along with the design of diffusion~~ tensor imaging (DTI) [1, 2], which is a feasible valuable technique ~~for in~~ identifying to model anisotropic diffusion as well as non-invasively to delineate ing the principle orientations of white matter tracts non-invasively [3-5]. However, the assumption ~~of~~ a single Gaussian diffusion compartment in the tensor model results in the ambiguous orientations of fibers in regions where they cross each other containing crossing fibers [6]. Thus, it with the typical resolution ~~of a MRI, it may be~~ becomes intricate difficult to interpret the complex neural connections between functional areas of athe human brain with under typical resolution of an MRI. ___

In recent years, various diffusion imaging strategies have been developed to improve the depiction of water diffusion and to resolve the intravoxel fiber orientations. Diffusion spectrum imaging (DSI) [7]; for example, utilizes the 3-D spectra of water displacements to characterize the heterogeneity ities of fiber architectures. DSI was is based on the ~~Its~~ theory established ~~on~~ by the q-space imaging technique ~~;~~ technique, which The theory describes the Fourier relationship between echo signal attenuation and the probability density function (PDF) of the displacement of water molecules s ~~displacements~~ with the prerequisite of a narrow pulse approximation

[8-10]. ~~The DSI technique was~~ has been used ~~into map the mapping~~ tissue architecture of biological systems ~~DSI has shown its capability of mapping tissues architectures in biological systems~~ [7,–11], providing information on the intravoxel compartment sizes ~~scales~~ of ~~the~~ neural fibers [12], ~~thus allowing and interpreting the~~ physiological and structural conditions of ~~the~~ neural tissues to be interpreted. In addition, 3-D tractography and comparative segmentation of human brain structures have been identified based on DSI and the proceeding orientation distribution function (ODF) [13].

~~Notwithstanding~~ ~~t~~The utility of DSI comes at a cost: a complete reconstruction of the diffusion PDF requires 515 q-value encoding points distributed on a Cartesian lattice across 3-D q-space. This involves long acquisition times as well as adequate ~~q~~-values for sufficient resolution. Since the available gradient strength in clinical systems are limited, ~~t~~The latter requirement is achieved by prolonging the diffusion gradient duration (δ) and the diffusion time (Δ) ~~since the available gradient strength in clinical systems is limited~~. Unfortunately, this ~~would accompany~~ leads to a long echo time (TE) and a decline in the SNR ~~level~~ due to a severe T2 decay in ~~an~~the echo planar imaging (EPI) sequence. As a consequence of this, ~~The~~ angular accuracy and discrimination ~~would be~~ unavoidably diminished ~~as a consequence~~ [11]. ~~Both~~ ~~t~~The lengthy acquisition times, ~~cost~~ and the requirements of the gradient system, ~~request~~ have ~~retarded~~hindered the further applications of DSI on clinical scanners.

–These limitations basically stems from the need to exhaustively sample on a 3-D Cartesian sampling lattice.

A hemispheric encoding scheme (half-q-DSI) ~~could~~ can be applied to halve the scan time in DSI since the ~~the~~ diffusion contrast is

Comment [TK2]: CHECK:

This sentence is confusing. It seems you are saying that tractography and comparative segmentation of brain structures were identified with DSI. Then you talk about what appears to be another technique, ODF. However, it is not clear how this technique is related to DSI or was it just another technique used in conjunction with DSI? Consider rewording to clarify. Also, if ODF is another technique (in addition to DSI), which has been helpful in reconstructing tissue structure, it may be good to point that out. At the moment, it doesn't quite tie into the rest of the paragraph but just appears all of a sudden at the end.

Comment [TK3]: CHECK:

Consider not using the word “adequate”. Perhaps use ‘large quantities of’ if that is appropriate.

positive and spherically symmetric [7, 14]. However, ~~uncorrected~~ cross-term interactions ~~ss~~ between diffusion and imaging gradients might result in ~~the a misunderstanding misinterpretation~~ of the q-space analysis and inaccurate ODFs in half-q-DSI [15, 16]. Instead of a Cartesian lattice, a body-centered ~~ed~~ cubic lattice (~~BB~~~~CB~~~~CC~~) sampling scheme ~~is~~~~was~~ proposed to ~~gain~~ ~~improve the~~ imaging efficiency of DSI by 30% [17]. ~~Another non-Cartesian q-space encoding scheme, Hybrid diffusion imaging (HYDI), was~~ ~~has also been employed for DSI-PDF reconstruction. This scheme consists~~ ~~ss of five concentric spherical shells and may be applied to multiple types of diffusion analyses~~ [18]. ~~, which is flexible for multiple diffusion analyses, employs a non-Cartesian q-space encoding scheme comprising five concentric spherical shells for DSI-PDF reconstruction. Although it was possible to shorten the~~ ~~the~~ acquisition times ~~could be shortened~~ with ~~all of the above~~ q-space sampling strategies ~~described above, the need~~ ~~requirement for~~ ~~of~~ a large number of ~~high~~ q-values to preserve ~~adequate spatial resolution~~ ~~acquisitions~~ could not be omitted ~~to preserve adequate spatial resolution.~~

