



Published in final edited form as:

IEEE J Biomed Health Inform. 2016 November ; 20(6): 1552–1561. doi:10.1109/JBHI.2015.2470682.

Single Anisotropic 3D MR Image Upsampling via Over-complete Dictionary Trained from In-plane High Resolution Slices

Yuanyuan Jia,

College of Computer Science, Chongqing University, Chongqing, China

Zhongshi He*,

College of Computer Science, Chongqing University, Chongqing, China

Ali Gholipour, and

Boston Children's Hospital, Harvard Medical School, 300 Longwood Ave. Boston, MA 02115 USA

Simon K. Warfield

Boston Children's Hospital, Harvard Medical School, 300 Longwood Ave. Boston, MA 02115 USA

Abstract

In Magnetic Resonance (MR), hardware limitation, scanning time, and patient comfort often result in the acquisition of anisotropic 3D MR images. Enhancing image resolution is desired but has been very challenging in medical image processing. Super resolution (SR) reconstruction based on sparse representation and over-complete dictionary has been lately employed to address this problem; however, these methods require extra training sets, which may not be always available. This paper proposes a novel single anisotropic 3D MR image upsampling method via sparse representation and over-complete dictionary that is trained from in-plane high resolution slices to upsample in the out-of-plane dimensions. The proposed method, therefore, does not require extra training sets. Abundant experiments, conducted on simulated and clinical brain MR images, show that the proposed method is more accurate than classical interpolation. When compared to a recent upsampling method based on the non-local means approach, the proposed method did not show improved results at low upsampling factors with simulated images, but generated comparable results with much better computational efficiency in clinical cases. Therefore, the proposed approach can be efficiently implemented and routinely used to upsample MR images in the out-of-planes views for radiologic assessment and post-acquisition processing.

Index Terms

Magnetic resonance imaging; Over-complete dictionary; Super resolution reconstruction; Sparse representation

I. Introduction

Spatial resolution in magnetic resonance imaging (MRI) depends on multiple factors, but is limited by the MRI hardware, tissue relaxation times and image contrast requirements,

*Corresponding author: Zhongshi He.

acquisition time, and patient comfort. In a trade-off to avoid prolonged scans to reduce the risk of subject motion and increase patient comfort, while maintaining high signal-to-noise ratio and contrast-to-noise ratio, many MRI scans are performed with relatively few slices but with rather large slice thickness. The acquired images often have higher in-plane resolution (i.e. in the phase-encoding and frequency-encoding dimensions) than the out-of-plane resolution (or the slice-select direction, also referred to as through-plane dimension), thus have anisotropic voxels (i.e. rectangular voxels with one direction longer than the other two) that are longer in the slice select direction. This results in significant partial voluming effect (PVE) in the out-of-plane views. Such low resolution (LR) images pose limitations on the performance of voxel-wise analysis, image segmentation, and other post-processing algorithms. Standard interpolation methods, such as nearest neighbor, bilinear, bicubic, and B-spline interpolations, may be used to scale up the LR images, but result in blocky edges.

Super-resolution (SR) techniques have emerged as efficient methods to improve the resolution of images. The idea behind SR is to reconstruct a high resolution (HR) image as accurately as possible based on single or multiple low-resolution images. In 2001 and 2002, initial attempts were made to adopt SR algorithms from the computer vision community to medical imaging with a focus on MRI [1]. The MRI framework is particularly well adapted to the application of SR techniques because of the control one has over the acquisition process [2]. The SR methods have shown to improve the trade-off between resolution, signal-to-noise ratio (SNR) and acquisition time of specific MR imaging sequences [3].

SR reconstruction can be performed both in the frequency domain and the spatial domain. The frequency-domain SR methods are simple, but the observation models are limited to global translation motion and linear space invariant blur. Besides, it is difficult to utilize the spatial prior information. On the other hand, in the spatial-domain SR methods, more comprehensive generative models, and spatial prior information may be used to achieve improved reconstruction accuracy. In the following discussion we focus on SR MRI methods in the spatial domain.

Previously in MRI multiple images of the same subject with small shifts were acquired to reconstruct the HR image aiming to improve both the in-plane and out-of-plane resolution. In [4] the authors tried to improve the in-plane resolution. This method was questioned by [5] since the in-plane images obtained by shifting the field-of-view (FOV) involved the acquisition of the same points in k-space (i.e. spatial frequency domain, where information about the frequency of a signal and where it comes from in the patient is stored), which means they contained the same information. In [6] variable demodulation frequency was used to obtain shifted sampling in the image space for in-plane SR. In [7] the authors compared the combination of images acquired at the same sample points in k-space but shifted afterwards, and the combination of the same number of images acquired at shifted positions through changes in demodulation frequency. They found that the HR images obtained by the latter method contained additional details in the form of image features, suggesting that new information was added through a denser sampling of the point-spread function (PSF). This study confirmed that the in-plane resolution in MRI was dependent on the effective width of the PSF and the extent of the k-space sampling.

Super-resolution in the out-of-plane directions of 2D MRI scans has been more promising as 2D MR acquisition is governed by slice-selective excitation in the image domain and Fourier encoding is only performed in the phase and frequency encoding directions. Shifted, rotated, or orthogonal anisotropic slice acquisitions therefore naturally contain additional information that can be used in SR reconstruction. Greenspan *et al.* [1] applied SR method to combine several orthogonal 2D MRI acquisitions to improve the out-of-plane resolution. The results were encouraging. Many methods from the SR reconstruction literature have been adopted to improve the out-of-plane resolution of MRI to generate isotropic (symmetrical) voxels. Typical reconstruction-based algorithms used for SR MRI are error back-projection [8], maximum a posteriori (MAP) estimation [9] and projection into convex sets [10]. Obviously, SR methods that combine multiple LR images for resolution enhancement rely on an exact correspondence between images. This may be achieved using image registration. The reconstruction results are highly dependent upon original alignment of images or the registration accuracy. These techniques have evolved into complex algorithms that correct for motion at the slice level and combine the image information in a robust fashion in an application like fetal MRI [11].

In another, more common scenario, improving resolution through upsampling is desired where no additional scans are available. The single-frame SR reconstruction has emerged to address this problem, with the specific aim of estimating the best possible HR image from only one single LR image. These techniques are naturally compared to interpolation algorithms that are routinely used for image upsampling (e.g. nearest neighbor, bilinear, bicubic, B-spline, and windowed Sinc interpolations). While high-order B-spline and windowed Sinc kernel functions provide good, practical estimations of the ideal Sinc interpolator, all these techniques are bounded by fundamental performance limits as they do not use any prior knowledge about image structure or appearance in upsampling.

A recent trend in SR reconstruction is learning-based methods which exploit the natural redundancy and self-similarity of images. These methods have shown competitive results compared to high order interpolation. The two most successful classes of techniques in this category use Non-local Means (NLM) and Sparse Representations. The NLM method was first proposed in [12] for image denoising. In [13], [14], the NLM approach was adopted to SR reconstruction in MRI. A feature-based multi-modality approach was proposed and generated better results in [15]. Sparse representation, which has shown great promise in processing natural images, has also been successfully applied to single-image SR [16], [17]. In [18], the algorithm from [16], [17] were applied to single anisotropic 3D brain MR image, with a knowledge-driven patch selection criteria based on brain tissue segmentation.

