
Simulated Diffusion-Weighted Imaging for the ITK Masses

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Abstract

A recent article by Van Hecke et al. [3] describes a framework for creating simulated diffusion-weighted images (DWI). The methodology allows for modeling intersubject variability, regional pathology, and noise and is quite useful for evaluation purposes. This article describes our ITK implementation of this work which we provide to the community as open source.

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Contents

1	Introduction	1
2	General Overview of Simulated DWI Creation	2
3	Implementation	2
4	Simulated DWI Example	4

1 Introduction

The popularity of diffusion tensor imaging (DTI) has resulted in a proliferation of corresponding analysis algorithms. The need for an evaluative framework for such algorithms motivated the developments of Van Hecke and colleagues in a recent paper [3] which encompasses modeling intersubject variability, regional pathology, and noise in creating tailored DWI from an existing DTI cohort. It should be noted that whereas

Van Hecke's paper also describes the registration processes to both align the DTI cohort to the common reference space and back to the individual subject space, we omit that part of the framework in this submission and allow the user to select their preferred method of image normalization. In addition, the user will need to reconstruct the DTI which can easily be performed with several publicly available packages such as Camino [2].¹

2 General Overview of Simulated DWI Creation

The DWI simulation allows for the creation of a control group and experimental group consisting of an arbitrary number of directional image sets which can then be fitted to create individual DTI.

Intersubject variability is modeled using an eigendecomposition of the aligned sample DTI cohort. Specifically, for each aligned DTI, the first (i.e. longitudinal) eigenvalue and the average of the second and third (i.e. transverse) eigenvalues of each voxel within a specified brain mask are concatenated within a single vector. The matrix constituting the grouping of image vectors of eigenvalues is then decomposed to create the projections describing the variation found within the aligned cohort. These projections are used to add random intersubject variations to each of the control and experimental subject images.

Based on recent literature, pathology is introduced by Van Hecke using a simultaneous decrease in the longitudinal eigenvalues and an increase in the transverse eigenvalues. In propagating the change in transverse diffusion, we enforce that the ratio of the second to the third eigenvalues be the same before and after a transverse diffusion decrease.

Finally, the application of standard Rician noise is also included as a possibility for enhancing the realism of the simulated DWI data.

For further details, we encourage the reader to consult the paper on which this work is based.

3 Implementation

This work was originally developed within our Advanced Normalization Tools software package² and uses our ANTs command line classes. After compilation, one can invoke the help menu which will list the set of options and corresponding descriptions:

```
[ntustison@Nietzschean-Numerics] $ CreateDTICohort --help
COMMAND:
  CreateDTICohort
    CreateDTICohort implements the work of Van Hecke et al. (On the construction of
    a ground truth framework for evaluating voxel-based diffusion tensor MRI analysis
    methods, Neuroimage 46:692-707, 2009) to create simulated DTI data sets. The
    only difference is that all registrations (both for the input population and for
    the output population) are assumed to take place outside of this program.

OPTIONS:
  -d, --image-dimensionality 2/3
    This option forces the image to be treated as a specified-dimensional image. If
    not specified, the program tries to infer the dimensionality from the input
    image.

  -a, --dti-atlas inputDTIAtlasFileName
```

¹<http://camino.org.uk/>

²<http://www.picsl.upenn.edu/ANTS/>

A diffusion tensor atlas image is required input for creating the cohort.

```
-x, --label-mask-image maskImageFileName
                        lowerThresholdValue
```

A mask image can be specified which determines the region(s). to which the simulated pathology operations are applied. See also the option '--pathology'. If no mask is specified one is created by thresholding the atlas FA map at 0.2 unless a lower threshold is specified.

```
-n, --noise-sigma <noiseSigma=18>
```

This parameter characterizes the Rician noise in the original DWIimages. Van Hecke uses the noise-estimation method of Sijbers et al. "Automatic estimation of the noise variance from the histogram of a magnetic resonance image", Phys. Med. Biol. 52:1335-1348, 2007.

```
-p, --pathology label[<percentageChangeEig1=-0.05>,<percentageChangeAvgEig2andEig3=0.05>,<numberOfVoxels=all or percentageOfvoxels>]
```

