DiffusionKit Documentation

Release 1.2.1

Brainnetome Center, CASIA

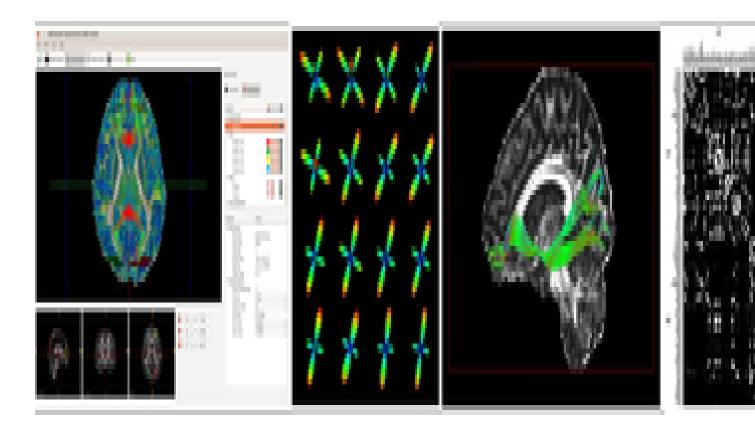
Apr 27, 2020

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CHAPTER 1

Welcome to Brainnetome DiffusionKit's Homepage



Note:

- a full pipeline for (pre-)processing and visualization for diffusion MRI data.
- cross-platform support and a small installation size without 3rd party dependency.
- utilize a graphical interface along with command-line programs that enables easy operation and batch process-

ing.

Brainnetome DiffusionKit is a light one-stop cross-platform solution to dMRI data analysis. The package delivers a complete pipeline from data format conversion to preprocessing, from local reconstruction to fiber tracking, and from fiber statistics to visualization.

It was developed as a cross-platform framework, using VTK [2] for visualization, and Qt for GUI design. Both GPU and CPU computing were implemented for visualization to achieve high frame-rate, for rendering complex scene like whole brain tractographs in particular. The project was managed using the compiler-independent CMake [3], which is compatible with gcc/g++ and MS Visual Studio, etc. Well-established algorithms, such as the DICOM conversion tool dcm2nii by Chris Rorden [4] and the constrained spherical deconvolution (CSD) for HARDI reconstruction in MRtrix [5], were adopted with improved interface and user experience.

- Visit Manual page for a complete list of usage instructions.
- Visit Tutorial page to take a look at how DiffusionKit can solve your practical problems.
- Visit Screenshot page to see how the GUI front-end looks like.
- Visit Support page to submit comments, or email us at diffusion.kit@nlpr.ia.ac.cn .

Please see the navigation sidebar to the left to begin.

Important: DiffusionKit v1.5 Released! Please Update your DiffusionKit.

- Fix bugs.
- Add features in 2D image display.
- Add functions to transform fiber track format between DiffusionKit and MRTrix.

The citation for DiffusionKit could be:

Sangma Xie, Liangfu Chen, Nianming Zuo and Tianzi Jiang, **DiffusionKit: A Light One-Stop Solution** for Diffusion MRI Data Analysis, *Journal of Neuroscience Methods*, vol. 273, pp. 107-119, 2016. [PDF].

1.1 Manual

1.1.1 Overview

Brainnetome DiffusionKit is a light one-stop cross-platform solution to dMRI data analysis. The package delivers a complete pipeline from data format conversion to preprocessing, from local reconstruction to fiber tracking, and from fiber statistics to visualization. DiffusionKit was developed as a cross-platform framework, using ITK [1] for computation, VTK [2] for visualization, and Qt for GUI design. Both GPU and CPU computing were implemented for visualization to achieve high frame-rate, for rendering complex scene like whole brain tractographs in particular. The project was managed using the compiler-independent CMake [3], which is compatible with gcc/g++ and MS Visual Studio, etc. Well-established algorithms, such as the DICOM conversion tool dcm2nii by Chris Rorden [4] and the constrained spherical deconvolution (CSD) for HARDI reconstruction in MRtrix [5], were adopted with improved interface and user experience.

Func- tions	Program	Description (use '-h' argument for more details)	
Preprocess	dcm2nii	Convert DICOM to unified 4D NIFTI files	
1	bneddy	Correct eddy current induced distortion and head motion	
	topup/eddy	Correct eddy current and susceptibility induced distortion and head movements	
	bet2	Extract brain tissue (Smith, 2002)	
	bnsplit,bnmerge	Split/merge the 4D image along the 4th dimension	
Modeling	bndti_estimate	Estimate tensor model, output FA, MD, tensor etc.	
	bnhardi_ODF_estimate	Estimate ODF by SPFI method	
	bnhardi_FOD_estimate	Estimate FOD by CSD method (Tournier et al., 2007)	
Tracking	bndti_tracking	Track white matter fiber based on tensor model	
	bnhardi_tracking	Track white matter fiber based on ODF/FOD	
Visualize	bnviewer	Visualize various kinds of data (.nii.gz, .trk)	
Tools	bncalc,bnroisplit	Numeric calculation and ROI generation	
	bninfo	Show the head information of DICOM and NIFTI files	
	reg_aladin,reg_f3d	Inter/intra-image registration across modalities	
	reg_resample,	Resample and apply transformation matrix	
	reg_transform		
	bnfiber_end, bn-	Prune fiber bundle, logical and/or/not based on ROIs	
	fiber_prune		
	bnfiber_stats	Export attributes of fiber bundles	
	bnfiber_map	Generate the fiber density going through each voxel	
	bnnetwork	Function to construct anatomical network	

Key functions of the software

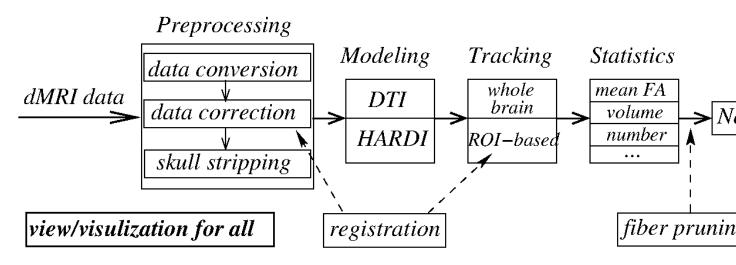


Fig. 1: Figure 1. The overall design framework of DiffusionKit

It should be noteworthy that, for all the computing steps provided in GUI, the called commands with the complete parameter list are displayed in a separate log window. Such a design is special for the users to keep in mind what he/she is doing, and furthermore, it could be directly copied into script (Bash, Python ...) for batch processing.

1.1.2 Data Processing Pipeline

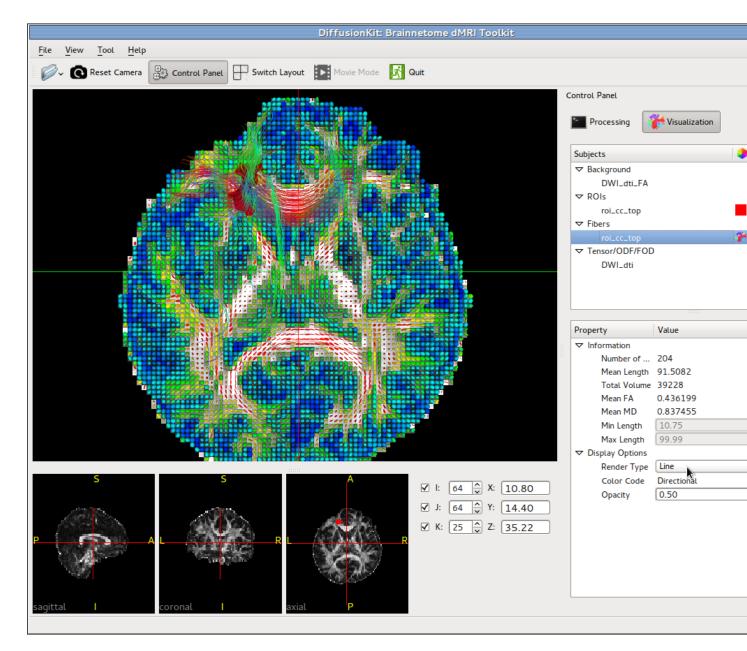


Fig. 2: Figure 2. The main window of the software.

Preprocessing

A set of command line tools for data preprocessing steps is described in this section, which include data format conversion, data correction and brain extraction.

DICOM to NIFTI Conversion

For saving space of data store, the nii.gz format was used throughout the whole pipeline. Firstly, we use the dcm2nii (by Chris Rorden, https://www.nitrc.org/projects/dcm2nii) to convert the data into a single 3/4D .nii.gz volume-series file, plus bval and bvec files. The format of bval file is

```
0 1500 1500 ... 1500
```

and the format of byec file is

```
00.999444842338560.002153856446970.002690417459230-0.00535333110020.998366951942440.6051847934722900.03288444131613-0.05708565562963-0.79608047008514
```

where the 0 in the first column indicates the b0 images in the scan and certainly the software also supports multib0 images. Since the DICOM formats from different scanner might be different, this function is not always able to successfully extract the bvec/bval files. If you encounter such a problem, please get back to us <link> and give the link of your data if it has big size beyond the email capability.

```
$ ./dcm2nii -h
Compression will be faster with /usr/local/bin/pigz
Chris Rorden's dcm2niiX version 24Nov2014
usage: dcm2nii [options] <in_folder>
Options :
      show help
-h
-f
     filename (%c=comments %f=folder name %p=protocol %i ID of
     patient %n=name of patient %s=series, %t=time; default 'DTI')
     output directory (omit to save to input folder)
 -0
     gz compress images (y/n, default n)
 -z
Defaults file : /home/ccm/.dcm2nii.ini
Examples :
dcm2nii /Users/chris/dir
dcm2nii -o /users/cr/outdir/ -z y ~/dicomdir
dcm2nii -f mystudy%s ~/dicomdir
dcm2nii -o "~/dir with spaces/dir" ~/dicomdir
Example output filename: '/DTI.nii.gz'
```

Data correction

During the MRI scanning, many factors can cause distortions and misalignment. The diffusion weighted imaging (DWI) volume series (as a 4D zipped NIFTI file) are then corrected for the distortions induced by off-resonance field and the misalignment caused by subject motion. The off-resonance effects are usually caused by the eddy currents of switching the diffusion encoding gradients and the susceptibility distribution of the imaged subjects, resulting in the deterioration of images due to blurring, spatial distortion, local signal artifacts, etc. The motion effects also cause image blurring and geometric misalignment [21] [22]. To correct the distortions induced by susceptibility and eddy current when the data is acquired with different phase-encode parameters, we include the correction mechanism using different phase-encode information. We exported the functions of topup, applytopup, eddy and eddy_combine

from FSL [21] [23] [24], compiled them on both Linux and Windows platforms, and packed the executable files into DiffusionKit. The detailed usage information, please refer to the website of FSL (http://fsl.fmrib.ox.ac.uk/fsl/fslwiki/).

```
$ ./topup -h
Part of FSL (build 504)
topup
Usage:
topup --imain=<some 4D image> --datain=<text file> --config=<text file with_
→parameters> --coutname=my_field
Compulsory arguments (You MUST set one or more of):
      --imain name of 4D file with images
      --datain
                    name of text file with PE directions/times
Optional arguments (You may optionally specify one or more of):
                    base-name of output files (spline coefficients (Hz) and
      --out
→movement parameters)
      --fout
                    name of image file with field (Hz)
                    name of 4D image file with unwarped images
      --iout
      --logout
                    Name of log-file
      --warpres (approximate) resolution (in mm) of warp basis for the
→different sub-sampling levels, default 10
      --subsamp sub-sampling scheme, default 1
      --fwhm
                    FWHM (in mm) of gaussian smoothing kernel, default 8
      --config
                  Name of config file specifying command line arguments
                    Max # of non-linear iterations, default 5
      --miter
      --lambda Weight of regularisation, default depending on --ssqlambda and
↔--regmod switches. See user documetation.
      --ssqlambda If set (=1), lambda is weighted by current ssq, default 1
      --reqmod
                     Model for regularisation of warp-field [membrane_energy_
→bending_energy], default bending_energy
      --estmov Estimate movements if set, default 1 (true)
                     Minimisation method 0=Levenberg-Marquardt, 1=Scaled Conjugate
      --minmet
⇔Gradient, default 0 (LM)
      --splineorder Order of spline, 2->Qadratic spline, 3->Cubic spline. Default=3
      --numprec
                    Precision for representing Hessian, double or float. Default.
⇔double
                     Image interpolation model, linear or spline. Default spline
       --interp
                     If set (=1), the images are individually scaled to a common_
      --scale
→mean, default 0 (false)
                    If set (=1), the calculations are done in a different grid,
      --regrid
→default 1 (true)
      -v,--verbose Print diagonostic information while running
      -h,--help
                     display help info
```

```
$ ./eddy -h
Part of FSL (build 504)
eddv
Copyright(c) 2011, University of Oxford (Jesper Andersson)
Usage:
eddy --monsoon
Compulsory arguments (You MUST set one or more of):
       --imain File containing all the images to estimate distortions for
       --mask Mask to indicate brain
       --index File containing indices for all volumes in --imain into --acqp and --
                                                                           (continues on next page)
```