Despite the overall satisfactory performance of the learning-based single-image SR methods, they require extra HR reference images and training sets, and are typically demanding on computational resources. These will significantly reduce the applicability of these techniques in clinical practice. Based on the above analysis, and to mitigate the limitations of the current techniques, this paper proposes a new approach to upsample a single anisotropic 3D MR image without extra training sets based on sparse representation and over-complete dictionary. The proposed method is compared with classical interpolation and a state-of-the-art NLM-based SR approach [13], which also does not require an HR

reference image but does not use the in-plane HR images either. Experiments were performed with both simulation and real clinical 3D brain MR images, showing that the proposed method achieves much better results than classical interpolation methods. In clinical MR images, this approach outperforms the NLM-based method both in terms of accuracy and the execution time and memory usage.

The rest of the paper is organized as follows. Section II provides a general formulation for SR reconstruction using sparse representation and over-complete dictionary. Section III describes the details of the proposed method. Extensive experimental results and analysis are presented in Section IV. Section V contains the concluding remarks.

II. FORMULATION

We begin our journey with a description of the theory behind SR reconstruction. The SR problem can be mathematically stated as:

$$Y_l = DBGX_h + \nu, \quad (1)$$

Where X_h is the original HR image, D is the down-sampling operator, B is the blurring operator, G is geometric transformation, ν is an additive noise, Y_l is the LR image. Equation (1) means the observed LR image Y_l is the downsampled, blurred, transformed and noisy version of X_h . The goal of this problem is to recover X_h as accurately as possible based on the LR observation Y_l . This is an ill-posed problem and has no unique solution, so we need regularizers to obtain a unique optimal solution for this problem. Since we perform the analysis at the level of small patches, ensuring to make use of the image redundancy, the formulation is rewritten in the following form:

$$P_l^k = D_k B_k G_k P_h^k + \nu_k, \quad (2)$$

Where P_l^k and P_h^k are patches respectively extracted from the LR and HR images at location k . P_h^k is with size $(n \times s) \times (n \times s)$ and P_l^k is with size $n \times n$, s is the upsampling scale. ν_k is the noise on the patch k . Without loss of generality, (2) can be written as

$$P_l^k = M_k P_h^k + \nu_k \quad (3)$$

The assumption of pattern redundancy of MR images means information can be sparsely coded. A patch can be sparsely represented by α over the dictionary A , namely:

$$P_h^k = A_h \alpha_h^k \text{ where } \|\alpha_h^k\| \ll sp, \quad (4)$$

$$P_l^k = A_l \alpha_l^k \text{ where } \|\alpha_l^k\| \ll sp, \quad (5)$$

$\|\alpha_h^k\| \ll sp$ and $\|\alpha_l^k\| \ll sp$ mean the number of non-zeros in vector α_h^k or α_l^k is much smaller than the sparsity sp , which is the number of the atoms of A_h and A_l we use to represent the P_h^k and P_l^k .

According to (3) and (4), we can get $P_l^k = M_k A_h \alpha_h^k + \nu_k$, that is $\|P_l^k - M_k A_h \alpha_h^k\| \leq \epsilon$, where ϵ is related to noise ν_k . So $P_l^k \approx M_k A_h \alpha_h^k$ which means P_l^k can be represented by sparse representation α_h^k over dictionary $M_k A_h$. So LR dictionary and HR dictionary can share the same sparse representation, i.e. $\alpha_h^k = \alpha_l^k$. Both LR and HR dictionaries are over-complete, with more atoms than signal dimensions, allowing to represent a wide range of signal phenomena.

Supported by the above theory, the SR method based on sparse representation and over-complete dictionary is defined as follows: LR and HR dictionaries are trained from the training set. Then the observed LR image is sparsely represented over the trained LR dictionary. As assumed above, the sparse representations of the LR and HR image are the same, so based on the obtained sparse representation and trained HR dictionary, HR image is reconstructed. The above model based on sparse representation can also be referred as sparse land. This model constructs a connection between HR patches and the corresponding LR patches, which is exploited to recover the HR image.

III. Proposed Method

A. Motivations

For the SR reconstruction of single anisotropic 3D MR image based on over-complete dictionary, one key point is the construction of relevant learning database, i.e. the training set. But sometimes no extra training set is provided. To solve this problem, we should construct the training set from the anisotropic 3D MR image itself.

The more similar the observed LR image is to the training examples, the better reconstruction results we may obtain. Furthermore, the training set should include HR images and their corresponding LR images to learn the relationship between them. So the training examples should be HR and similar to the out-of-plane slices of anisotropic 3D MR image. It has been proven in [19] that the local self-similarity of anatomical features occurs both within the same plane and across the planes. That means the out-of-plane patches are similar to the in-plane patches. Meanwhile, the in-plane slices from anisotropic 3D MR image are HR. This is the rationale behind constructing the training set based on the in-plane patches.

Based on the above analysis, this paper utilizes in-plane HR patches from anisotropic 3D MR image to construct the training set for the over-complete dictionaries.

B. The Proposed Algorithm

The proposed SR method includes two main phases: training and reconstruction. The procedure is shown in Algorithm 1. The details are described in the following.

Algorithm 1

The proposed Super-resolution approach.

Input:

I_o : Single anisotropic 3D MR image with a slice dimension $a \times b$ and c slices.

s : In-plane to out-of-plane resolution ratio.

n : Size of patches extracted from LR images.

o : The overlap of LR patches.

Output:

I_i : Upsampled isotropic 3D MR image with size $a \times b \times (c \times s)$.

Step 1 Dictionary training

Step 1.1 Training set construction

1. Collect in-plane HR 2D slices $\{T_h^j\}_j, j=1,2,\dots,c$ with size $a \times b$ from I_o , where $T_h^j = I_o(:, :, j)$.
2. Produce the corresponding LR images $\{T_l^j\}_j, j=1,2,\dots,c$ via averaging the near rows or columns.
3. Extract overlapped patches $\{P_h^k\}_k$ and $\{P_l^k\}_k: P_h^k = R_h^k * T_h^j, P_l^k = R_l^k * T_l^j$, where k is the location of the patches, R_h^k/R_l^k means extract the HR/LR patches from corresponding HR/LR slices at location k . P_h^k is with size $(n \times s) \times (n \times s)$, P_l^k is with size $n \times n$, overlapped voxels number for P_h^k and P_l^k are respectively $(o \times s)$ and o .

The patch pairs $\{P_h^k, P_l^k\}_k$ construct the training set.

Step 1.2 Pre-processing

1. Remove the low-frequencies from $\{P_h^k\}_k$, and extract features $\{P_{fl}^k\}_k$ from $\{P_l^k\}_k: P_{fl}^k = filters * P_l^k, filters = \{G, G^T, L, L^T\}$, where $G = [1, 0, -1]$, $L = [1, 0, -2, 0, 1]/2$.
2. Perform dimensionality reduction by principal component analysis (PCA) over $\{P_{fl}^k\}_k$.

Step 1.3 Dictionary training: Based on the processed training set, train over-complete dictionary A_l and sparse representation $\{\alpha_{fl}^k\}_k$ for $\{P_{fl}^k\}_k$ by K singular value decomposition

(K-SVD) algorithm companied with orthogonal matching pursuit (OMP) method, so that

$$P_{fl}^k = A_l * \alpha_{fl}^k. \text{ Then obtain corresponding } A_h \text{ for HR patches based on } \{\alpha_{fl}^k\}_k \text{ to satisfy}$$

$$P_h^k = A_h * \alpha_{fl}^k.$$

Step 2 Isotropic 3D MR image reconstruction Step

Step 2.1 Pre-processing

1. Interpolate each image of $\{Z_l^i, i=1, 2, \dots, b\}_i$ with a slice dimension $a \times c$ to the destination size $\{X_l^i\}_i$ using bicubic interpolation algorithm by factor s .
2. Cut every LR slice X_l^i into patches $\{P_x^k\}$ with size $n \times n$ and overlapped voxels o .
3. Extract features $\{P_{fx}^k\}$ from $\{P_x^k\}$ by the method used in the training phase.
4. Dimensionality reduction is performed again by PCA over $\{P_{fx}^k\}$.