The user can specify the simulated pathology in a given area using a label mask. If no label is prepended to parameters, the specified parameters are applied to all labels. Pathology is simulated by changing the eigenvalues. Typically this involves a decrease in the largest eigenvalue and an increase in the average of the remaining eigenvalues. Change is specified as a percentage of the current eigenvalues. However, care is taken to ensure that diffusion direction does not change. Additionally, one can specify the number of voxels affected in each region or one can specify the percentage of voxels affected. Default is to change all voxels. Note that the percentages must be specified in the range [0,1]. For dimension=3 where the average transverse diffusion eigenvalues are altered, this change is propagated to the distinct eigenvalues by forcing the ratio to be the same before the change.

```
-w, --dwi-parameters [B0Image,directionFile,bvalue]
                        [B0Image,schemeFile]
```

This option specifies the parameters of the output diffusion-weighted images including the directions and b-values. The directions are specified using a direction file which has as its first line the number of directions. Each successive three lines contains the x, y, and z directions, respectively, and a single b-value. Note that several direction files of this format are distributed with the Camino DTI toolkit (<http://web4.cs.ucl.ac.uk/research/medic/camino/pmwiki/pmwiki.php>). Alternatively, one can specify a scheme file where each direction is specified followed by a b-value for that direction, i.e. <x1> <y1> <z1> <bvalue1> ... <xN><yN><zN><bvalueN>.

```
-r, --registered-population textFileWithFileNames.txt
```

If one wants to introduce inter-subject variability a registered DTI population to the DTI atlas is required. This variability is modeled by a PCA decomposition on a combination of the first eigenvalue image and the average of the second and third eigenvalues. The registered image file names are specified using a text file where each line is the name of an individual DTI.

```
-o, --output [outputDirectory,fileNameSeriesRootName,
              <numberOfControls=10>,<numberOfExperimentals=10>]
```

The output consists of a set of diffusion-weighted images for each subject. Each file name is prepended with the word 'Control' or 'Experimental'. The number of control and experimental subjects can be also be specified on the command line. Default is 10 for each group.

```
-h
```

Print the help menu (short version).
<VALUES>: 0

```
--help
```

Print the help menu.

Listing 1: CreateDTICohort long command line menu which is invoked using the '--help' option. The short command line menu is obtained by typing '-h'.

4 Simulated DWI Example

We have included a sample data set derived from the NKI/Rockland INDI³ data set consisting of 10 DTI from normal subjects, a B0 image, and a label mask modified from the popular ICBM template consisting of 52 regions where the additional 2 regions are 1) unlabeled regions from the original ICBM mask where $FA \geq 0.2$ and 2) the remainder of the brain.⁴ This allows for pathology to be individually prescribed for each labeled region. All images are aligned to the MNI152 template space [1].

Below we give the first few lines of the terminal output from a sample call to the routine `CreateDTICohort` contained in the script `createDWI.sh`, which is also included.

```
[ntustison@Nietzschean-Numerics] $ sh createDWI.sh

Creating DTI cohort for 3-dimensional images.

--- Modeling intersubject variability ---

Processing DTI/dti_01.nii.gz (1 of 10).
Processing DTI/dti_02.nii.gz (2 of 10).
Processing DTI/dti_03.nii.gz (3 of 10).
Processing DTI/dti_04.nii.gz (4 of 10).
Processing DTI/dti_05.nii.gz (5 of 10).
Processing DTI/dti_06.nii.gz (6 of 10).
Processing DTI/dti_07.nii.gz (7 of 10).
Processing DTI/dti_08.nii.gz (8 of 10).
Processing DTI/dti_09.nii.gz (9 of 10).
Processing DTI/dti_10.nii.gz (10 of 10).

--- Calculating regional average FA and MD values (original and pathology) ---
```