⇔topup

```
--acqp File containing acquisition parameters
      --byecs File containing the b-yectors for all volumes in --imain
      --bvals File containing the b-values for all volumes in --imain
             Basename for output
      --out
Optional arguments (You may optionally specify one or more of):
       --session File containing session indices for all volumes in --imain
      --topup Base name for output files from topup
      --flm First level EC model (linear/quadratic/cubic)
      --fwhm FWHM for conditioning filter when estimating the parameters
      --niter Number of iterations (default 5)
      --resamp
                     Final resampling method (jac/lsr)
      --repol Detect and replace outlier slices
      -v,--verbose switch on diagnostic messages
      -h,--help
                      display this message
```

Unfortunately, most clinical acquisitions do not currently meet the requirement (two or more acquisitions where the parameters are different so that the mapping fields for distortion correction are different.) of topup. To handle this issue, we implemented a function called bneddy to correct eddy-current induced distortion and head movements efficiently. bneddy applies rigid and affine registrations to amend the distortions and misalignment.

```
$ ./bneddy -h
bneddy: Eddy Currents and Head Motion Correction.
(Mar 10 2016, 16:20:52)
-i Input file, 4D .nii.gz file.
-o Prefix for output file, and it will also output the log file as_
-Prefix.txt which will be called by bnrotate_bvec.
-ref 0 Reference image.
-omp 2 Max number of threads
```

Skull Stripping (Brain Extraction)

This module is to strip the brain skull and extract the brain tissue, including gray matter, white matter, cerebrospinal fluid (CSF) and cerebellum. It largely benefits the following processing and analysis, offering better registration/alignment results and reducing computational time by excluding non-brain tissue. Therefore, although this step is not compulsory, we strongly recommend enforcing it. This module is adapted from the FSL/BET functions (http://fsl.fmrib.ox.ac.uk/fsl/fslwiki/BET) for its excellent efficacy and efficiency.

```
$ ./bet2 -h
Part of FSL (build 504)
BET (Brain Extraction Tool) v2.1 - FMRIB Analysis Group, Oxford
Usage:
bet2 <input_fileroot> <output_fileroot> [options]
Optional arguments (You may optionally specify one or more of):
       -o, --outline generate brain surface outline overlaid onto original image
       -m,--mask <m> generate binary brain mask
                      generate approximate skull image
       -s,--skull
       -n,--nooutput don't generate segmented brain image output
       -f <f>
                      fractional intensity threshold (0->1); default=0.5; smaller_
\mathop{\hookrightarrow}\mathsf{values} give larger brain outline estimates
       -g <g> vertical gradient in fractional intensity threshold (-1->1);
\rightarrow default=0; positive values give larger brain outline at bottom, smaller at top
       -r, --radius <r> head radius (mm not voxels); initial surface sphere is set to...
  half of this
```

```
-w,--smooth <r> smoothness factor; default=1; values smaller than 1 produce_

-more detailed brain surface, values larger than one produce smoother, less detailed_

-surface

-c <x y z> centre-of-gravity (voxels not mm) of initial mesh surface.

-t,--threshold -apply thresholding to segmented brain image and mask

-e,--mesh generates brain surface as mesh in vtk format

-v,--verbose switch on diagnostic messages

-h,--help displays this help, then exits
```

Reconstruction of the diffusion model

The reconstruction for diffusion model within pixels is one of the key topics in diffusion MRI research and it is also one of key modules of the software. At the current stage, we have integrated three modeling methods: one is the traditional Gaussian model (commonly known as DTI, diffusion tensor imaging), and the other two are for high angular resolution diffusion imaging (HARDI). For more detailed information please refer to our review paper [11] and [13].

DTI Reconstruction

The classical diffusion gradient sequence used in dMRI is the pulsed gradient spin-echo (PGSE) sequence proposed by Stejskal and Tanner. This sequence has 900 and 1800 gradient pulses with duration time δ and separation time Δ . To eliminate the dependence of spin density, we need at least two measurements of diffusion weighted imaging (DWI) signals, i.e. S(b) with the diffusion weighting factor b in Eq. (1) introduced by Le Bihan etc, and S(0) with b = 0 which is the baseline signal without any gradient.

In the b-value in Eq. (1), $\qquad I \in \mathbb{C}^{1}$ is the proton gyromagnetic ratio, $\left|\left\{G\right\}\right\|$ is the diffusion sensitizing gradient pulse with norm $\left\|\left\{bf G\right\}\right\|$ and direction $\left\{bf u\right\}$. $\left|\left\{a\right\}\right\}$ is normally used to describe the effective diffusion time. Using the PGSE sequence with S(b), the diffusion weighted signal attenuation E(b) is given by Stejskal-Tanner equation,

where D is known as the apparent diffusion coefcient (ADC) which reects the property of surrounding tissues. Note that in general ADC D is also dependent on G in a complex way. However, free diffusion in DTI assumes D is only dependent on the direction of G, i.e. The early works in dMRI reported that ADC D is dependent on gradient direction u and used two or three DWI images in different directions to detect the properties of tissues. Then Basser et al. introduced diffusion tensor [12] to represent ADC as $D(bf{u}) = {bf u^{T}}{bf D}bf{u}s$, where ${bf D}$ is called as the diffusion tensor, which is a 3×3 symmetric positive definite matrix independent of ${bf u}$. This method is called as diffusion tensor imaging (DTI) and is the most common method nowadays in dMRI technique. In DTI, the signal E(b) is represented as

The diffusion tensor D can be estimated from measured diffusion signal samples through a simple least square method or weighted least square method [12], or more complex methods that consider positive denite constraint or Rician noise. If single b-value is used, the optimal b-value for tensor estimation was reported to in the range of $(0.7 \sim 1.5) \times 10^{43}$ s/mm², and normally about twenty DWI images are used in DTI in clinical study. ome useful indices can be obtained from tensor D. The most important three indices are fractional anisotropy (FA), mean diffusivity (MD) and relative anisotropy (RA) dened as

where, are the three eigenvalues of D and is the mean eigenvalue. MD and FA have been used in many clinical applications. For example, MD is known to be useful in stroke study. For more detailed information please refer to our review paper [13].

```
$ ./bndti_estimate -h
bndti_estimate: Diffusion Tensors Estimation.
DiffusionKit (v1.3), http://diffusion.brainnetome.org/.
(Jun 24 2016, 09:40:15)
```

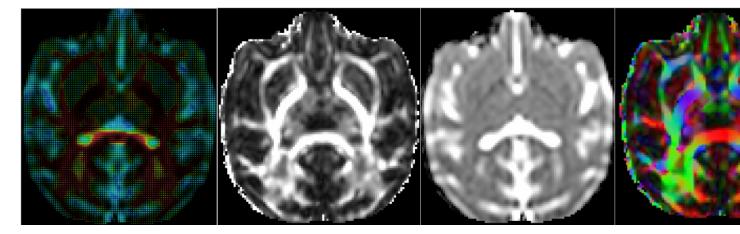


Fig. 3: Figure 4. Tensor field and the scalar maps estimated from a monkey data with b = 1500s/mm2.

general argu	uments	
-d		Input DWI Data, in NIFTI/Analyze format (4D)
-g		Gradients direction file
-b		b value file
-m		Brain mask : filein mask OPTIONAL
-0	dti	Result DTI : fileout prefix; 'dti' by default
-tensor	0	Save tensor : 0 - No; 1 - Yes; (Default: 0)
-eig	0	Save eigenvalues and eigenvectors : 0 - No; 1 - Yes;
\hookrightarrow (Default:	0)	

SPFI Reconstruction

It was proposed that the SPFI method has more powerful capability to identify the tangling fibers [8]. In SPFI [8], the diffusion signal is represented by spherical polar Fourier (SPF) basis functions in Eq. 7.

The SPF basis denoted by $R_{n}(\|q\|)Y_{1}^{m}(u)\$ is a 3D orthonormal basis with spherical harmonics in the spherical portion and the Gaussian-Laguerre function in the radial portion. Furthermore, Cheng and his colleagues proposed a uniform analytical solution to transform the coefficients $a_{mn}\$ of $E(q)\$ to the coefficients $c_{mn}\$ of $E(q)\$ to the coefficients $c_{mn}\$ of Phi_w of ODF (orientation distribution function) represented by the spherical harmonics basis, as in Eq. 8.

It is a model-free, regularized, fast and robust reconstruction method which can be performed with single-shell or multiple-shell HARDI data to estimate the ODF proposed by Wedeen et al. [14]. The implementation of analytical SPFI includes two independent steps. The first estimates the coefficients of E(q) with least squares, and the second transforms the coefficients of E(q) to the coefficients of ODF.

(continues on next page)

(continued from previous page)

-scale	-1	The scale parameter of Spherical Polar Fourier Basis.
→The default i	s calculated	by the program.
-tau	0.02533	The diffusion time of DWI data.
-outGFA	false	Whether output generalized FA. True means Yes.
-rdis	0.015	The radius value for EAP profile.
-sh	4	Order of spherical basis
-ra	1	Order of radial basis
-lambda_sh	0	Regularization parameter for spherical basis
-lambda_ra	0	Regularization parameter for radial basis

CSD Reconstruction

The CSD method was proposed by Tournier et al. [9], which expresses the diffusion signal as in Eq. 9,

where \$F(\theta,\Phi)\$ is called the fiber orientation density function (fODF), which needs to be estimated, and \$R(\theta)\$ is the response function, which is the typical signal generated from one fiber. The response function can be directly estimated from diffusion weighted image (DWI) data by measuring the diffusion profile in the voxels with the highest fractional anisotropy values, indicating that a single coherently oriented fiber population is contained in these voxels. When the response function is obtained, we can utilize the deconvolution of \$R(\theta)\$ from \$F(\theta,\Phi)\$ to estimate the fiber ODF. The computation of the fiber ODF was carried out using the software MRtrix (J-D Tournier, Brain Research Institute, Melbourne, Australia, http://www.brain.org.au/software/). We thank Dr. Jacques-Donald Tournier for sharing MATLAB code of the CSD method, which inspired an efficient C/C++ implementation.

```
$ ./bnhardi_FOD_estimate -h
bnhardi_FOD_estimate: Constraind Spherical Deconvolution (CSD) based HARDI.
→reconstruciton.
DiffusionKit (v1.2), http://diffusion.brainnetome.org/.
(Jul 15 2015, 11:50:20)
general arguments
   -d
                                 Input DWI Data, in NIFTI/Analyze format (4D)
   -q
                                 Gradients direction file
   -b
                                 b value file
                                 Brain mask : filein mask | OPTIONAL
   -m
  -out.FA
                 1
                                 Whether to output the FA of DTI
  -0
                                 Result CSD Estimate (.nii.gz)
                                 6/8/10, Max order of the adopted harmonical base
  -lmax
                 8
  -fa
                 [0.75,0.95]
                               The FA thesshold considered as single fiber
  -erode
                 -1
                                 The unit is voxel: Remove the garbage near the
\rightarrow boundary of FA image,
                                 for better estimating response function
  -nIter
                 50
                                 Max iteration number before aborting
                                 The regularization weight for optimization
  -lambda
                 1
  -tau
                 0.1
                                 The threshold on the FOD amplitude used to identify.
→negative lobes
  -hr
                 300
                                 300/1000/5000. The later get more acurate estimation.
→while more
                                 time consuming, so use the first one unless your_
⇔computer
                                 is powerful !
bnhardi_FOD_estimate: Constraind Spherical Deconvolution (CSD) based HARDI,
→reconstruciton.
DiffusionKit (v1.3), http://diffusion.brainnetome.org/.
(Jun 24 2016, 09:40:29)
general arguments
   -d
                                Input DWI Data, in NIFTI/Analyze format
                                                                         (4D)
```