Step 2.2 Reconstruction: For each $X_h^i, i=1, 2, \dots, b$,

1. Sparse code $\{P_{fx}^k\}$ by OMP method using the trained LR dictionary A_l and get the corresponding sparse representations $\{\alpha_{fx}^k\}$.
2. Recover the HR patches P_{hx}^k by multiplying $\{\alpha_{fx}^k\}$ and HR dictionary A_h .
3. Add low frequency to HR patches.
4. Merge the HR patches by averaging the overlapped parts, and get the final reconstructed slice X_h^i . I_u is the 2D slice stack of X_h^i .

1) Dictionary Training Phase: The dictionary training stage can be divided into three parts: training set construction, image pre-processing, sparse representation and dictionary training.

a. Training Set Construction: The training set is constructed by the HR in-plane slices. When constructing the training set, we obtain the corresponding LR images by averaging adjacent rows or columns. This simulates PVE (Partial Volume Effects) as a consequence of LR image acquisition. Besides, we cut the images into small patches to form the training set. This process is to utilize the redundancy of the images and reduce the computation time.

b. Image Pre-Processing: We subtract low frequency information from HR patches and extract the structure features for each patch, so that the dictionary represents image textures rather than absolute intensity. Here, the low frequency is the mean pixel value, which is the same with the LR patches. As mentioned before, the HR image loses its high-frequency information through the acquisition process, and our task is to recover the high-frequency information. That is why we use the high frequency features as the examples to train the dictionary. Another pre-processing operation is dimensionality reduction over the feature vectors by PCA. This simultaneously reduces computations and improves the reconstruction accuracy.

c. Dictionary Training and Sparse Representation: The dictionary and the corresponding sparse representation encode the connection between the LR patches and the corresponding HR patches. Based on this connection the HR patches can be reconstructed. The objective of this step is to express the LR patches by dictionary and sparse representation as accurately as possible based on the sparsity prior. This is an optimization problem which can be mathematically expressed in the following way:

$$A_l, \{\alpha_{fl}^k\} = \underset{A_l, \{\alpha_{fl}^k\}}{\operatorname{argmin}} \sum_k \|P_{fl}^k - A_l \alpha_{fl}^k\|^2$$

$$s.t. \|\alpha_{fl}^k\|_0 \leq sp \forall k, \quad (6)$$

An efficient algorithm is needed to obtain the best A_l and α_{fl}^k . In this paper, we choose K-SVD [20] to train the dictionary for the sparse representation. The K-SVD algorithm is an efficient iterative method that alternates between sparse coding based on the current dictionary and updating the dictionary atoms to better fit the examples. This method is generalized from the K-means clustering process. It is flexible and can work with any pursuit method. In this paper, we use the OMP (orthogonal matching pursuit) [21] algorithm as it is simple and only involves the computation of inner products of matrices.

2) Up-sampling LR Anisotropic 3D MR Image: Based on the HR and LR dictionaries trained from the anisotropic 3D image itself, we up-sample the image through the following steps. Firstly, feature extraction and dimensionality reduction are performed again as in the dictionary training phase. The next step is reconstruction. For each X_l^i , we get the sparse code $\{\alpha_{fx}^k\}$ for $\{P_{fx}^k\}$ by OMP method using the trained LR dictionary A_l based on (6). Because HR patches and corresponding LR patches share the same sparse representation, we can get the HR patches by multiplying $\{\alpha_{fx}^k\}$ and the trained HR dictionary A_h . In the training phase, we subtract low frequency from the HR patches. Therefore, the recovered HR patches so far do not contain the low frequency information. We should add the low frequency to the reconstructed HR patches. In this method, we process small patches not the whole image, so we should connect all the small patches into the whole image. The final HR image is constructed by solving the following minimization problem with respect to X_h^i :

$$X_h^i = \underset{X_h^i}{\operatorname{argmin}} \sum_k \|R_k(X_h^i - X_l^i) - P_{hx}^k\|_2^2, \quad (7)$$

where R_k means an extractor to extract patches at location k from high frequency resulting image, $(X_h^i - X_l^i)$. The extracted patches should be as close as possible to the reconstructed patches P_{hx}^k . This problem can be solved by the following equation:

$$X_h^i = X_l^i + [\sum_k R_k^T R_k]^{-1} \sum_k R_k^T P_{hx}^k, \quad (8)$$

It is equivalent to putting P_{hx}^k in their proper location, averaging the overlap regions, and adding the low frequency content of X_l^i to generate the final image X_h^i .

IV. Experiments

To demonstrate the advantages of the proposed approach, we conducted extensive experiments. This section has been divided into 4 parts. The first part describes implementation details including the selection of the parameters in the proposed SR reconstruction process. In the second part we introduce the experimental data sets. Then, in part 3, we talk about the quantitative and qualitative evaluation methods. Finally, in part 4 we have compared the proposed method with classical interpolation and a state-of-the-art upsampling method based on the Non-Local Means approach [13] to demonstrate the superiority and impact of the proposed method. Based on the results of experiments, we analyze how the slice thickness, noise, and pathology affect the accuracy of the proposed method.

A. Implementation Details

All algorithms were implemented in MATLAB R2014a, running on a Windows machine with 2 3.10 GHZ Intel Core i5 CPUs and 4.00GBytes of RAM. Since the proposed methodology may be implemented in a number of different manners, we clarify the following implementation details: K-SVD was chosen as the dictionary training method and OMP as the sparse representation method; Based on Table I and Table II, which present the reconstruction results of simulated axial T2W MR image with 2mm slice thickness, in a trade-off between accuracy and efficiency, we always used 3×3 LR patches with 1 pixel overlap between adjacent patches, corresponding to $(3 \times scale) \times (3 \times scale)$ patches with overlap of $(1 \times scale)$ for the HR patches. Feature extraction was done using gradient and Laplacian filters. For initial upsampling we used bicubic interpolation; for the dictionary training phase, 40 iterations was experimentally found to provide a good trade-off between the efficiency and accuracy; number of dictionary atoms and the maximum sparsity for the sparse representation were set to 512 and 3, respectively, following [16].

B. Brain MR Data Sets

To validate the proposed method, a synthetic dataset and several real MR images were used. Various simulated T2-weighted (T2W) brain MR images were obtained from the publicly available BrainWeb database [22], including normal and pathologic (multiple sclerosis) MR images, non-noisy and noisy ones. The HR T2W volumes had $181 \times 217 \times 181$ voxels with a resolution of $1 \text{ mm} \times 1 \text{ mm} \times 1 \text{ mm}$. Different percentage noise (0%, 1%, 3%, 5%, 7% and 9%) levels were used to investigate the noise influence. The noise in the simulated images has Rayleigh statistics in the background and Rician statistics in the signal regions. The “percent noise” number represents the percent ratio of the standard deviation of the Gaussian

white noise versus the signal for a reference tissue. For T2W images, the reference tissue is CSF (Cerebrospinal Fluid).