Region	FA (original)	FA (path.)	FA (%change)	MD (original)	MD (path.)	MD (%change)
1	0.426666	0.392867	-0.0792165	0.0010568	0.00101915	-0.0356265
2	0.387891	0.385816	-0.00534951	0.000799096	0.000827288	0.0352805
3	0.610309	0.618394	0.013248	0.00087893	0.000985969	0.121783
4	0.562708	0.572108	0.0167038	0.000992207	0.00111096	0.119684
5	0.641522	0.646897	0.00837787	0.00091601	0.00103919	0.13448
6	0.450288	0.462858	0.0279156	0.00154076	0.00185622	0.204742
7	0.504297	0.562074	0.114571	0.000824662	0.00090193	0.0936977
8	0.455938	0.452463	-0.0076219	0.0008713	0.000884652	0.0153244
9	0.458622	0.447742	-0.0237231	0.000859755	0.000851439	-0.00967258
10	0.553985	0.568811	0.0267624	0.000894992	0.000954038	0.0659738
11	0.560088	0.57298	0.0230185	0.000869192	0.000948747	0.0915286
12	0.455487	0.440347	-0.0332389	0.00151854	0.00143839	-0.0527812
13	0.450941	0.436025	-0.0330789	0.00232649	0.00220849	-0.050718
14	0.531836	0.555623	0.0447257	0.00104972	0.00119604	0.139394
15	0.558183	0.57513	0.0303612	0.000980834	0.00110763	0.129277
16	0.604708	0.619218	0.023996	0.000831829	0.000947156	0.138643
17	0.620054	0.649983	0.0482687	0.000799265	0.00089297	0.117239
18	0.44215	0.455869	0.0310285	0.000782263	0.00085731	0.0959354
19	0.468299	0.495326	0.0577122	0.000743647	0.000816076	0.0973973
20	0.59195	0.63697	0.0760543	0.000770545	0.000860518	0.116765
21	0.603748	0.645064	0.0684337	0.000746336	0.000831687	0.114361
22	0.509597	0.554742	0.0885913	0.000837227	0.000934504	0.11619
23	0.359127	0.397746	0.107537	0.000830458	0.000907076	0.0922604
24	0.37626	0.419576	0.115122	0.000813638	0.000879125	0.080486
25	0.401218	0.440621	0.098208	0.000789715	0.000861153	0.0904596
26	0.392857	0.426172	0.0848036	0.000757251	0.000823564	0.0875718
27	0.367774	0.427039	0.161146	0.000873593	0.000956149	0.0945015
28	0.395989	0.449048	0.133992	0.000865832	0.000947765	0.0946285
29	0.467424	0.483241	0.0338396	0.000879305	0.00097071	0.103951
30	0.459866	0.486898	0.058784	0.000899325	0.000982584	0.0925787