-g	Gradients direction file
-b	b value file
-m	Brain mask : filein mask OPTIONAL
-outFA 1	Whether to output the FA of DTI
-0	Result CSD Estimate (.nii.gz)
-lmax 8	6/8/10, Max order of the adopted harmonical base
-fa [0.75,0.	95] The FA thesshold considered as single fiber
-erode -1	The unit is voxel: Remove the garbage near the
→boundary of FA image, for	better estimating response function
-nIter 50	Max iteration number before aborting
-lambda 1	The regularization weight for optimization
-tau 0.1	The threshold on the FOD amplitude used to identify_
⇔negative lobes	
-hr 300	300/1000/5000. The later get more acurate estimation_
→while more time consuming	, so use the first one unless your computer is powerful !
-omp 2	Max number of threads

Fiber tracking and attributes extraction

Fiber tracking is a critical way to construct the anatomical connectivity matrix. For the tracking based on tensors from DTI, an intuitive way is to link the neighboring voxels following their main directions (e.g. V1 in the eigenvector of the DTI) given a set of some stop criteria, such as maximum bending angle of the curve and minimum FA value, which is to ensure the target voxel is indeed white matter microstructure. This is the so called deterministic streamline tractography [15], as illustrated in Figure 5.

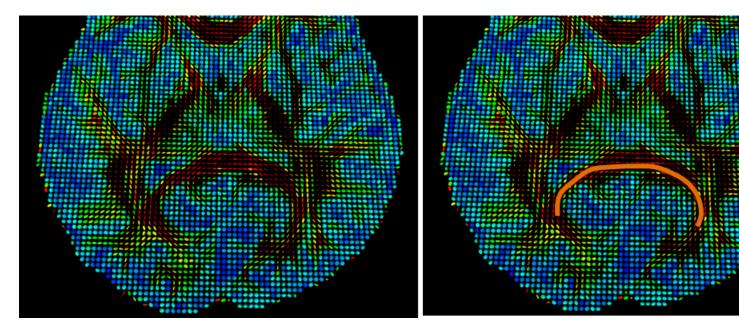


Fig. 4: Figure 5. Illustration for deterministic streamline tractography.

-m		Mask Image.
-s		Seeds Image.
-fa		FA Image.
-invert	0	Invert x: 1; Invert y: 2; Invert z: 3.
-swap	0	Swap x/y: 1; Swap x/z: 2; Swap y/z: 3.
-ft	0.1	FA Threshold.
-at	45	Angular Threshold.
-sl	0.5	Step Length (Voxel).
-min	10	Threshold the fiber (mm). (remove fibers shorter than the $\!\!\!\!\!$
→number)		
-max	5000	Upper-threshold the fiber (mm). (remove fibers longer than \Box
→the number)		
-0		Output Fibers Filename (.trk file).

The tracking module in the software for HARDI estimation is similar to the streamline method as described above. It should be kept in mind that, for HARDI estimation, usually there are more than one main direction (which is what we desired since it possibly identifies tangling fibers), so we should consider these kissing/branching cases. Meanwhile, since there is no explicit dominant directions for each voxel, searching algorithm should be applied to locate the main directions. The searching should run for each voxel, so it is not as fast as the traditional DTI tracking.

```
$ ./bnhardi_tracking -h
bnhardi_tracking: HARDI Deterministic Fibertracking.
DiffusionKit (v1.3), http://diffusion.brainnetome.org/.
(Jun 24 2016, 09:40:28)
                              HARDI spherical harmonic image.
   -d
   -fa
                              FA image.
   -m
                              Mask image.
                              Seeds image.
   -s
   -sl
                 0.5
                              Step length (Voxel): float.
                 0.1
                              FA threshold: float.
   -ft
                  60
                              Angle threshold: float.
   -at
                              Threshold the fiber (mm). (remove fibers shorter than \_
                  10
   -min
\rightarrowthe number)
                  5000
                              Upper-threshold the fiber (mm). (remove fibers longer,
   -max
\rightarrow than the number)
                              Invert x: 1; Invert y: 2; Invert z: 3.
   -invert
                 0
                              Swap x/y: 1; Swap x/z: 2; Swap y/z: 3.
   -swap
                  0
   -threshold
                 0.25
                              Threshold for the peaks of ODF.
                  1
                              Max number of threads.
   -omp
                              Tract file (.trk file).
   -0
```

Visualization for various images

The software provides a variety of entries for visualizing different data types, including 3D/4D image, Tensor/ODF/FOD image and white mater fibers. The views of different data could be superimposed for a precise anatomical localization. It should be noted that, if one uses the GUI, these view functions could be called in each processing steps for inspecting the results.

Figure 6. Illustrations for the capability of the software to show many types of images.

```
bnviewer [[-volume] DTI_FA.nii.gz]
          [-roi ROI/roi_cc_top.nii.gz]
          [-fiber ROI/roi_cc_top.trk]
          [-tensor DTI.nii.gz]/[-odf HARDI.nii.gz]
options:
 -help
                       show this help
 -volume
         .nii.gz
                       set input background data
 -roi
          .nii.gz
                      set input ROI data
 -fiber
          .trk
                      set input fiber data
 -tensor .nii.gz
                     set input DTI data (conflict with -odf args)
 -odf
          .nii.gz
                       set input ODF/FOD data (conflict with -tensor args)
```

Several other useful tools

Image registration

This module is implemented by NiftyReg, which is an open-source software for efficient medical image registration. It has been mainly developed by members of the Translational Imaging Group with the Centre for Medical Image Computing at University College London, UK [10]. In our software, the registration module is customized to autoconfigure the parameter settings for different image modalities, e.g., two DWI images (for eddy current correction in current version), standard space and T1 space (for mapping ROIs to the individual space), DWI space and standard space (for statistical comparisons across subjects). This module contains several commands reg_aladin is the command for rigid and affine registration which is based on a block-matching approach and a Trimmed Least Square (TLS) scheme [18] [19]. reg_f3d is the command to perform non-linear registration which is based on the Free-From Deformation presented by Rueckert et al. [20]. reg_resample is been embedded in the package. It uses the output of reg_aladin and reg_f3d to apply transformation, generate deformation fields or Jacobian map images for example.

```
$ ./reg_aladin -h
Block Matching algorithm for global registration.
Based on Ourselin et al., "Reconstructing a 3D structure from serial histological.
→sections", Image and Vision Computing, 2001
Usage: reg_aladin -ref <filename> -flo <filename> [OPTIONS].
       -ref <filename> Reference image filename (also called Target or Fixed)...
\rightarrow (mandatory)
       -flo <filename> Floating image filename (also called Source or moving)...
\rightarrow (mandatory)
OPTIONS
                               The symmetric version of the algorithm is used by
       -noSvm
→default. Use this flag to disable it.
                               To perform a rigid registration only. (Rigid+affine by,
       -rigOnly
→default)
       -affDirect
                              Directly optimize 12 DoF affine. (Default is rigid,

→initially then affine)

       -aff <filename>
                              Filename which contains the output affine
→transformation. [outputAffine.txt]
                         Filename which contains an input affine transformation.
       -inaff <filename>
→ (Affine*Reference=Floating) [none]
                          Filename of a mask image in the reference space.
       -rmask <filename>
       -fmask <filename>
                             Filename of a mask image in the floating space. (Only_
→used when symmetric turned on)
```

```
-res <filename>
                              Filename of the resampled image. [outputResult.nii]
      -maxit <int>
                              Maximal number of iterations of the trimmed least,
\leftrightarrow square approach to perform per level. [5]
      -ln <int>
                              Number of levels to use to generate the pyramids for
\rightarrow the coarse-to-fine approach. [3]
      -lp <int>
                              Number of levels to use to run the registration once_
→the pyramids have been created. [ln]
      -smooR <float>
                              Standard deviation in mm (voxel if negative) of the
\hookrightarrowGaussian kernel used to smooth the Reference image. [0]
      -smooF <float> Standard deviation in mm (voxel if negative) of the
\hookrightarrowGaussian kernel used to smooth the Floating image. [0]
      -refLowThr <float> Lower threshold value applied to the reference image.
→ [0]
      -refUpThr <float>
                             Upper threshold value applied to the reference image.
→ [0]
      -floLowThr <float>
                            Lower threshold value applied to the floating image.
→ [0]
      -floUpThr <float> Upper threshold value applied to the floating image.
→ [0]
      -nac
                              Use the nifti header origin to initialise the,
→transformation. (Image centres are used by default)
      -coa
                              Use the input masks centre of mass to initialise the
→transformation. (Image centres are used by default)
                              Interpolation order to use internally to warp the
      -interp
\rightarrow floating image.
      -iso
                              Make floating and reference images isotropic if
⇔required.
      -pv <int>
                              Percentage of blocks to use in the optimisation scheme.

→ [50]

      -pi <int>
                              Percentage of blocks to consider as inlier in the
→optimisation scheme. [50]
      -speeeed
                              Go faster
      -omp <int>
                               Number of thread to use with OpenMP. [4]
      -voff
                              Turns verbose off [on]
```