To test the proposed approach on real clinical data, three FSE (Fast Spin Echo) T2W brain MR images were collected, with different slice-selection directions but for the same subject. Those three-plane MR images all had a slice thickness of 2.0 mm, and a pixel size of 0.46875 mm \times 0.46875 mm. The axial slice stacks had a slice dimension of 408 \times 512 and 80 slices, the coronal scan had a slice dimension of 408 \times 512 and 100 slices, while the sagittal scan had a slice dimension of 512 \times 512 and 83 slices.

C. Evaluation Method

To quantitatively and qualitatively evaluate the performance of the proposed method over different brain data sets, we introduce four different methods in this section for two scenarios:

1) Images with ground truth—In the experiments, if we have an original HR image, considered as the ground truth, comparing the reconstruction with the original image is a good way to evaluate the results. The following two performance metrics are calculated when the ground truth is available:

Peak Signal-to-Noise Ratio (PSNR) is defined as:

$$\text{PSNR}(X_o, X_h) = 10 \cdot \log_{10} \left(\frac{d}{\text{MSE}(X_o, X_h)} \right), \quad (9)$$

Where $\text{MSE}(X_o, X_h)$ stands for means square error, quantifies the pixel intensity difference between the original HR image X_o and the corresponding SR reconstruction X_h , using

$\text{MSE}(X_o, X_h) = \frac{1}{|\Omega|} \sum_{k \in \Omega} |x_o^k - x_h^k|$, x_o^k and x_h^k are the image intensity at location k , d is the dynamic range of the intensity value, i.e. $d = \max(X_o) - \min(X_o)$. Typically, the PSNR values are between 25 dB and 50 dB. A higher value of PSNR indicates a better performance of the reconstruction method.

Structural Similarity Image Metric (SSIM) [23]: It measures the similarity between two images, with a definition that is more consistent with the human visual perception of image quality. Under the assumption that human visual perception is highly adapted to extracting structural information from a scene, SSIM is formulated as:

$$\text{SSIM}(X_o, X_h) = \frac{(2\mu_x\mu_y + C_1)(2\sigma_{xy} + C_2)}{(\mu_x^2 + \mu_y^2 + C_1)(\sigma_x^2 + \sigma_y^2 + C_2)}, \quad (10)$$

Where μ_o and μ_h are the mean intensity of images X_o and X_h , respectively; σ_o and σ_h are the standard deviation of images X_o and X_h , which are estimates of the signal contrast; σ_{oh} is the covariance of X_o and X_h , $C_1 = (K_1L)^2$ and $C_2 = (K_2L)^2$, $K_1 \ll 1$ and $K_2 \ll 1$ are small

constants and L is the dynamic range of the intensity values. In this paper we use $K_1 = 0.01$ and $K_2 = 0.03$. SSIM values are between 0 and 1, where a higher value indicates the better reconstruction results.

2) Images without ground truth—In reality, for example in clinical data, no original HR reference image is available, so we cannot evaluate the reconstruction results by comparing the similarity with a ground truth image. Alternative methods to evaluate the results are as follows:

Visual inspection: visual assessment of images is also a precious method to compare and judge the benefit of proposed methods; however, it is obviously a subjective method, and also may not be easy when large datasets should be evaluated and compared. In this paper, we display several 2D reconstructed slices and evaluate the slices by viewing the image details.

Intensity profile: the intensity profile of an image is the set of intensity values taken from regularly spaced points along a line segment or multiline path in an image. The fundamental problem of SR reconstruction can be stated as restoring some high-frequency information (like edges) that has been lost during the acquisition process. An effective SR reconstruction technique should be able to recover these high-frequencies. Intensity profile can show intensity value changes at the interfaces between different tissues, thus may be used as a surrogate measure of how edge features appear and are distinguished in the image. We also evaluate the reconstruction results of our clinical MR experiments based on image intensity profiles in this paper.

D. Experimental Results and Analysis

In this part, we compare our proposed method with classical interpolation algorithms and a state-of-the-art single-image SR approach on both simulated database and real medical image. Furthermore, we analyze the influence of different factors to the proposed method based on the experiment results.

1) Comparison with Classical Interpolation Methods—To evaluate the efficacy of the proposed method, we perform comparisons with classical interpolation algorithms, including the nearest neighbor, bilinear, bicubic, B-spline interpolation. Different MR images have various features: different slice thickness, noise and lesions. So we compared the proposed method with the classical interpolation method on 3D T2W MR images with different features respectively.

Firstly, we constructed a down-sampled version of a normal non-noisy simulated HR T2w image. Axial slice stacks with different slice thickness (2 mm, 3 mm, 4 mm, 5 mm, 6 mm and 7 mm) were simulated. Adjacent slices were averaged to produce different slice thicknesses. This simulates the Partial Volume Effect (PVE). PVE increases as the slice thickness increases. For example, three adjacent slices along the Z direction were averaged into one slice to simulate an anisotropic acquisition at $1 \text{ mm} \times 1 \text{ mm} \times 3 \text{ mm}$ resolution. The slice thickness became 3 mm and the matrix size was $180 \times 216 \times 60$. The out-of-plane slices were reconstructed based on the dictionary trained from the in-plane slices in the axial

direction. These simulated LR 3D MR images were upsampled to $1\text{ mm} \times 1\text{ mm} \times 1\text{ mm}$. To see the relative degree of improvements, in this section we report the results of comparing the classic interpolation algorithm and the proposed method over normal non-noisy axial 2D slice stacks with slice thickness of 2 mm, 3 mm, 4 mm, 5 mm, 6 mm, 7 mm. Table III shows that the bicubic and B-spline interpolations generate very similar results and the proposed method generates the best results in terms of both PSNR and SSIM values. The improvements achieved in these metrics by using the proposed method are comparable to the amount of improvement obtained from higher-order interpolation methods (bicubic and B-spline) as compared to the nearest neighbor interpolation. This indicates major improvements. The PSNR/SSIM values obtained from the proposed method are 31.555 dB/0.9849 in 2 mm and 26.791 dB/0.9544 in 3 mm. The PSNR/SSIM values drop rapidly as the slice thickness increases from 2 mm to 3 mm. The reason for the influences is that the up-sampling scalar increases as the slice becomes thicker. The limitation of most SR algorithms is that their performance deteriorates quickly when the magnification factor is only moderately large.

It is clear that the 0% noise case is an idealization of the real MR image acquisition. To compare the classic interpolation method and proposed method on noisy 3D MR images, another experiment was performed on T2W simulated MR images from BrainWeb with different noise levels (0%, 1%, 3%, 5%, 7% and 9%). Noisy axial MR images with voxel size $1\text{ mm} \times 1\text{ mm} \times 2\text{ mm}$ and matrix size $180 \times 216 \times 90$ were simulated. The resolution in the slice-selected direction is improved based on the trained dictionary. The results are shown in Table IV. In addition, similar experiment is repeated using the MS (Multiple Sclerosis) T2W MR images, as shown in Table V. Again, the proposed method obtains the best results in all cases, including the noisy and pathological images. The PSNR/SSIM values drop as the noise level increases.

Experiments were also performed on real clinical data. Axial 2D slice stacks (voxel size: $0.46875\text{ mm} \times 0.46875\text{ mm} \times 2.0\text{ mm}$) was up-sampled to $1\text{ mm} \times 1\text{ mm} \times 1\text{ mm}$ using bicubic interpolation and the proposed method. In Fig. 1, a visual comparison of the results is shown. One can see that the reconstruction using the proposed approach shows a better anatomical content. A close up of the image clearly shows the reconstruction using the proposed method is significantly less blocky and blurry. The proposed method is visually superior in particular near the image edges.