³http://fcon_1000.projects.nitrc.org

⁴http://www.loni.ucla.edu/Atlases/Atlas_Detail.jsp?atlas_id=5

```

31 0.468384 0.520791 0.111889 0.000892105 0.000971572 0.0890779
32 0.475016 0.504796 0.0626932 0.000931563 0.00104899 0.126052
33 0.330981 0.372975 0.12688 0.000834499 0.000914815 0.0962437
34 0.335543 0.372813 0.111073 0.000800139 0.000874732 0.0932242
35 0.474078 0.510758 0.0773704 0.000793743 0.000879568 0.108127
36 0.406761 0.459163 0.128828 0.000771591 0.000858159 0.112193
37 0.305497 0.314823 0.0305267 0.000867374 0.00097527 0.124394
38 0.320879 0.34981 0.0901605 0.000847713 0.00095333 0.124591
39 0.476671 0.483099 0.0134837 0.000928494 0.0010066 0.0841179
40 0.474913 0.495017 0.042332 0.000925391 0.00102012 0.102363
41 0.378974 0.409393 0.0802644 0.000779719 0.000843773 0.0821508
42 0.398861 0.420382 0.0539565 0.000765444 0.000833252 0.0885867
43 0.360756 0.36034 -0.00115312 0.000769285 0.00083562 0.0862293
44 0.367618 0.386133 0.0503644 0.000755733 0.000823069 0.089101
45 0.408336 0.432339 0.0587838 0.000865836 0.000952377 0.09995
46 0.450344 0.473607 0.0516559 0.000839035 0.000934961 0.114329
47 0.352407 0.335938 -0.046734 0.000862954 0.000947891 0.0984256
48 0.479243 0.482496 0.00678764 0.000806836 0.000892731 0.106459
49 0.390706 0.409486 0.0480653 0.00127162 0.00148426 0.167222
50 0.420062 0.456485 0.0867099 0.00140615 0.00156606 0.11372
51 0.296425 0.326624 0.101875 0.000876206 0.000949146 0.0832462
52 nan nan nan 0.00901245 0.00884544 -0.0185311

--- Writing images ---

Writing control 1 (of 11) DWI images.
Applying direction 0 (of 30): [0 0 0], bvalue = 0
Applying direction 1 (of 30): [-0.546405 0.619202 0.563943], bvalue = 1000
Applying direction 2 (of 30): [-0.398931 -0.600006 0.693432], bvalue = 1000
Applying direction 3 (of 30): [0.587973 0.521686 0.618168], bvalue = 1000
Applying direction 4 (of 30): [0.055894 -0.991971 -0.113444], bvalue = 1000
Applying direction 5 (of 30): [-0.666933 -0.677984 0.309094], bvalue = 1000
Applying direction 6 (of 30): [0.163684 0.533013 0.830123], bvalue = 1000
Applying direction 7 (of 30): [0.542826 0.133898 0.829102], bvalue = 1000
Applying direction 8 (of 30): [-0.074751 -0.350412 0.933608], bvalue = 1000
Applying direction 9 (of 30): [0.845751 -0.478624 -0.235847], bvalue = 1000
Applying direction 10 (of 30): [0.767148 -0.610673 0.196372], bvalue = 1000
Applying direction 11 (of 30): [-0.28381 0.381633 0.879663], bvalue = 1000
Applying direction 12 (of 30): [0.537228 -0.616249 0.575868], bvalue = 1000
Applying direction 13 (of 30): [-0.711387 0.197778 0.674398], bvalue = 1000
Applying direction 14 (of 30): [0.886511 0.219025 0.407586], bvalue = 1000
Applying direction 15 (of 30): [0.296061 0.842985 0.449136], bvalue = 1000
Applying direction 16 (of 30): [-0.93754 -0.34099 0.068877], bvalue = 1000
Applying direction 17 (of 30): [0.398833 0.917023 0.000835], bvalue = 1000
Applying direction 18 (of 30): [0.097278 -0.711949 0.69546], bvalue = 1000
Applying direction 19 (of 30): [-0.311534 0.908623 -0.278121], bvalue = 1000
Applying direction 20 (of 30): [-0.432043 -0.089758 0.897375], bvalue = 1000
Applying direction 21 (of 30): [-0.94998 0.03081 0.310788], bvalue = 1000
Applying direction 22 (of 30): [0.146722 -0.811981 -0.564942], bvalue = 1000
Applying direction 23 (of 30): [-0.172201 -0.908573 0.38058], bvalue = 1000
Applying direction 24 (of 30): [0.507209 -0.848578 -0.150513], bvalue = 1000
Applying direction 25 (of 30): [-0.730808 -0.654136 -0.194999], bvalue = 1000
Applying direction 26 (of 30): [0.077744 0.094961 0.992441], bvalue = 1000
Applying direction 27 (of 30): [0.383976 -0.293959 0.8753], bvalue = 1000
Applying direction 28 (of 30): [0.788208 -0.213656 0.57713], bvalue = 1000
Applying direction 29 (of 30): [-0.752333 -0.301447 0.585769], bvalue = 1000
Applying direction 30 (of 30): [-0.975732 0.165497 -0.143382], bvalue = 1000

Writing control 2 (of 11) DWI images.
Applying direction 0 (of 30): [0 0 0], bvalue = 0
Applying direction 1 (of 30): [-0.546405 0.619202 0.563943], bvalue = 1000
Applying direction 2 (of 30): [-0.398931 -0.600006 0.693432], bvalue = 1000
Applying direction 3 (of 30): [0.587973 0.521686 0.618168], bvalue = 1000
. . .
. . .
. . .

```

References

- [1] D. L. Collins A. C. Evans and B. Milner. An MRI-based stereotactic atlas from 250 young normal subjects. In *Society for Neuroscience Abstracts*, volume 18, page 408, 1992. [4](#)
- [2] P. A. Cook, Y. Bai, S. Nedjati-Gilani, K. K. Seunarine, M. G. Hall, G. J. Parker, and D. C. Alexander. Camino: Open-source diffusion-MRI reconstruction and processing. In *Proceedings of the 14th Scientific Meeting of the International Society for Magnetic Resonance in Medicine*, 2006. [1](#)
- [3] Wim Van Hecke, Jan Sijbers, Steve De Backer, Dirk Poot, Paul M Parizel, and Alexander Leemans. On the construction of a ground truth framework for evaluating voxel-based diffusion tensor mri analysis methods. *Neuroimage*, 46(3):692–707, Jul 2009. ([document](#)), [1](#)