\$./reg f3d -h Fast Free-Form Deformation algorithm for non-rigid registration. Based on Modat et al., "Fast Free-Form Deformation using graphics processing units", →CMPB, 2010 Usage: reg_f3d -ref <filename> -flo <filename> [OPTIONS]. -ref <filename > Filename of the reference image (mandatory) -flo <filename> Filename of the floating image (mandatory) OPTIONS Initial transformation options (One option will be considered): -aff <filename> Filename which contains an affine transformation → (Affine*Reference=Floating) -incpp <filename> Filename ofloatf control point grid input Output options: Filename of control point grid [outputCPP.nii] -cpp <filename> -res <filename> Filename of the resampled image [outputResult.nii] Input image options: -rmask <filename> Filename of a mask image in the reference space (continues on next page)

```
-smooR <float>
                                    Smooth the reference image using the specified.
⇔sigma (mm) [0]
      -smooF <float>
                                    Smooth the floating image using the specified
⇔sigma (mm) [0]
                                    Lower threshold to apply to the reference_
      --rLwTh <float>
--rUpTh <float>
                                   Upper threshold to apply to the reference,
--fLwTh <float>
                                   Lower threshold to apply to the floating image_
→intensities [none]. Identical value for every timepoint.*
      --fUpTh <float>
                                   Upper threshold to apply to the floating image_
⇔intensities [none]. Identical value for every timepoint.*
      -rLwTh <timepoint> <float> Lower threshold to apply to the reference.

→image intensities [none] *

      -rUpTh <timepoint> <float> Upper threshold to apply to the reference_

→image intensities [none] *

      -fLwTh <timepoint> <float> Lower threshold to apply to the floating image_
→intensities [none] *
      -fUpTh <timepoint> <float> Upper threshold to apply to the floating image_
⇔intensities [none] *
Spline options:
      -sx <float>
                            Final grid spacing along the x axis in mm (in voxel if.
→negative value) [5 voxels]
      -sy <float>
                            Final grid spacing along the y axis in mm (in voxel if
→negative value) [sx value]
      -sz <float>
                            Final grid spacing along the z axis in mm (in voxel if.
→negative value) [sx value]
Regularisation options:
                            Weight of the bending energy penalty term [0.005]
      -be <float>
      -le <float> <float>
                            Weights of linear elasticity penalty term [0.0 0.0]
      -12 <float>
                            Weights of L2 norm displacement penalty term [0.0]
      -jl <float>
                            Weight of log of the Jacobian determinant penalty term.
\rightarrow [0.0]
      -noAppJL
                            To not approximate the JL value only at the control.
→point position
Measure of similarity options:
NMI with 64 bins is used expect if specified otherwise
      --nmi
                            NMI. Used NMI even when one or several other measures,
→are specified.
                          NMI. Number of bin to use for the reference image_
      --rbn <int>
⇔histogram. Identical value for every timepoint.
      --fbn <int> NMI. Number of bin to use for the floating image_
⇔histogram. Identical value for every timepoint.
      -rbn <tp> <int> NMI. Number of bin to use for the reference image.
→histogram for the specified time point.
                           NMI. Number of bin to use for the floating image_
      -rbn <tp> <int>
→histogram for the specified time point.
      --lncc <float>
                           LNCC. Standard deviation of the Gaussian kernel.
→ Identical value for every timepoint
      -lncc <tp> <float>
                          LNCC. Standard deviation of the Gaussian kernel for
→the specified timepoint
      --ssd
                           SSD. Used for all time points
      -ssd <tp>
                           SSD. Used for the specified timepoint
      --kld
                           KLD. Used for all time points
```

```
KLD. Used for the specified timepoint
      -kld <tp>
                              To use the additive NMI for multichannel data,
       -amc
\hookrightarrow (bivariate NMI by default)
Optimisation options:
      -maxit <int>
                              Maximal number of iteration per level [300]
       -ln <int>
                              Number of level to perform [3]
                              Only perform the first levels [ln]
      -lp <int>
      -nopy
                             Do not use a pyramidal approach
                              To not use the conjuage gradient optimisation but a
      -noConj
⇔simple gradient ascent
      -pert <int>
                             To add perturbation step(s) after each optimisation
⇔scheme
F3D2 options:
      -vel
                              Use a velocity field integration to generate the
⇔deformation
      -fmask <filename>
                         Filename of a mask image in the floating space
OpenMP-related options:
      -omp <int>
                              Number of thread to use with OpenMP. [4]
Other options:
      -smoothGrad <float> To smooth the metric derivative (in mm) [0]
      -pad <float>
                             Padding value [nan]
      -voff
                             To turn verbose off
$ ./reg_resample -h
Usage: reg_resample -ref <filename> -flo <filename> [OPTIONS].
      -ref <filename>
                         Filename of the reference image (mandatory)
      -flo <filename>
                           Filename of the floating image (mandatory)
OPTIONS
      -trans <filename> Filename of the file containing the transformation_
→parametrisation (from reg_aladin, reg_f3d or reg_transform)
      -res <filename> Filename of the resampled image [none]
                         Filename of the resampled blank grid [none]
       -blank <filename>
      -inter <int> Interpolation order (0, 1, 3, 4)[3] (0=NN, 1=LIN; 3=CUB,
\rightarrow 4 = SINC)
      -pad <int> Interpolation padding value [0]
-tensor The last six timenoints of the
                          The last six timepoints of the floating image are
      -tensor
→considered to be tensor order as XX, XY, YY, XZ, YZ, ZZ [off]
      -psf
                          Perform the resampling in two steps to resample an image.
→to a lower resolution [off]
```

Need further help?

-voff

Check the website http://cmictig.cs.ucl.ac.uk/wiki/index.php/NiftyReg.

Image calculation and ROI generation

The module bncalc provides simple image calculations, such as add/minus/multiply/divide operations. Meanwhile, it can also generate user-defined ROIs given the origin and radius in a user-specified image space.

Turns verbose off [on]

```
$ ./bncalc -h
DiffusionKit (v1.3), http://diffusion.brainnetome.org/.
This funciton provide basic process for the input data (NIFTI/Analyze format)
Usage of bncalc:
                          The original file you want to manage.
  - i
         image
  -add image/value Add to the data from the last step.
-sub image/value Subtract data from the last step.
  -mul image/value Multiply the data from the last step.
  -div
          image/value Divide the data from the last step.
        x,y,z,r
  -roi
                          Generate a ROI centered at [x,y,z] (MNI mm) with radius r,...
\rightarrow in the input data space.
  -roi_rect x1,x2,y1,y2,z1,z2 OR i
                           Generate a cuboid ROI by specifying left-bottom
                           corner [x1,y1,z1] and right-up corner [x2,y2,z2].
                           If only one integer is specified, we assume you
                           want to get the i-the volume along the 4th dimension (from
\rightarrow 0).
                           Mask the data from last step by this input one.
  -mask
            image
                           If this input is a binary, then it is the same as
                           -mul, otherwise it keep the voxels from the last step
                           when the new input is nonzero.
   -bin
                          set 1 if >value, otherwise 0.
            value
   -ut.hr
            value
                          Set voxel=0 when it>value.
   -dthr
            value
                          Set voxel=0 when it<value.
   -0
            image
                          output a NIFTI (.nii.gz) file
```

Fiber manipulation

After obtaining a large bundle of white matter fibers, you may want to prune the fibers that go through specified locations (ROIs). Here, we provide several tools to manipulate the fiber bundles. bnfiber_prune, to split fiber bundles based on given ROIs.

```
$ ./bnfiber_prune
bnfiber_prune: Prune fiber bundles based on given ROIs.
DiffusionKit (v1.3), http://diffusion.brainnetome.org/.
(Jun 24 2016, 09:40:29)
  -fiber Tract file (.trk file)
  -and AND ROI file: rol.nii.gz,roi2.nii.gz
  -or OR ROI file: rol.nii.gz,roi2.nii.gz
  -not NOT ROI file: rol.nii.gz,roi2.nii.gz
  -o Output Tract file (.trk file).
```

bnfiber_end, to cut the fiber bundles given start/stop ROIs, which is useful to get the exact connections between two ROIs in constructing connectivity matrix.

```
$ ./bnfiber_end -h
bnfiber_end: Extract fibers which end in the two given rois.
DiffusionKit (v1.3), http://diffusion.brainnetome.org/.
(Jun 24 2016, 09:40:29)
    -fiber Tract file (.trk file).
    -roi1 ROI1 file
    -roi2 ROI2 file
    -o Output tract file (.trk file)
```

bnfiber_stats, to extract statistical properties of the fiber bundle, such as mean FA/MD and fiber numbers.

```
$ ./bnfiber_stats -h
bnfiber_stats: Show fiber stats.
DiffusionKit (v1.3), http://diffusion.brainnetome.org/.
(Jun 24 2016, 09:40:33)
        -fiber Input tract file (.trk file).
```

bnfiber_map, to compute the fiber density map which is used in track density imaging [16].

bnmerge / bnsplit, to merge the 3D volumes to 4D or split 4D to 3D volumes.

```
$ ./bnmerge -h
bnmerge: Merge 3D NIfTI files to 4D NIfTI file.
This program is to merge 3D NIfTI files to 4D NIfTI file.
Usage: bnmerge filein1 filein2 ... fileout
options: -h : show this help
```

bninfo, to display a short head information of the input image. Supported input image format includes NIFTI/DICOM.

1.1.3 GUI Front-end of Data Processing Pipeline

Preprocessing

A 'Confirmation' option is provided in each of the following preprocessing step. When confirmation is required, the program displays the processing result after the target file is generated to ensure the correctness the final result.

DICOM to NIFTI Conversion

This is the GUI front end for dcm2nii program. Details can be found at DICOM to NIFTI Conversion section.

DICOM to NIFTI	
Input Data Dir	rdiffusion/data/NC_01_0050/DWI
Output Data File:	sion/data/NC_01_0050/DWI.nii.gz
Confirmation	

Eddy Current Correction

This is the GUI front end for bneddy program. Details can be found at Data correction section.

Eddy Current Correction				
Input Data File:	sion/data/NC_01_0050/DWI.nii.gz			
Output Data File:	ata/NC_01_0050/DWI_eddy.nii.gz			
Confirmation	Reference 0			

Since this is a time-consuming step in preprocessing stage, one can skip this step ignoring potential image registration error in original DWI data by unchecking the option. Then the converted NIFTI image would be input of brain extraction tool directly if the skull stripping option is enabled.

Skull Stripping

FSL's BET (Brain Extraction Tool) is integrated as part of the program, so that one can verify automatic skull stripping result by visualizing overlaying mask image upon original image. See *Volume Image Overlay* on how to generate overlaid background images.

✓ Skull Stripping	
Input File:	ata/NC_01_0050/DWI_eddy.nii.gz
Brain File:	C_01_0050/DWI_eddy_brain.nii.gz
Mask File:	[_01_0050/DWI_eddy_mask.nii.gz]
Confirmation	Threshold 0.50 🗘

A threshold value that indicate 'fractional intensity' is exposed on the GUI, so that one can adjust the threshold to get smaller/larger brain outline estimates. Details on the implement can be found at BET's UserGuide.

Diffusion Model Reconstruction

Starting from this section, we provide three different approaches for diffusion MR data processing:

This is the GUI front end for *Reconstruction of the diffusion model*, which generates required model for *Fiber tracking and attributes extraction* from DWI image and related b-values.

Click one of the options above to see guidelines on using the specific method for data processing.

Output Type: DTI 😂 🗹 Eigen Data				
Data Input				
DWI [e/mrdiffusion/data/NC_01_0050/DWI.nii.gz]				
Bvals ce/mrdiffusion/data/NC_01_0050/DWI.bval				
Bvecs ce/mrdiffusion/data/NC_01_0050/DWI.bvec				
Mask [iffusion/data/NC_01_0050/DWI_mask.nii.gz]				
Prefix ace/mrdiffusion/data/NC_01_0050/DWI_dti				
Reset				

Fig. 5: GUI front-end for DTI Reconstruction using bndti_estimate

Once DWI data is selected through file selection dialog, all other input and output options would be automatically generated. This simplifies the usage of command-line program, which requires a list of options to be entered.

Fiber Tracking

To generate fiber tracking result using previously generated diffusion model, we provide the GUI front-end for using bndti_tracking and bnhardi_tracking, which generate tract files from DTI and HARDI respectively. The details of the program are described on *Fiber tracking and attributes extraction* section.

Fiber tracking require seed(ROI) files to initialize a tracking process. One can either generate the seed files from the **Registration Tool** we provided from **Tools** menu, or split a brain atlas image (e.g. AAL) using bncalc or bnroisplit into multiple ROI files. To generate whole brain fiber, just use the brain mask image generated during *Skull Stripping* as seed input. This may cost a few minutes when HARDI is used as input diffusion model.

Miscellaneous Tools

Image Registration

This is basically the GUI front-end of NiftyReg, while we perform reg_aladin on intra-subject images, and addtionally run reg_f3d for inter-subject cases.

The details of the tool is described in Image Registration section in UserGuide page.