2) Comparison with Non-Local Means SR method—Furthermore, as mentioned before, we compared the proposed method with a recent SR method based on NLM (Non-local Means) without a reference image [13]. This technique was also shown to outperform the standard interpolation methods. First, simulated normal non-noisy axial 2D slice stacks, with 2 mm-7 mm slice thickness, were up-sampled by NLM and the proposed method. The results can be observed in Table VI. For slice thickness of 2 mm and 3 mm, the NLM method performs slightly better than the proposed approach. That is because the NLM algorithm optimizes the reconstruction results by mean preservation constraint, which has not been implemented in our proposed method. Although this step could have slightly improved the accuracy of our proposed method, it is time-consuming. Considering the trade-off between accuracy and computation cost, we omit this optimization step to make the

proposed method more suitable for clinical applications. In addition, those two methods generate very similar results at higher upsampling factors (i.e. for slice thicknesses of 4 mm, 5 mm, 6 mm, and 7 mm). In general, the PSNR/SSIM values drop rapidly as the slice thickness increases.

Next we reconstructed normal and noisy axial 2D slice stacks with 2 mm slice thickness by NLM and the proposed method. Table VII shows that the NLM method generates slightly better results when the simulated MR images have 0%, 1% and 3% noise. However, when the noise percentages are 5%, 7% and 9%, the proposed method generates comparable results as NLM does. Again, the PSNR/SSIM values drop as the noise level increases.

Based on the above two experiments, we conclude that the proposed method generates results that are comparable to the NLM approach in clinical cases. In clinical applications, the in-plane and out-plane resolution ratio of FSE T2W images is often bigger than 3, which means that the anisotropic MR images should be upsampled into isotropic volumes by a factor of 3 or more. The noise percentage is also usually more than 3%. The proposed method is therefore, practically as effective as the NLM method, but is more efficient.

To verify our conclusion, the NLM and the proposed methods were respectively applied to clinical FSE T2W MR images, including axial, coronal and sagittal scans. The detail information of the clinical MR images were discussed in Section IV-B. The results are displayed in Fig. 2, 3, and 4. Visual comparison shows that these two methods generate similar results, and our proposed method generates slightly shaper intensity profiles which indicate better delineation of image edge features. Table VIII shows that our proposed method needs significantly less computation time and memory than the NLM method. Average computation time for the proposed method was about 2.9 minutes whereas the average computation time for NLM was 6 minutes. Moreover, average peak memory for the proposed method was about 107 Mb, while the average peak memory for the NLM was 245Mb in this experiment. Overall, we conclude that the proposed method outperforms NLM over the clinical MR images.

3) Impact of the training set size—Based on the idea of the proposed method, the training set is extracted from the LR 3D MR image itself, i.e. from the HR in-plane slices. There are two choices to construct the training set: one is to select all the HR in-plane slices as the training examples, so the training set size is related to the number of slices; the other is to select only part of them to construct the training set. To test how the training set size impact the proposed method performance, we selected all the HR slices and randomly extracted part of the HR slices (including 80%, 60%, 40% and 20%) from the in-plane direction to construct the training set respectively. The reconstruction results of non-noisy axial 3D MR based on training sets of different size have been shown in Fig. 5, 6, 7. We repeated the experiments 10 times and calculated the average. Based on the results, we observed the PSNR and SSIM values decrease slightly as the training set size reduces, while the training time drops rapidly. This shows that the algorithm is robust to the training set size. But when the slice thickness is 7mm, if 20% of the available data were extracted, the proposed method is worse than the classic interpolation method (bicubic method). That is because while slice thickness increasing, the available data decreases rapidly. If part of the

available data were extracted, the training set would be extremely small. That will affect the performance of the proposed method. Therefore in practice, to ensure the best performance, all available HR slices from the in-plane direction should be extracted to train the dictionary, which is already very efficient and fast than the state-of-art algorithm (NLM method).

V. Conclusion

This paper presents a novel SR approach towards single anisotropic 3D MR image reconstruction based on sparse representation and over-complete dictionary without extra training sets. We train the dictionary from the in-plane HR slices. Our proposed method outperforms the classical interpolation algorithms. Furthermore, the proposed SR approach is compared with a recent single MR image SR method based on the NLM approach. Experiments show both methods can generate similar results in clinical applications, but our proposed algorithm is more efficient than the NLM-based method in terms of computation time and memory usage. The proposed approach may be used as an efficient method for upsampling anisotropic MR images in the out-of-plane dimensions.

Acknowledgments

This work was supported in part by the China Science and Technology Project of Ministry of Transport under Grant 2011318740240, by the Chongqing graduate education reformation research project under the No.yjg133005, by the Scientific and Technological Research Program of Chongqing Municipal Education Commission under Grant no. KJ1400409, and in part by the National Institutes of Health grant R01 EB018988, R01 EB013248, and R03 DE022109.

We are very thanks to the anonymous reviewers for their useful comments.

REFERENCES

1. Greenspan H, Oz G, Kiryati N, Peled S. MRI inter-slice reconstruction using super-resolution. *Magn. Reson. Imaging*. 2002 Jun.20(5):437–446. [PubMed: 12206870]
2. Reeth EV, Tham IWK, Tan CH, Poh CL. Super-resolution in Magnetic Resonance Imaging: A review. *Concept in Magn. Reson*. 2012 Nov.40A(6):306–325.
3. Plenge E, Poot DHJ, Bernsen M, Kotek G, Houston G, Wielopolski P, Weerd LVD, Niessen WJ, Meijering E. Super-resolution methods in MRI: Can they improve the trade-off between resolution, signal-to-noise ratio, and acquisition time? *Magn. Reson. Med*. 2012 Nov.68(6):1983–1993. [PubMed: 22298247]
4. Peled S, Yeshurun Y. Superresolution in MRI: application to human white matter fiber tract visualization by diffusion tensor imaging. *Magn. Reson. Med*. 2001 Jan.45(1):29–35. [PubMed: 11146482]
5. Scheffler K. Superresolution in MRI? *Magn. Reson Med*. 2002 Aug.48(2):408–408. [PubMed: 12210953]
6. Carmi E, Liu SY, Alon N, Fiat A, Fiat D. Resolution enhancement in MRI. *Magn. Reson. Imaging*. 2006 Feb.24(2):133–154. [PubMed: 16455402]
7. Tieng QM, Cowin GJ, Reutens DC, Galloway GJ, Vegh V. MRI resolution enhancement: how useful are shifted images obtained by changing the demodulation frequency? *Magn. Reson. Med*. 2011 Mar.65(3):664–672. [PubMed: 20928827]
8. Ziye, Y., Yao, L. Super resolution of MRI using improved ibp. *Int. Conf. on Computational Intelligence and Security (CIS)*; London, England. 2009. p. 643-647.
9. Gholipour, Estroff, JA., Sahin, M., Prabhu, SP., Warfield, SK. *Medical Image Computing and Computer-Assisted Intervention - MICCAI, Lecture Notes in Computer Science*. Vol. 13. Beijing,