Another category of diffusion imaging ~~techniques~~ ~~methods~~ utilizes an encoding scheme formed by a single spherical shell with a constant diffusion weighting; as opposed to ~~athe~~ 3-D Cartesian lattice with multiple diffusion weightings. These ~~se~~ techniques include high angular resolution diffusion imaging (HARDI) [19, 20], q-ball imaging (QBI) [21, 22], persistent angular structure MRI (PAS-MRI) [23], ~~fiber orientation estimation~~ ~~eed~~ using continuous axially symmetric ~~al~~ tensors (FORECAST) [24], diffusion orientation transformation (DOT) [25], and spherical deconvolution methods [26, 27]. These approaches

Comment [TK4]: CHECK:
Are you trying to say that the cross-term interactions would not be corrected (uncorrected) or that they would not be correct (incorrect).

Comment [WL5]: CHECK:
You don't use this term anywhere else, so there is no need for an abbreviation.

Comment [TK6]: CHECK:
You don't use this term anywhere else, so there is no need for an abbreviation.

Comment [TK7]: CHECK:
Would it be appropriate to say: 'estimation of fiber orientation using continuous axially symmetrical tensors'?

Comment [WL8]: CHECK:
The abbreviations used in this sentence (eg. HARDI, QBI, PAS-MRI) are not needed as the terms are only used here and nowhere else in the paper.

provide ~~information on the orientation~~directional information of complex neural fiber ~~networks~~ within ~~a a-feasible-reasonable~~ scan time ~~and may be for~~ routinely ~~implemented.~~ ~~implementation.~~ The substantial~~ly~~ increase in imaging efficiency mainly results from ~~the~~ fewer numbers of diffusion-weighted images (DWIs) ~~needed-required~~ for data analysis. In addition, the shortened TEs ~~following on a moderate b-value~~ could enhance the SNR of DWIs. ~~These conditions; however, may be insufficient to characterize the 3-D diffusion function that is derived from the multiple q-value diffusion measurements, and would thus would-be unable to For-inferring tissue-microstructural tissue conditions~~shape and orientations. ~~-, however, they might be insufficient to characterize the 3-D diffusion function derived from the multiple q-values diffusion measurements.~~

Comment [WL9]: CHECK:

Do you mean “are routinely implemented” or only “may be routinely implemented”? “May be” indicates that they are not currently routinely implemented but “may be”.

Comment [TK10]: CHECK:

‘following a moderate b value’ OR ‘followed by a moderate b value’ would make more sense. If neither of those fits, consider revising that part of the sentence. Perhaps ‘following’ is not an appropriate word choice here.

Comment [WL11]: CHECK:

Do you mean “can enhance” or “could enhance”? If they always enhance it, use “can enhance”.

In this study, ~~it is proposed that the~~ reduced-encoding DSI ~~implemented-complemented~~ with a bi-Gaussian model (RE-DSI), ~~is proposed~~be used to trim down the drawbacks of DSI ~~as well as~~while ~~to-retaining~~q-space information. In RE-DSI, a reduced Cartesian sampling scheme, where ~~high q-value~~ acquisitions are omitted, is used to bypass long acquisition times and gradient system demands in DSI. To achieve sufficient resolution ~~for-to resolving-determining~~determine ~~the fiber-orientations~~of fibers, the 1-D bi-Gaussian model ~~fitting is performed-onis applied to~~ the sampled data at low ~~q-space~~ to ~~regain~~ all diffusion signals at high ~~q-space~~. Previous studies on animal and human brains have demonstrated that diffusion-attenuated curves ~~could~~ can be characterized as a bi-exponential function [28-31]. Accordingly, we hypothesized that ~~the~~ diffusion signal attenuation along each radial direction in ~~q-space~~ was a bi-Gaussian function. ~~The-This~~ assumption

Comment [TK12]: CHECK:

Throughout the paper, you refer to high q-values. However, it is not clear what you mean by that. Is it many q-values? Or, a high value of q? It would be helpful if this term was explained more clearly early on in the paper to avoid confusion.

Comment [TK13]: CHECK:

Do you mean ‘regain’ or ‘retain’?

is similar ~~to with~~ that used in the DOT technique [25], which ~~straightforwardly~~ directly converts the diffusivity function into displacement probabilities at a particular distance away from the origin, while RE-DSI tends to reconstruct a diffusion PDF from q-space signals.