Output Type: HARDI 😂				
Data Input				
DWI	e/mrdiffusion/data/NC_01_0050/DWI.nii.gz			
Bvals	ce/mrdiffusion/data/NC_01_0050/DWI.bval)		
Bvecs	ce/mrdiffusion/data/NC_01_0050/DWI.bvec)		
Mask	iffusion/data/NC_01_0050/DWI_mask.nii.gz)		
Data Output Prefix ce/mrdiffusion/data/NC_01_0050/DWI_spfi				
Reconstrut Parameters				
Method SPFI (ODF) C Generate Direction				
SH Order 4 🗘 SH Lambda 0.00 🗘				
Rad Ord	er 1 🗘 Rad Lambda 0.00)		
Amptitud	de 0.02533 🗘 Radius 0.015 🚭)		
🗌 Gen	erate GFA			

Fig. 6: GUI front-end for SPFI Reconstruction using bnhardi_ODF_estimate

Output Type: HARDI 😂					
Data Inpu	t				
DWI	e/mrdiffusion/data/NC_01_0050/DV	VI.nii.gz 📖			
Bvals	ce/mrdiffusion/data/NC_01_0050/D	WI.bval			
Bvecs	Bvecs ce/mrdiffusion/data/NC_01_0050/DWI.bvec				
Mask	Mask [iffusion/data/NC_01_0050/DWI_mask.nii.gz]				
Data Outp	Data Output				
Prefix ce/mrdiffusion/data/NC_01_0050/DWI_fod					
Reconstrut Parameters					
Method	Method CSD (FOD) C Generate Direction				
SH Order	r 8 🗘 FA Range	0.75			
Max Itera		0.95 🗘			
Regulariz	ation 1.00 🔶 Resolution	300 🛛 🗘			
Amptitud	e 0.10 🗘				

Fig. 7: GUI front-end for CSD Reconstruction using bnhardi_FOD_estimate

Data Type	HARDI I I			
Data Inpu	Data Input			
Data	sion/data/NC_01_0050/DWI_spfi_ODF.nii.gz			
Mask	diffusion/data/NC_01_0050/DWI_mask.nii.gz)			
FA	ffusion/data/NC_01_0050/DWI_spfi_FA.nii.gz			
Seeds	diffusion/data/NC_01_0050/DWI_mask.nii.gz			
Data Out	put			
Fiber (Fiber [/mrdiffusion/data/NC_01_0050/DWI_mask.trk]			
Tracking I	Parameters			
Angle thr	reshold 45 🗘 FA Threshold 0.10 🗘			
Step size	Step size (voxel) 0.5			
Min Lengt	th (mm) 10 🗘 Max length 5000 🗘			

Image Registration 🗙			
Options Intr	a-Subject O Inter-Subject		
Source	ome/liangfu/workspace/mrdiffusion/data/NC_01_0050/DWI_brain.nii.gz		
Output	ne/liangfu/workspace/mrdiffusion/data/NC_01_0050/DWI_brain_r.nii.gz		
Reference	/home/liangfu/workspace/mrdiffusion/data/NC_01_0050/T1_brain.nii.gz		
	<u>Cancel</u> <u>O</u> K		

Create ROI

The details of the tool is described in Image calculation and ROI generation section in UserGuide page.

Create ROI			
ROI Shape	ROI Location		
O Circular ROI	Xmin (mm) 0.00 \$\$\$ Ymin (mm) 0.00 \$\$\$ Zmin (mm) 0.00 \$\$\$\$		
Rectangular ROI	Xmax (mm) 0.00 \$\$\$ Ymax (mm) 0.00 \$\$\$ Zmax (mm) 0.00 \$\$\$\$		
Source	/home/liangfu/workspace/mrdiffusion/data/NC_01_0050/T1_brain.nii.gz		
Output	/home/liangfu/workspace/mrdiffusion/data/NC_01_0050/T1_brain_roi.nii.gz		
	<u>Cancel</u> <u>O</u> K		

Fiber Pruning

The details of the tool is described in Fiber manipulation section in UserGuide page.

	Fiber Pruning		
Input Fiber	(/home/liangfu/workspace/mrdiffusion/data/NC_01_0050/DWI_mask.trk		
ROIs: AND	<pre>/path/roi_and1.nii.gz,/path/roi_and2.nii.gz</pre>		
ROIs: OR	/path/roi_or1.nii.gz,/path/roi_or2.nii.gz		
ROIs: NOT	<pre>/path/roi_not1.nii.gz,/path/roi_not2.nii.gz,/path/roi_not3.nii.gz</pre>		
Output Fiber	me/liangfu/workspace/mrdiffusion/data/NC_01_0050/DWI_mask_prune.trk		
	<u>Cancel</u> <u>O</u> K		

1.1.4 Guidelines on Data Visualization

In this page, the general functionality implemented in the bnviewer program is described in detail. This covers methods that a user can visualize almost all kinds of diffusion MRI data.

Except those specially noticed, all the data should be formatted in either NIFTI or ANAYLYZE image. NIFTI format is recommended, see *Data Format* page for more details.

Command Line Interface

Typing bnviewer -h in command line would given following information, which help users to load a list of files without click them one by one. The can be extremely useful when one wants to contantly load a large list of data files to get screen capture images.

```
basic uage:
    bnviewer [[-volume] DTI_FA.nii.gz]
        [-roi ROI/roi_cc_top.nii.gz]
```

```
[-fiber ROI/roi_cc_top.fiber]
           [-tensor DTI.nii.gz]/[-odf HARDI.nii.gz]
           [-atlas DTI.nii.gz]
options:
 -help
                        show this help
 -volume
          .nii.gz
                       set input background data
 -roi
          .nii.gz
                       set input ROI data
 -fiber
          .fiber/.trk set input fiber data
 -tensor .nii.gz set input DTI data (conflict with -odf args)
 -odf
          .nii.gz
                        set input ODF/FOD data (conflict with -tensor args)
```

Todo: create interface for generating high quality images and save them into specified location.

Image Data Navigation

3D Navigation

When an image data is loaded, one can change view angle by dragging *left* mouse button. This is the default interactor style implemented in *vtkInteractorStyleTrackballCamera*, which is used internally in our program. According to VTK's documentation:

vtkInteractorStyleTrackballCamera allows the user to interactively manipulate (rotate, pan, etc.) the camera, the viewpoint of the scene. In trackball interaction, the magnitude of the mouse motion is proportional to the camera motion associated with a particular mouse binding. For example, small left-button motions cause small changes in the rotation of the camera around its focal point. For a 3-button mouse, the left button is for *rotation*, the right button for *zooming*, the middle button for *panning* (translation), and ctrl + left button for *spinning*. (With fewer mouse buttons, ctrl + shift + left button is for zooming, and shift + left button is for panning.)

We simplify the trackball interaction to used middle button for both panning and zooming, in order to spare right mouse click to popup option menu for more actions.

The background color for 3D view is defined as black by default. This can be changed from context menu popped up when *right* mouse click event is captured. By selecting the color from the popup color dialog, the background color is changed instantly.

Also, saving screenshots as PNG image is enabled from the context menu. The default image is named after current timestamp to avoid naming duplication.

Screenshot Options			
Location: WI/bnviewer-160225-143612.png			
Screenshot options			
Apply Transparent Background			
Set Magnification Rate 1			
<u>C</u> ancel <u>O</u> K			

To customize generated image for different requirements, two additional options are provided, as shown in the above screenshot. The higher magnification value indicate higher resolution of generated image. By enabling transparent background in output image, one can alternate the background color in a later stage.

2D Navigation

Three 2D slice image views are placed below 3D view by default. The three slice views from left to right are *sagittal plane, coronal plane* and *axial plane* respectively.

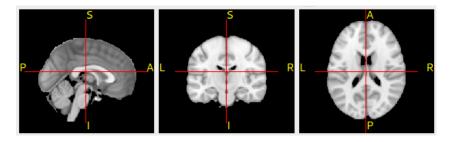


Fig. 8: *sagittal plane, coronal plane* and *axial plane* (left to right)

2D slice view widget sizes can be enlarged by switching the overall layout. This functionality is implemented in a single button, named **Switch Layout**, available on the toolbar. By toggling the button, the main view panels switches between 3D view port and a combination of sliced 2d views.

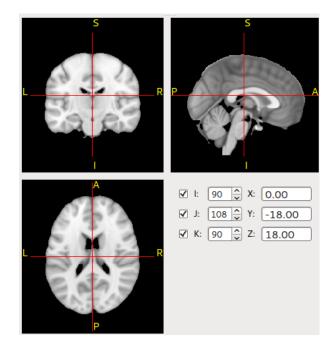


Image slice index can be changed by clicking on 2d image slice views. Besides, we perform radiological/neurological (or RAS/LAS) conversion intantly when slice index values are changed.

For people interested in more details about the radiological/neurological conversion, please refer to FSL's Orientation Explained .

Background Volume Data Layer

Beginning from this section, several kinds of 'layers' are described. Each 'layer' may be contrain one or more data that holds the same property. By default, the layers manager is located at top-right side of the window.

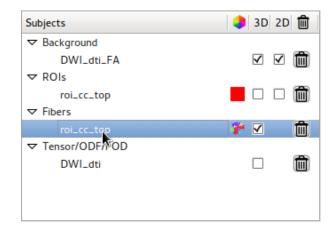


Fig. 9: The layers manager

We refer to *Background* data as plain 3D or 4D volume data that user wants to visualize its sliced views. The content should be either a typical 3D image (e.g. T1/T2 image) or a 4D image such as DWI, DTI, ODF and even fMRI data.

To load a Background Data, click **Load Background** in the menu or toolbar, and select a single volume image. Note that selecting multiple image is not valid here, because the program doesn't aware which background image is laid that the bottom that might be overlaid by subsequently loaded images.

Once background image is loaded, several basic data information is shown in the data property panel.

Property	Value		
✓ Information			
File name	mni152.nii.gz		
Data type	short		
Volumes	1		
Dimensions	181 x 217 x 181		
Spacing	1 x 1 x 1		
Value range	0~8364		
Volume Rendering	\checkmark		
Opacity	1.00		
Volume Index	0		
Intensity	0		
Min Value	0.0000		
Max Value	8364.0000		
	20		

Fig. 10: property panel showing mni152 as background image

The widget below the data property panel is the color table for *Volume Rendering*. The horizontal axis indicates intensity values of a target image, while the vertical axis of the widget indicate the opacity value that maps the intensity values onto visible planes in volume rendering. One can adjust opacity map (by dragging blue opacity nodes) as well as color map to get proper renderering results.

Image Contrast Enhancement

During loading a Background image, the minimum and maximum intensity values are calculated and displayed in the data property panel. By narrow down the range of image intensity values, one can the enhance image contrast.

Note that this functionality is currently applied for 2d sliced views only.

Volume Image Overlay

By loading multiple image throught **Load Background** menu, one can place one image over another, Therefore, it's easy to make slice-to-slice comparison to find out image registration errors by adjusting opacity values of the upper level image.

This feature can be quite useful for verifying Skull Stripping results. After performing skull stripping, ...

Todo: PUT OVERLAY IMAGE HERE!

Volume Rendering

Volume rendering of the background image can be enabled in *Data Property* panel. Once volume rendering is enabled, a color table is shown to enable users to adjust color map and opacity values that maps intensity values onto visible planes. This functionality is particularly useful for visualizing skull-stripped T1 image, where the folds that increase the surface area of the cortex can be directly visible without threshold-based surface extraction.

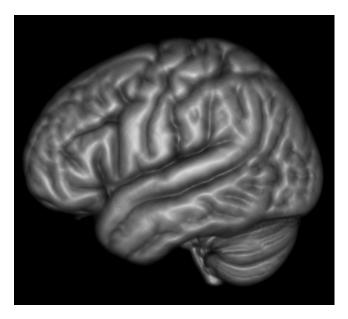


Fig. 11: volume rendering of mni152 data

Region of Interest (ROI) Layer

To describe the location of a specific region, we employ the ROI layer. It outlines surface of the region on 3d view, along with a filled area on 2d slice views.

To open a ROI file, click on 'Load ROI' from either menu or toolbar. Note that loading multiple ROI data files that located within the same directory is allowed. Therefore, one don't have load splitted brain atlas files one-by-one.

To change the color of a selected ROI files, set Color to different value from the data properties panel. This is useful when more than one ROI file is loaded, and it's hard to distinguish them by appearance.

To adjust transparency of the surface or the filled area in 2d views, set opacity value of the selected fiber file to a value range form 0 to 1.

Todo: compute the voxel-level volume size of a region, as well as its size in millimeters (by multiplying spacing).

Fiber Layer

To display deterministic tractography results, the program generates and displays .fiber/.trk files in either lines or tubes. One can load fiber files by selecting *Load Fiber* either from menu or toolbar. The definition of .fiber files is available at *Data Format* page. The specification of .trk file format is available at TrackVis.