- China: 2010. Maximum A Posteriori estimation of isotropic high-resolution volumetric MRI from orthogonal thick-slice scans; p. 109-116.
10. Shilling RZ, Robbie TQ, Bailloeu T, Mewes K, Mersereau RM, Brummer ME. A super-resolution framework for 3-d high-resolution and high-contrast imaging using 2-d multislice MRI. *IEEE Trans. Med. Imaging*. 2009 May; 28(5):633–644. [PubMed: 19272995]
 11. Gholipour, Estroff JA, Warfield SK. Robust super-resolution volume reconstruction from slice acquisitions: application to fetal brain MRI. *IEEE Trans. Med. Imaging*. 2010 Oct.29(10):1739–1758. [PubMed: 20529730]
 12. Buades, Coll B, Morel JM. A review of image denoising algorithms, with a new one. *Multiscale Model. Sim.* 2005; 4(2):490–530.
 13. Manjon JV, Coupe P, Buades A, Fonov V, Collins DL, Robles M. Non-Local MRI upsampling. *Med. Image Anal.* 2010 Dec.14(6):784–792. [PubMed: 20566298]
 14. Rousseau F. A non-local approach for image super-resolution using intermodality priors. *Med. Image Anal.* 2010 Aug.14(4):594–605. [PubMed: 20580893]
 15. Jafari-Khouzani K. MRI upsampling using feature-based Non-local means approach. *IEEE Trans. Med. Imaging*. 2014 Jun.33(10):1969–1985. [PubMed: 24951680]
 16. Zeyde, R., Eald, M., Protter, M. On single image scale-up using sparse-representations. *Proc. Of the 7th Inter. Conf. Curves and Surfaces; Berlin, Heidelberg.* 2012. p. 711-730.
 17. Yang JC, Wright J, Huang TS, Ma Y. Image super-resolution via sparse representation. *IEEE Trans. Image. Process.* 2010 May; 19(11):2861–2873. [PubMed: 20483687]
 18. Rueda, Malpica N, Romero E. Single-image super-resolution of brain MR images using overcomplete dictionaries. *Med. Image Anal.* 2013 Jan.17(1):113–132. [PubMed: 23102924]
 19. Plenge, E., Poot, DHJ., Niessen, WJ., Meijering, E. Super-resolution reconstruction using cross-scale self-similarity in multi-slice MRI. *16th Intern. Conf. MICCAI; Nagoya, Japan.* 2013. p. 123-130.
 20. Aharon M, Elad M, Bruckstein A. K-SVD: an algorithm for designing overcomplete dictionaries for sparse representation. *IEEE Trans. Signal Process.* 2006 Nov.54(11):4311–4322.
 21. Tropp JA. Greed is good: Algorithmic results for sparse approximation. *IEEE Trans. Inf. Theory.* 2004 Oct.50(10):2231–2242.
 22. Cocosco CA, Kollokian V, Kwan RK-S, Evans AC. Brain Web: online interface to a 3D MRI simulated brain database. *Proc. of Third Intern. Conf. on Functional Mapping of the Human Brain.* 1997; 5(4)
 23. Wang Z, Bovik AC, Sheikh HR, Simoncelli EP. Image quality assessment: from error visibility to structural similarity. *IEEE Trans. Image Process.* 2004 Apr.13(4):600–612. [PubMed: 15376593]

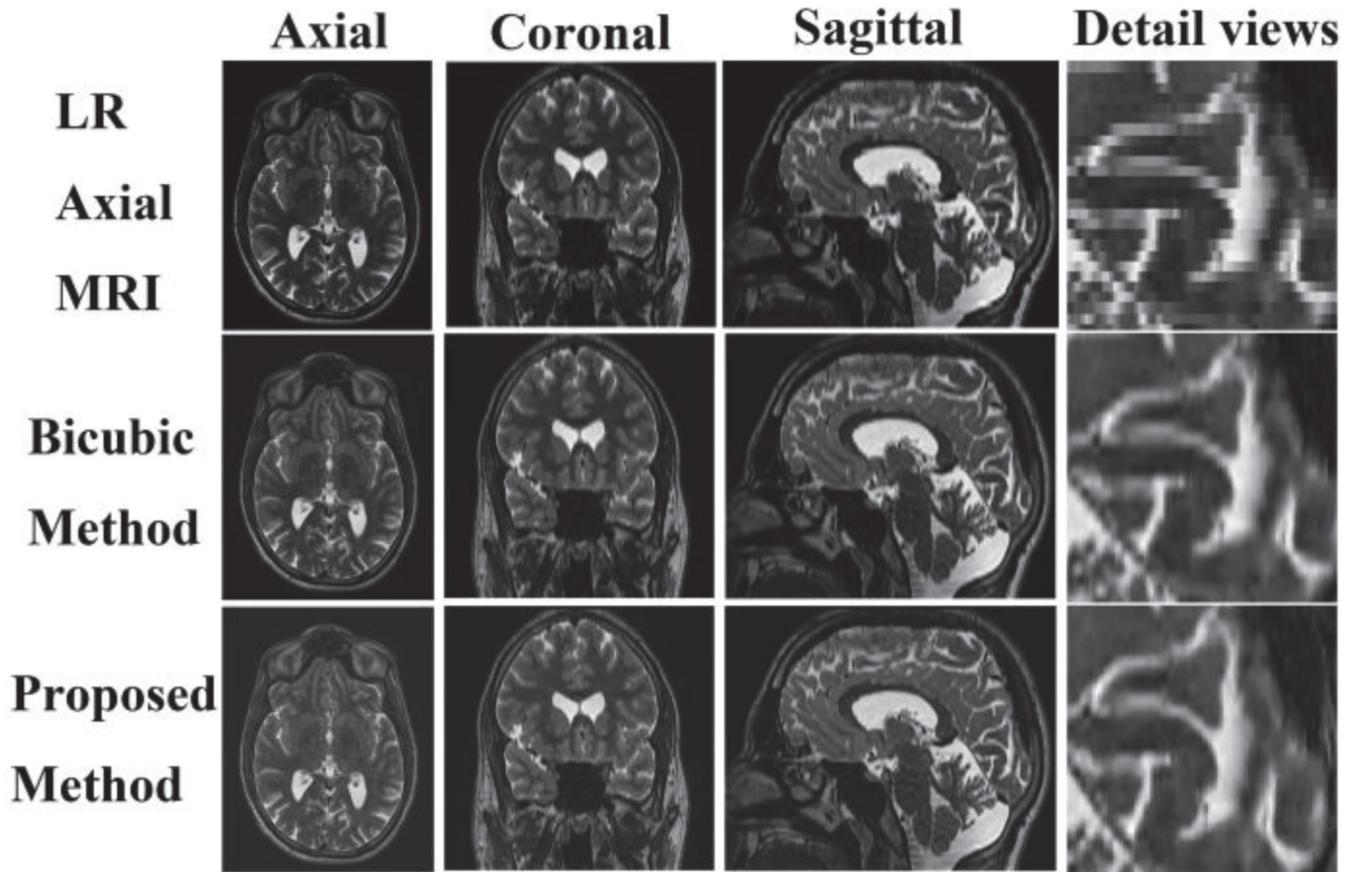


Fig. 1. Clinical axial slice stacks reconstruction using bicubic interpolation and the proposed method