Property	Value
Number of Fibers	180
Mean Length	112.086
Total Volume	33504
Mean FA	0.456426
Mean MD	n/a
Min Length	23.71
Max Length	99.99
Render Type	Line
Color Code	[0,255,0]
Opacity	1.00

Fig. 12: Fiber data properties panel

Basic fiber data statistics is available on data properties panel once an arbitrary tract file is loaded. The statistics information include number of fibers in a tract, average length of fibers and total volume covered by the tract etc. These information are generated during fiber tracking process, and there is extra computation during online visualization.

Several display options are available to provided high quality rendering results. For performance reasons, tracts are rendered as slim lines by default. Therefore, detail shapes of the fibers are not clearly visible. However, location

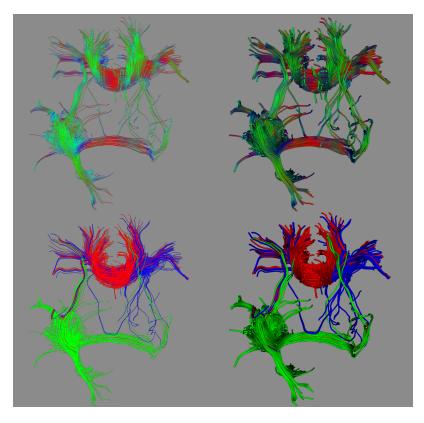


Fig. 13: A comparison of fibers rendered in different render types.

and orientation information of the lines are quite clear. By adjusting **Color Code** and **Render Type**, one can easily visualize multiple fiber files. On the above figure, two images on the left are fibers rendered as lines, while the two images on the right are rendered as tubes. Images on the top row are fibers rendered with directional coloring, while fibers on the bottom row are signed to a single color.

Note: Note that rendering whole brain fiber in 'Tube' rendering type can be quite slow for low performance computers. However, a progress bar would popup to show the current loading progress. Also, we recommand storing whole brain fiber files as .trk format instead of .fiber file to increase file loading efficiency.

Tensor/ODF/FOD Layer

Both diffusion tensor and ODF can be visualized within this layer. After adjusting rendering parameters, the **Render** button should be pressed in order to regenerate the scene. This prevents the time-consuming processing, every time when users are trying to change the configuration.

Diffusion tensor (DTI) file input should be a typical 4D NIFTI image, with exactly six volumes. The six values on each voxel represents entries in a 3x3 symmetric positive definite matrix. Numerical definition can be found at *DTI Reconstruction* section.

Diffusion orientation distribution function (ODF) file input should be a 4d image, too. However, it's fourth dimension should be one of 15,28,45,66,91 etc. If the format is incorrect, the program would refuse to load the image to avoid potential errors. Numerical definition can be found at *SPFI Reconstruction* and *CSD Reconstruction* section.

Due to performance reasons, ODF/FOD is rendered in low resolution by default. For researchers who need high-resolution images for publication quality results, an option in the properties panel is provided.

Property	Value	
✓ Information		
Render Type	ODF/FOD	10)
Slice Plane	Z Plane	10
Slice Index	25 🗘 Render	
Filp X	\checkmark	
Filp Y		
Filp Z		
Resolution	High	10
Color Code	Spherical	10
Direction	DWI_spfi_DIR	-

Fig. 14: Tensor/ODF/FOD data properties panel

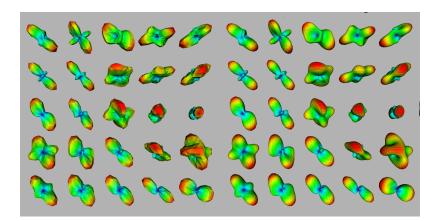


Fig. 15: A comparison of low/high resolution ODF rendering

1.1.5 Data Format

Input

Since we apply the dcm2nii to unify the format of input data, it can support most types of DICOM files (in folders). Please refer to the author's webpage for more information [4]. Please also get back to us if you encounter any problems.

Output

All the intermediate files are in compressed NIFTI format (.nii.gz). The final track file is in .trk format. The .trk format is from the TrackVis [17].

Name	Data type	Bytes	Comment
id_string[6]	char	6	ID string for track file, "TRACK"
dim[3]	short	6	Dimension of the image volume.
voxel_size[3]	float	12	Voxel size of the image volume.
origin[3]	float	12	Origin of the image volume.
n_scalars	short	2	Number of scalars saved at each track point
s_name[10][20]	char	200	Name of each scalar.
n_properties	short	2	Number of properties saved at each track.
p_name[10][20]	char	200	Name of each property.
vox_to_ras[4][4]	float	64	4x4 matrix for voxel to RAS
reserved[444]	char	444	Reserved space for future version.
voxel_order[4]	char	4	Order of the original image data.
pad2[4]	char	4	Paddings.
orient_p[6]	float	24	Image orientation of the original image.
pad1[2]	char	2	Paddings.
invert_x	uchar	1	Inversion/rotation flags
invert_y	uchar	1	As above.
invert_x	uchar	1	As above.
swap_xy	uchar	1	As above.
swap_yz	uchar	1	As above.
swap_zx	uchar	1	As above.
n_count	int	4	Number of tracks stored in this track file.
version	int	4	Version number. Current version is 2.
hdr_size	int	4	Size of the header, should be 1000.

Briefly, header section contains following information:

with data section in following format:

Track	Data type	Bytes	Comment
Track #1	int	4	Number of points in this track, as m.
	float	(3+n_s)*4	Track Point #1.
	float	(3+n_s)*4	Track Point #2. Same as above.
	float	(3+n_s)*4	Track Point #m. Same as above.
	float	n_p*4	n_p float numbers
Track #2	Same as above.		
Track #n	Same as above.		

By default, we prefined two scalars (FA,MD) and three properties (length,FA,MD) in the generated .trk file, along with

six additional values stored in reserved section. These are *version_num*, *num_fibers*, *mean_length*, *total_volume*, *tractFA*, *tractMD* values in float type (single precision).

Again, please visit DataFormat section on TrackVis.org for more detail.

1.1.6 Future Improvements

TO-DO list

- 1. To smooth the ODF/FOD/EAP for a smoother tract;
- 2. To add more efficient tracking algorithms;
- 3. To optimize the 3D rendering function;

1.1.7 Reference

1.2 Tutorials

1.2.1 Programming in Bash/Python

Basically the DiffusionKit is a well self-contained package which implements most of the required modules for diffusion MRI processing and analysis. Additionally, if you want to use script for batch processing for a cohort of subjects, we recommend using Python or Bash. Both of them are inherited for Linux system, and for MS/Windows one can use Anaconda and Git-bash, separately.

The Python is easy to learn for basic use as a script language, although its powerful functions largely depend on 3rd party packages. To this end, we have several suggestions to begin with it. First, take a couple of hours to go through the primary Python grammars. There is a large bundle of free but kind tutorials from internet (Google "Python tutorial" or related keywords). You can choose the websites according to your preference. If you are a newbie of Python, don't think about which version is appropriate for you and just use the latest version (Python>3.0) (In the current stage, you only need to note the difference of "print" function in different versions, which I think is not a smart change from version 3.0); and don't waste money to buy a book since the materials from the internet are largely beyond your capacity. Several tutorial links are listed here:

For English users

- 1. https://en.wikibooks.org/wiki/A_Beginner%27s_Python_Tutorial, a short tutorial
- 2. http://askpython.com/, a short tutorial
- 3. http://www.learnpython.org/, an interactive sandbox

For Chinese users

- 1. http://www.runoob.com/python3/python3-tutorial.html, a nice tutorial
- 2. http://www.cnblogs.com/vamei/archive/2012/09/13/2682778.html, python and advanced
- 3. http://woodpecker.org.cn/abyteofpython_cn/chinese/, a complete reference

If you prefer to use shell script, like Bash, you can also find some entrances for tutorials. The shell script itself is easy to follow and it is a powerful tool to concatenate the underlying execution functions. It should be noted that the Bash script is only for *nix system, so it is not suitable form cross-platforms. Several tutorial links are listed here:

For English users

1. http://linuxconfig.org/bash-scripting-tutorial, a short introduction

- 2. http://www.tldp.org/LDP/abs/html/, a complete tutorial
- 3. http://www.learnshell.org/, an interactive sandbox

For Chinese users

- 1. http://blog.jobbole.com/85183/, a short tutorial
- 2. https://serholiu.com/bash-by-example, several simple examples
- 3. http://c.biancheng.net/cpp/view/6998.html, a complete tutorial

1.2.2 Getting Started

For your convenience, we've created two bash scripts the Advanced and the Primary, which enable processing multiple data within a loop. The instructions for using the Advanced bash script are provided in Download Page.

In this tutorial, however, we would go through the processing pipeline with the Primary bash script, which contains more features and options. It should be noted that both the Advanced and the Primary scripts can be run in simple Bash environment or emulation, since they utilize extremely basic Bash commands. So, even in MS Windows, you can run these two scripts by a Bash emulation, such as git for Windows (STRONGLY recommend), which provides a light but awesome Bash emulation. When you install this emulation, the Primary script can ben run directly once you prepared all the required data (DO remember to add the DiffusionKit installation path to your system search path). And the Advanced script need an additional Makefile-based environment.

Preprocessing

Starting from this section, we would go throught the pipeline provided in DiffusionKit, assuming DiffusionKit is properly installed following Installation Instructions, along with the Example Dataset downloaded on your hard disk.

Data Format Conversion

Analyze format and Nifti format are the data formats that supported by DiffusionKit. If your image data is in DICOM format, you need to convert the data into Nifti format.

To convert DICOM format to Nifti format, please check the checkbox "DICOM to NIfTI" in the main window and input the data directory that contains DICOM files.

GUI-Frontend Usage

Preprocessing	Reconstruction		Tractography	
JICOM to N	IIFTI			
Input Data Di	r /s	ub01/DT	1	
Output Data File:		ub01/DT	I.nii.gz	
Confirma	ation			

Fig. 16: Figure 1. Convert data format.

Command-Line Usage:

\$ dcm2nii -o /path/sub01 -f DTI /path/sub01/DTI

Eddy Current Correction

DiffusionKit has motion correction function which is implemented with a affine registration method. To correct the effect of head motion, please check the checkbox "Eddy Current Correction" and import the original diffusion-weighted imaging data.

GUI-Frontend Usage

Eddy Current Cor	rection
Input Data File:	/sub01/DTI.nii.gz
Output Data File:	/sub01/DTI_eddy.nii.gz
Confirmation	Reference 0

Fig. 17: Figure 2. Eddy current correction.

Command-Line Usage:

\$ bneddy -i /path/sub01/DTI.nii.gz -o /path/sub01/DTI_eddy.nii.gz -ref 0

Skull Stripping

The next step is removal of extra-meningeal tissues from the MRI image of the whole head. The function of Skull Stripping is to delete non-brain tissues from an image of the whole head. An accurate brain mask will accelerate the following reconstruction.

To achieve the mask of brain, please check the checkbox "Skull Stripping" and input the image (e.g. the b0 image) you want to operate.

GUI-Frontend Usage:

Skull Stripping	
Input File:	/sub01/DTI_eddy.nii.gz
Brain File:	/sub01/DTI_eddy_brain.nii.gz
Mask File:	/sub01/DTI_eddy_mask.nii.gz
Confirmation	Threshold 0.50

Fig. 18: Figure 3. Skull stripping.

Command-Line Usage:

Diffusion Model Estimation

DTI Estimation

When we complete the preprocessing, we can do the job of DTI estimation. In this step, you can get diffusion tensor image and derived diffusion indexes (e.g. fractional anisotropy, mean diffusivity, radial anisotropy etc.) from the reconstruction. For GUI-frontend, please change to the "Reconstruction" label and make sure the "Output Type" is DTI. Input the files into the boxes in the "Reconstruction" Panel according to the illustrations in Figure 4.

GUI-Frontend Usage:

Preprocessing	Reconstruction Tractography		
Output Type:	DTI 🍨 👿 Eigen Data		
Data Input			
DWI /su	b01/DTI_eddy.nii.gz		
Bvals /su	/sub01/DTI.bval		
Bvecs /su	/sub01/DTI.bvec		
Mask /su	/sub01/DTI_eddy_mask.nii.gz		
Data Output Prefix /su	b01/DTI_eddy_dti		
Reset	Run		

Fig. 19: Figure 4. DTI estimation.

Command-Line Usage:

```
$ bndti_estimate -d /path/sub01/DTI_eddy.nii.gz -g /path/sub01/DTI.bvec -b /path/

$ sub01/DTI.bval -m /path/sub01/DTI_eddy_mask.nii.gz -o /path/sub01/DTI_eddy_dti -

$ $ tensor 1 -eig 1
```

HARDI Estimation (Optional)

DiffusionKit also includes advanced HARDI reconstructions which implemented with two types of HARDI methods, the Spherical Polar Fourier Imaging (SPFI) method and Constrained Spherical Deconvolution (CSD) methods. If you have DWI data with considerable number of gradient directions (more than 45 directions) and high b-value (larger than 2000) [ref], you can apply HARDI estimation to resolve crossing fibers in the reconstruction. To call the HARDI reconstruction, change the "Output Type" to HARDI and select the HARDI method (SPFI or CSD) you want in the "Reconstruct Parameters" section. The compulsory parameters for HARDI estimation is data input and output. The example of parameters setting is illustrated in Figure 5&6.

GUI-Frontend Usage:

Command-Line Usage:

Preprocessing Reconstruction Tractography
Output Type: HARDI
Data Input
DWI /sub01/DTI_eddy.nii.gz
Bvals /sub01/DTI.bval
Bvecs /sub01/DTI.bvec
Mask /sub01/DTI_eddy_mask.nii.gz
Data Output
Prefix /sub01/DTI_eddy_spfi
Reconstrut Parameters Method SPFI (ODF) Generate Direction
SH Order 4 🗘 SH Lambda 0.00 🗘
Rad Order 1 Rad Lambda 0.00
Radius 0.015 📮 🗌 Generate GFA
Reset

Fig. 20: Figure 5. HARDI estimation using SPFI method.

Preprocessing	Reconstruction	Tractograp	hy
Output Type:	HARDI 🌲		
Data Input			
DWI /sub	01/DTI_eddy.nii.g	Z	
Bvals /sub	01/DTI.bval		
Bvecs /sub	01/DTI.bvec		
Mask /sub	01/DTI_eddy_ma	sk.nii.gz	
Data Output Prefix /sut Reconstrut Par Method CSE		Generate Dire	ection
SH Order Max Iteration Regularization Amptitude	50 ‡	FA Range Resolution	0.75 ÷ 0.95 ÷ 300 ÷
Reset]		Run

Fig. 21: Figure 6. HARDI estimation using CSD method.

```
$ bnhardi_ODF_estimate -d /path/sub01/DTI_eddy.nii.gz -g /path/sub01/DTI.bvec -b /

→path/sub01/DTI.bval -m /path/sub01/DTI_eddy_mask.nii.gz -o /path/sub01/DTI_eddy_spfi

$ bnhardi_FOD_estimate -d /path/sub01/DTI_eddy.nii.gz -g /path/sub01/DTI.bvec -b /

→path/sub01/DTI.bval -m /path/sub01/DTI_eddy_mask.nii.gz -o /path/sub01/DTI_eddy_fod
```

Tractography

DiffusionKit provides the function of fibertracking which is implemented with a deterministic streamline method. You can perform tractography using the diffusion tensors reconstructed with DTI or diffusion/fiber ODFs reconstructed with SPFI/CSD. To call the function of fibertracking, change the "Processing" Panel to "Tractography". Select the data type (DTI or HARDI) according to the data you used (tensor or diffusion/fiber ODF). Input the FA map acquired in DTI estimation to the box of "FA". Provide the seeds image based on your research. For instance, if you want to do whole brain fibertracking, you can input a whole brain mask in the box of "Seeds". You can adjust other tracking parameters to satisfy your study. Please refer to the illustration in Figure 7&8 for more details.

GUI-Frontend Usage

Preprocessing	Reconstruction Tractography
Data Type	DTI 🛟
Data Input	
Data /sul	001/DTI_eddy_dti.nii.gz
Mask /su	001/DTI_eddy_mask.nii.gz
FA /sut	001/DTI_eddy_dti_FA.nii.gz
Seeds /sul	001/DTI_eddy_mask.nii.gz
Data Output	
Fiber /sub	01/DTI_wb.trk
Tracking Parar	neters
Angle thresho	A Threshold 0.10
Step size (vox	el) 0.5 🗘
Min length (m	m) 10 🗘 Max length 5000 🗘
Invert	disabled 🛟 Swap disabled 🛟
Reset	Run

Fig. 22: Figure 7. Fibe tracking based on DTI.

Command-Line Usage

```
$ bndti_tracking -d /path/sub01/DTI_eddy_dti.nii.gz -m /path/sub01/DTI_eddy_mask.nii.

$ path/sub01/DTI_eddy_mask.nii.gz -fa /path/sub01/DTI_eddy_dti_FA.nii.gz -o /

$ path/sub01/DTI_wb.trk

$ bnhardi_tracking -d /path/sub01/DTI_eddy_dti.nii.gz -m /path/sub01/DTI_eddy_mask.

$ inii.gz -s /path/sub01/DTI_eddy_mask.nii.gz -fa /path/sub01/DTI_eddy_dti_FA.nii.gz -

$ o /path/sub01/DTI_wb.trk

$ (continues on next page)
```

Preprocessing	Reconstruction Tractography		
Data Type	HARDI 🔹		
Data Input			
Data /sul	001/DTI_eddy_spfi_EAP_profile.nii.gz		
Mask /sul	001/DTI_eddy_mask.nii.gz		
FA /sut	001/DTI_eddy_dti_FA.nii.gz		
Seeds /sul	Seeds /sub01/DTI_eddy_mask.nii.gz		
Data Output			
Fiber /sub	01/HARDI_wb.trk		
Tracking Parar	neters		
Angle thresho	old 45 🗘 FA Threshold 0.10 🗘		
Step size (vox	el) 0.5 ‡		
Min length (m	m) 10 🗘 Max length 5000 🗘		
Invert	disabled 🔹 Swap disabled 🔹		
Reset	Run		

Fig. 23: Figure 8. Fiber tracking based on HARDI.

Construction of brain networks

When we get connectivity data from tractography, we can consider construction of brain networks. A network is a collection of nodes and links (edges) between pairs of nodes. In structural brain networks, we select specific ROIs of brain as nodes and the number of fibers which end in the pair of ROIs or other derived diffusion indices as edges. DiffusionKit provides the function of bn_network to construct brain network using given ROIs and whole brain tractography achieved from last step. Only command-line usage is available for bn_network. A text file which contains the paths and ROI filenames is needed to input when performing bn_network. Please refer to the example script and example data for details.

Command-Line Usage

1.2.3 Generate a ROI file from a given ROI list

If you want to generate a roi.nii.gz file in the space of a reference ref.nii.gz, with a given list of ROI coordinates and according weight value, where the radii are unified by the input radius r, you can call the bncalc function in-loop as the file gen_ROI_from_list.sh. The content is as following:

```
#!/bin/bash
# This is to call the bncalc function to create a .nii.gz mask image
# from a list of ROI centers and radii
# Jan 7, 2016, by NMZUO
# The list.txt has the format as:
                  weightValue # this could be used for color; set 1 by default
#
  X
        V
              Z
#
  -2
        38
              36
                  3.6
#
  65
        -31
              -9 -2.7
  12
        36
              20 4
#
#
  . . . . . .
Usage() {
   echo ""
   echo "This is to call the bncalc function to create"
   echo "a .nii.gz mask image from a list of ROI centers."
   echo "Usage:"
   echo "bash gen_ROI_from_list.sh list.txt r ref.nii.gz outmask.nii.gz"
   echo -e "
                 list.txt\t\t The input list containing ROI coordinates (MNI mm)"
   echo -e "
                  r\t\t\t The radius (mm) of each ROI"
   echo -e "
                 ref.nii.gz\t The reference image where the ROIs stay"
   echo -e "
                  outmask.nii.gz\t The output .nii.gz image"
}
if [ $# -lt 3 ]; then
   Usage
    exit 1
fi
tmpfile=$$"_tmp_"
```

```
iCount=0
while read line; do
   roi=($line)
    idx=`printf "%06d" $iCount`
   bncalc -i $3 -roi ${roi[0]},${roi[1]},${roi[2]},$2 -o ${tmpfile}${idx}.nii.gz
   if [ ${#roi[@]} -gt 3 ]; then
       bncalc -i ${tmpfile}${idx}.nii.gz -mul ${roi[3]} -o ${tmpfile}${idx}.nii.gz
   fi
    iCount=$((iCount+1))
done < $1
cp ${tmpfile}"000000".nii.gz $4
for i in `seq 1 $(($iCount-1))`
do
   echo -e "$i \c"
   idx=`printf "%06d" $i`
   if [ ${#roi[@]} -lt 4 ]; then
       bncalc -i ${tmpfile}${idx}.nii.gz -mul $(($i+1)) -o ${tmpfile}${idx}.nii.gz
    fi
   bncalc -i $4 -add ${tmpfile}${idx}.nii.gz -o $4
done
echo "Done!"
rm -f ${tmpfile}?????.nii.gz
```

1.2.4 Map ROIs defined in one space to another individual space

This is an example to map the ROI definitions (in the initial space, e.g. MNI 2mm space) to an individual space. Fox example, the source folder could be:

where the fold for each subject could be:

dti_L3.nii.gz
— dti_MD.nii.gz
— dti.nii.gz
— dti_RA.nii.gz
dti_V1.nii.gz
dti_V2.nii.gz
dti_V3.nii.gz
L t1
DP_0001_t1_brain_mask.nii.gz
DP_0001_t1_brain.nii.gz
DP_0001_t1.nii.gz

and *list_dp.txt* is the name list of the subjects as (the column other than the first one is the possible comment for the subject):

```
nmzuo@:dp_classify$ cat list_dp.txt
DP_0001
DP_0002 #no cerebellum
.....
```

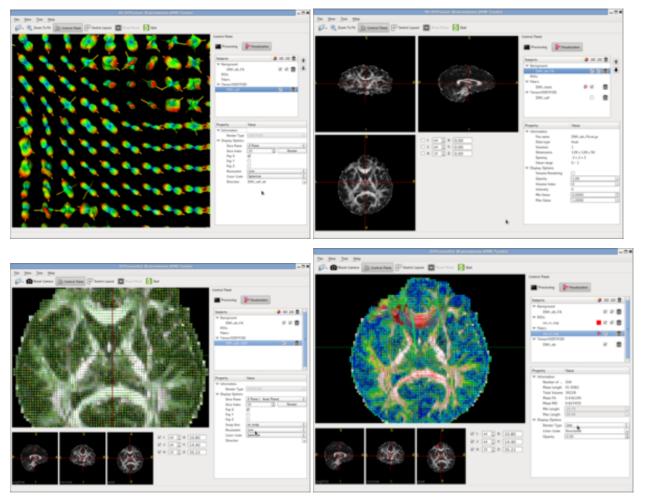
Then the following *bash code* is able to map the ROI in the standard space to the subject space (for each subject in *list_dp.txt*). Certainly the initial ROI does not require to be in the standard space in this code.