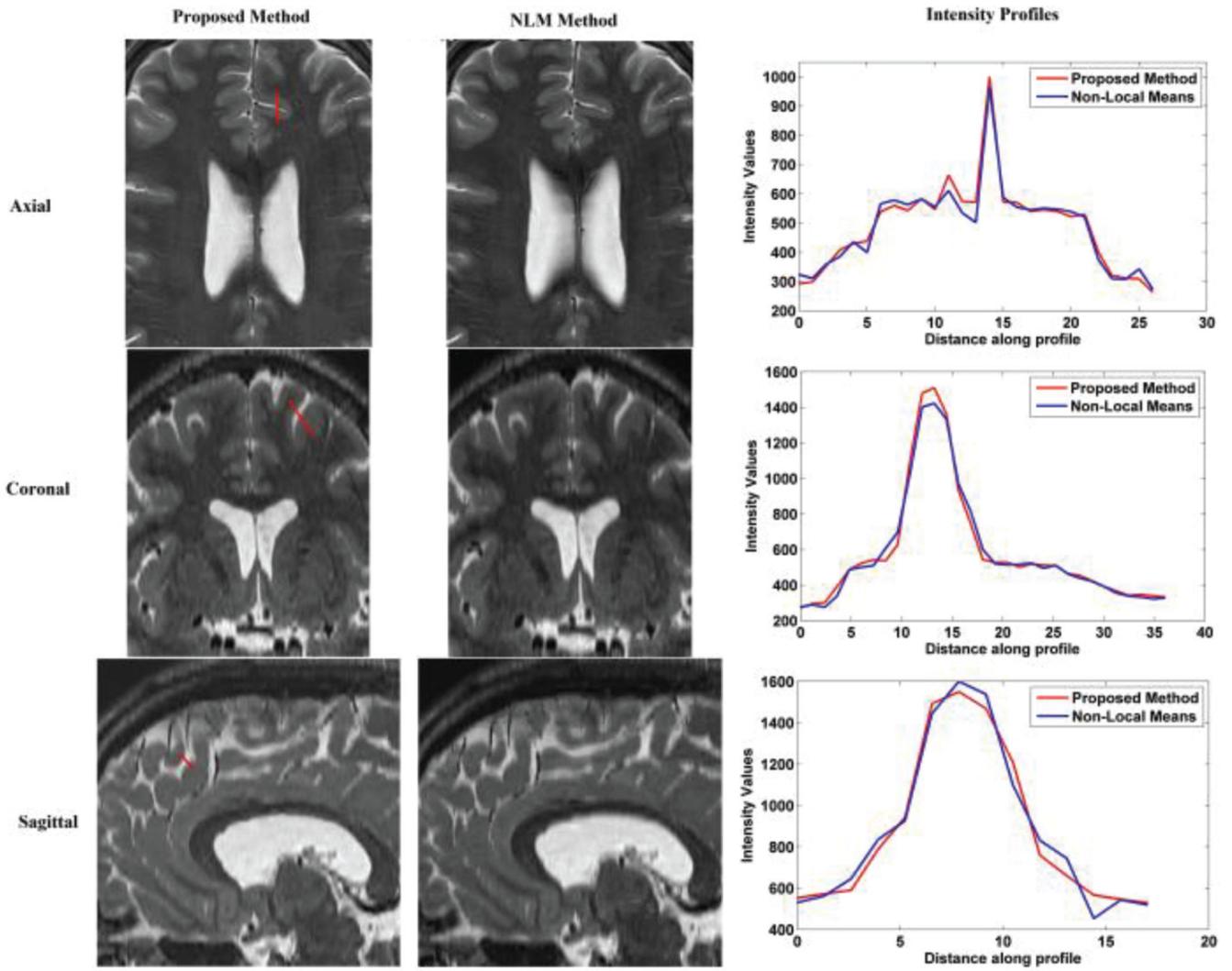


Fig. 2. Reconstruction using NLM and the proposed method over axial slices stacks

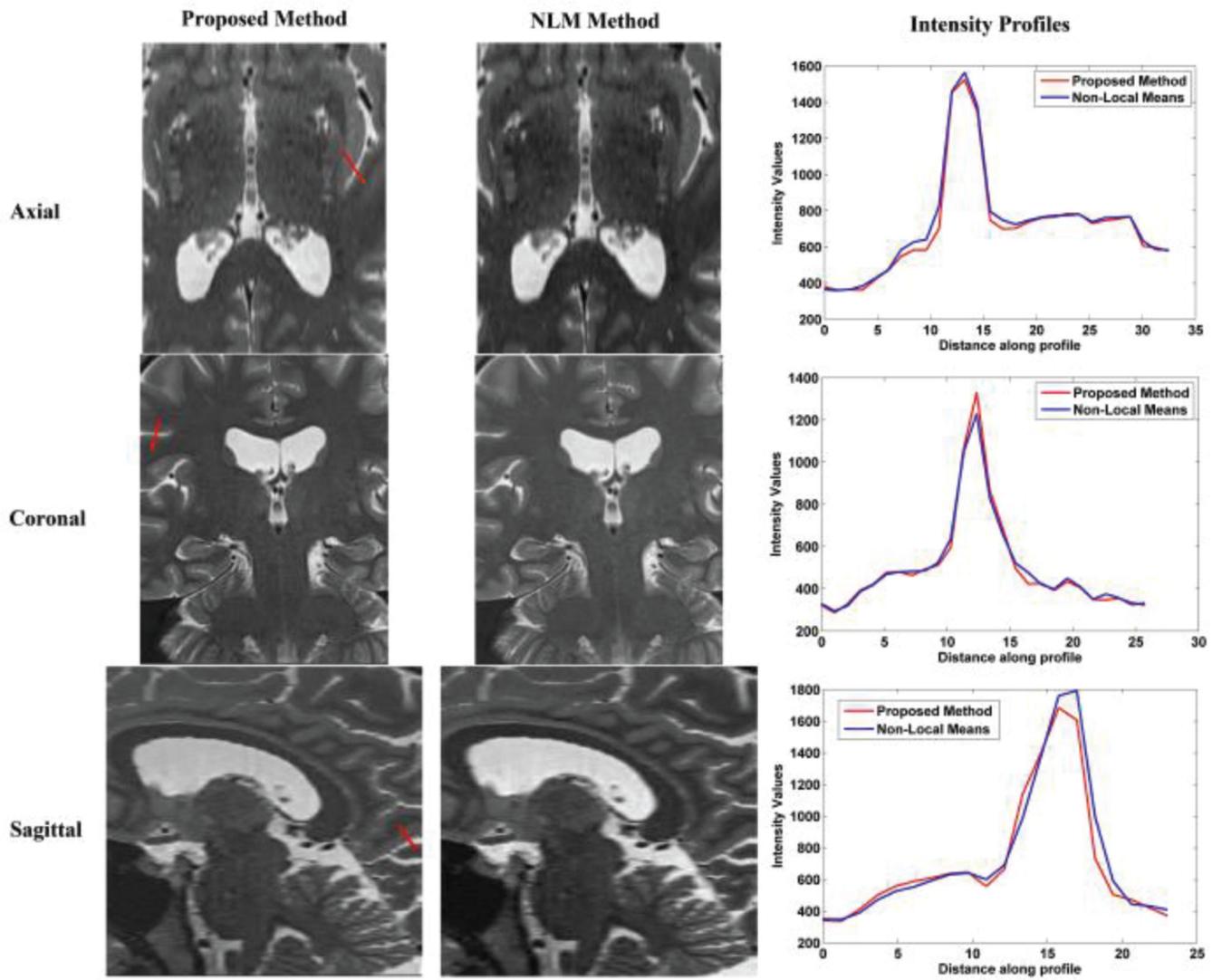


Fig. 3. Reconstruction using NLM and the proposed method over coronal slices stacks