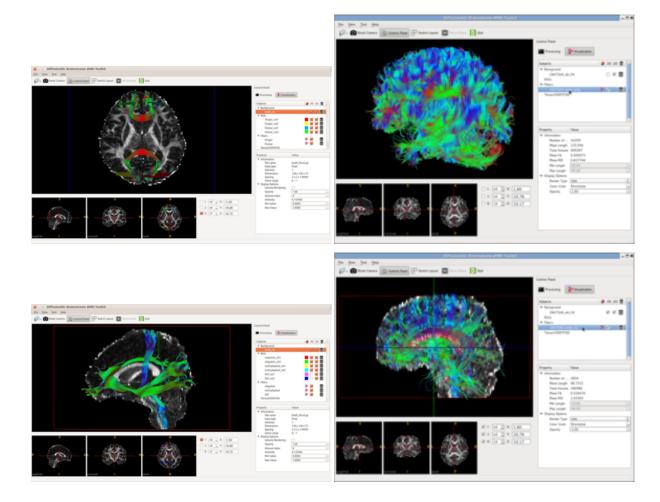
```
atlas='/datc/software/Brainnetome_Atlas/BN_Atlas_246_2mm.nii.gz' # Brainnetome Atlas.
→in MNI space
mni='/datc/software/fsl5.0/data/standard/MNI152_T1_2mm_brain.nii.gz'
for dat in `cat list_dp.txt |awk '{print $1}' ` #only read the first column
do
   echo $dat
   cpath='/datd/dp_classify'
   cpath=$cpath/${dat%%_*}/$dat/
   oldp=`pwd`
   cd $cpath
   echo 'aladin'
   reg_aladin -ref t1/$dat'_t1_brain.nii.gz' -flo $mni -aff aff.txt -voff
   echo 'f3d'
   reg_f3d -ref t1/$dat'_t1_brain.nii.gz' -flo $mni -aff aff.txt -cpp cpp.nii.gz,
→-maxit 3 -ln 2 -voff
   rm -f aff.txt
   echo 'aladin'
   reg_aladin -ref dwi/$dat'_dwi_eddy_b0.nii.gz' -flo t1/$dat'_t1_brain.nii.gz'
→aff aff0.txt -voff
   reg_resample -ref t1/$dat'_t1_brain.nii.gz' -flo $atlas -trans cpp.nii.gz -
→inter 0 -res jhu_t1.nii.gz -voff
   reg_resample -ref dwi/$dat'_dwi_eddy_b0.nii.gz' -flo jhu_t1.nii.gz -trans aff0.
→txt -inter 0 -res jhu_b0.nii.gz -voff
   rm -f aff0.txt cpp.nii.gz outputResult.nii jhu_t1.nii.gz
   mv jhu_b0.nii.gz dwi/atlas_b0.nii.gz
   cd $oldp
done
```

1.3 Screenshots

1.3.1 Tensor and ODF/FOD Visualization

[click to see high resolution images]





1.3.2 Tractography Visualization

1.4 Download and Install

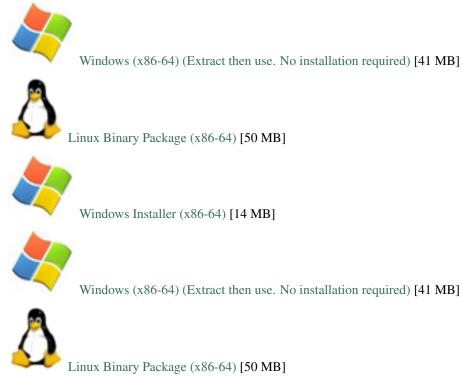
1.4.1 Download Links

Latest Release (Sept. 28, 2018)

Previous Release

See Releases for our previously compiled packages.





Example Data and Test Script

- List File: list.txt [12 Bytes]
- Subject 01: sub01.tar.gz [87.1 MB]
- Subject 02: sub02.tar.gz [85.5 MB]
- Brain Atlas: atlas.tar.gz [1.57 MB]
- Test Script [process_advanced.sh] [process_primary.sh]
- List File: list.txt [12 Bytes]
- Subject 01: sub01.tar.gz [87.1 MB]
- Subject 02: sub02.tar.gz [85.5 MB]
- Brain Atlas: atlas.tar.gz [1.57 MB]
- Test Script [process_advanced.sh] [process_primary.sh]

And this is what we have in the data above:

```
atlas
aal.nii.gz
aal.nii.lut
aal.nii.txt
aal_roi_024.txt
aal_roi_090.txt
BN_Atlas_246_1mm.nii.gz
BN_Atlas_246.txt
ch2bet.nii.gz
atlas.tar.gz
```

list.txt
process_advanced.sh
<pre>process_primary.sh</pre>
sub01
dwi.bval
dwi.bvec
dwi.nii.gz
Lt1.nii.gz
sub01.tar.gz
sub02
dwi.bval
dwi.bvec
— dwi.nii.gz
t1.nii.gz
sub02.tar.gz

We recommand users to follow the Tutorial Page for a step-by-step introduction of the functions within DiffutionKit.

IN A HURRY? Download all the data files (REQUIRED) above and run the Bash script process_advanced.sh or process_primary.sh. These two scripts have exactly the same functions, except that the primary one is for those who don't have too much background in Bash scripting and the advanced one is Makefile-based which avoids repeatly compiling the unchaged data in multiple compiles. Or simply run the following commands to do everything (for Linux ONLY).

```
# install DiffusionKit
kitv='v1.4-r161127' ## This line should be changed according to the version you.
→ desired.
kitname='DiffusionKitSetup-x86 64-'${kitv}
wget https://github.com/brainnetome/diffusionkit/releases/download/${kitv}/${kitname}
↔'.tar.gz'
tar zxvf ${kitname}'.tar.gz'
export PATH=$PATH:$(pwd) / ${kitname}/bin
# get the data and run!
wget https://github.com/brainnetome/diffusionkit/releases/download/v1.4-r161127/list.
→t.xt.
wget https://github.com/brainnetome/diffusionkit/releases/download/v1.4-r161127/sub01.
⇔tar.gz
wget https://github.com/brainnetome/diffusionkit/releases/download/v1.4-r161127/sub02.
→tar.gz
wget https://github.com/brainnetome/diffusionkit/releases/download/v1.4-r161127/atlas.
\rightarrowtar.gz
wget https://raw.githubusercontent.com/brainnetome/diffusionkit/master/source/static/
\hookrightarrow process_advanced.sh
chmod +x process_advanced.sh
./process_advanced.sh
```

```
# install DiffusionKit
kitv='v1.3-r160923' ## CAUTION: This line should be changed according to the version.
kitname='DiffusionKitSetup-x86_64-'${kitv}
wget http://ddl.escience.cn/ff/emKy?func=download -0 ${kitname}'.tar.gz'
tar zxvf ${kitname}'.tar.gz'
export PATH=$PATH:$(pwd)/${kitname}/bin
# get the data and run!
wget https://github.com/brainnetome/diffusionkit/releases/download/v1.3-r160923/list.

txt
```

And more simpler way is to copy the code in the above panel (choose the server close to you) and save it as a file, such as test-diffusionkit.sh, or test-diffusionkit-china.sh. and run to take a quick look how DiffusionKit can **BUILD A BRAIN NETWORK JUST A FEW MINUTES**!

1.4.2 System requirement

Basically this software can run in any system, including 64-bit MS Windows/Linux OS, although currently we only tested and released the binary packages for Windows/Linux OS. The software is developed based on C/C++, and some platform independent packages, including VTK, and OpenCV, etc. However, for high-performance data processing and visualization, we recommend using 64-bit OS with multi-core CPU and standalone graphics card.

1.4.3 Install/Uninstall

Please download the package from http://diffusion.brainnetome.org, according to your own OS. Unpack the files to where you want and you can enjoy the software. The 64-bit OS is recommended for high-performance data processing. Each installation package is completely standalone so you DO NOT need to install ANY other dependency to run the software. If you encounter any dependency problem please DO contact us.

For MS Windows OS

Double click the DiffusionKitSetup-WIN64-v1.4-r161127.exe file and then choose the destination path according to the wizard. You may need to provide administrator permission if you want to put the files into the system path. Similarly, to uninstall you only need to hit the menu of "uninstall" in the MS Windows start menu.

For Linux OS

Glibc>=2.2 is required. Download the DiffusionKitSetup-x86_64-v1.4-r161127.tar.gz, and then

```
tar zxvf DiffusionKitSetup-x86_64-v1.4-r161127.tar.gz
export PATH=$PATH:`pwd`/DiffusionKitSetup-x86_64-v1.4-r161127/bin
```

You could add the path to the \$PATH in the ~/.bashrc file, by adding the following line,

export \$PATH=\$PATH:/your/path/to/diffusionkit

To uninstall the software, just manually remove the entire folder where you untar-ed the .tar.gz file.

The citation for DiffusionKit could be:

Sangma Xie, Liangfu Chen, Nianming Zuo and Tianzi Jiang, **DiffusionKit: A Light One-Stop Solution** for Diffusion MRI Data Analysis, *Journal of Neuroscience Methods*, vol. 273, pp. 107-119, 2016. [PDF].

1.5 License

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1.5.1 The licenses of the third party toolboxes

BET (Brain Extraction Tool) v2.1 - FMRIB Analysis Group, Oxford

FMRIB Software Library, Release 5.0 (c) 2012, The University of Oxford (the 'Software')

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dcm2nii - Chris Rorden's dcm2niix, version 24Nov2014

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1.6 FAQs

1.6.1 1. Where can I download example data?

We provide a group of test data to test the software, and the data was also used in [11].

1.6.2 2. Why images and fibers are not properly aligned in my case?

This typically happens when nifti image headers are modified before generating fiber.

In DiffusionKit, a command named bninfo is provided to print out the header information inside a NIfTI image. To check whether an NIfTI data is canonical:

```
$ bninfo data.nii.gz
```

you would get:

```
qform_i_orientation = 'Left-to-Right'
qform_j_orientation = 'Posterior-to-Anterior'
qform_k_orientation = 'Inferior-to-Superior'
...
```

This indicate that you are lucky. And the matrix that transforms voxels coordinates (I/J/K) to physical locations (X/Y/Z) is orthogonal. However, it doesn't matter if this isn't true, the image can be converted with bnconvert. With --reorient argument, it transforms the input image into orthogonal one with head information modified as well.

\$ bnconvert data_brain.nii.gz data_brain_r.nii.gz -r

Warning: However, there would be problems if bnconvert converts a 4D vector image, e.g. tensor data, ODF/FOD data etc.

Usually, the reorient step should be taken when DICOM image is converted to NIFTI image, with argument -r y.

For more information, we recommand reading FSL's Orientation Explained and Docs on qform and sform to get a good understanding of orientation layout within NIfTI data format.

1.6.3 3. How to extract the gradient table if the dcm2nii/MRIcron fails?

The dcm2nii/MRIcron are useful to extract the gradient table for the DWI data. Occasionally, they fails to get the gradient table, though it indeed get the NIFTI data from the DICOM images. Herein we provide a temporary solution by calling the MATLAB/dicominfo which depends on a specified keyword dictionary.

```
function findGrad(dcmfile)
% This script is special to locate the individual grad directions
% for each DICOM file, if the dicom2nii tool failed to extract them.
% by NMZUO, Sept. 12, 2014
% Change the following if still fails
%prestr = 'DiffusionGradientOrientation';
prestr = 'DiffusionGradient';
myhdr = dicominfo(dcmfile);
myfind = fieldnamesr(myhdr, prestr);
iLen = length(myfind);
iCount = 0;
for i=1:iLen
    mystr = myfind{i};
```

```
if strfind(mystr, prestr)
    iCount = iCount + 1;
    disp(['Field containing ' prestr ': ' num2str(iCount) ]);
    disp(mystr);
    % to extract the grad directions
    eval(strcat('myhdr.', mystr))
  end
end
end
```

Please download the two files fieldnamesr.m and findGrad.m. The first one is to generate the full string of the keyword dictionary and second is to find the locations. If your problem remains unsolved, please send us your data (only one 2D/3D DICOM image is enough) and then we will update our DICOM dictionary.

1.7 Support

1.7.1 Feedback

If you encounter any problems or have any suggestions, please feel free to get back to us, diffusion.kit@nlpr.ia.ac.cn .

Note: Document last updated on Apr 27, 2020.