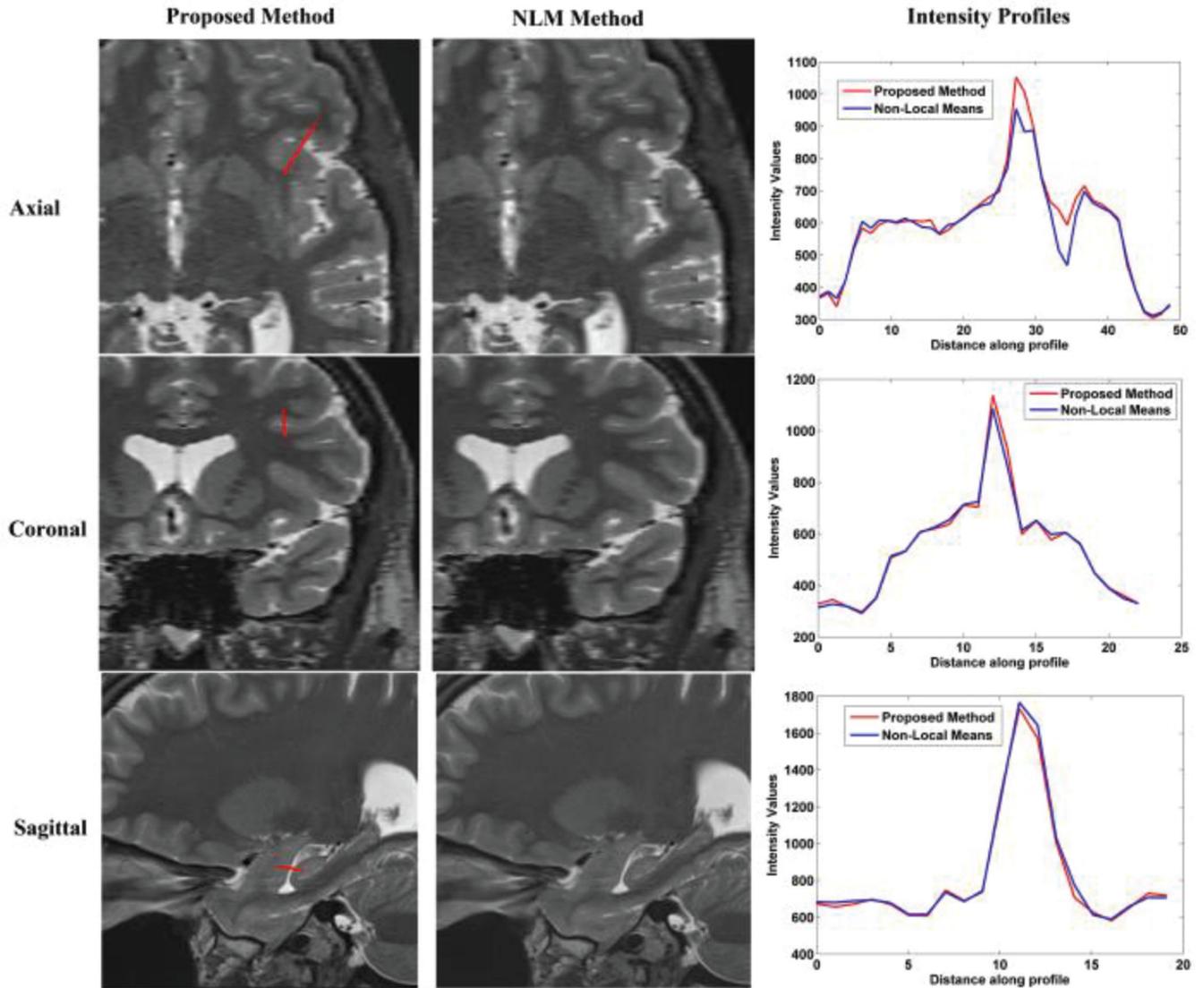


Fig. 4. Reconstruction using NLM and the proposed method over sagittal slices stacks

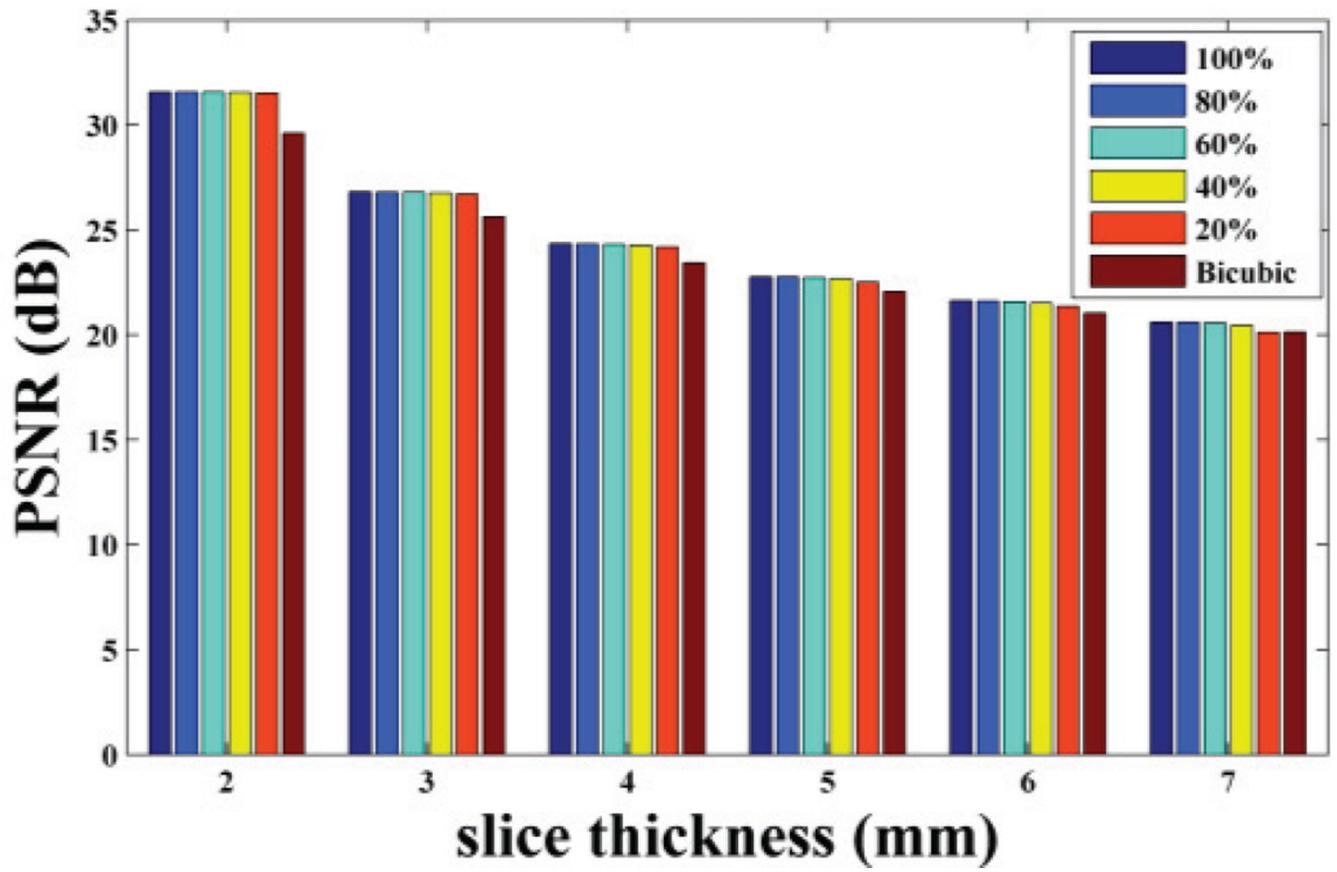


Fig. 5.
PSNR value of the reconstruction results based on training set with different sizes

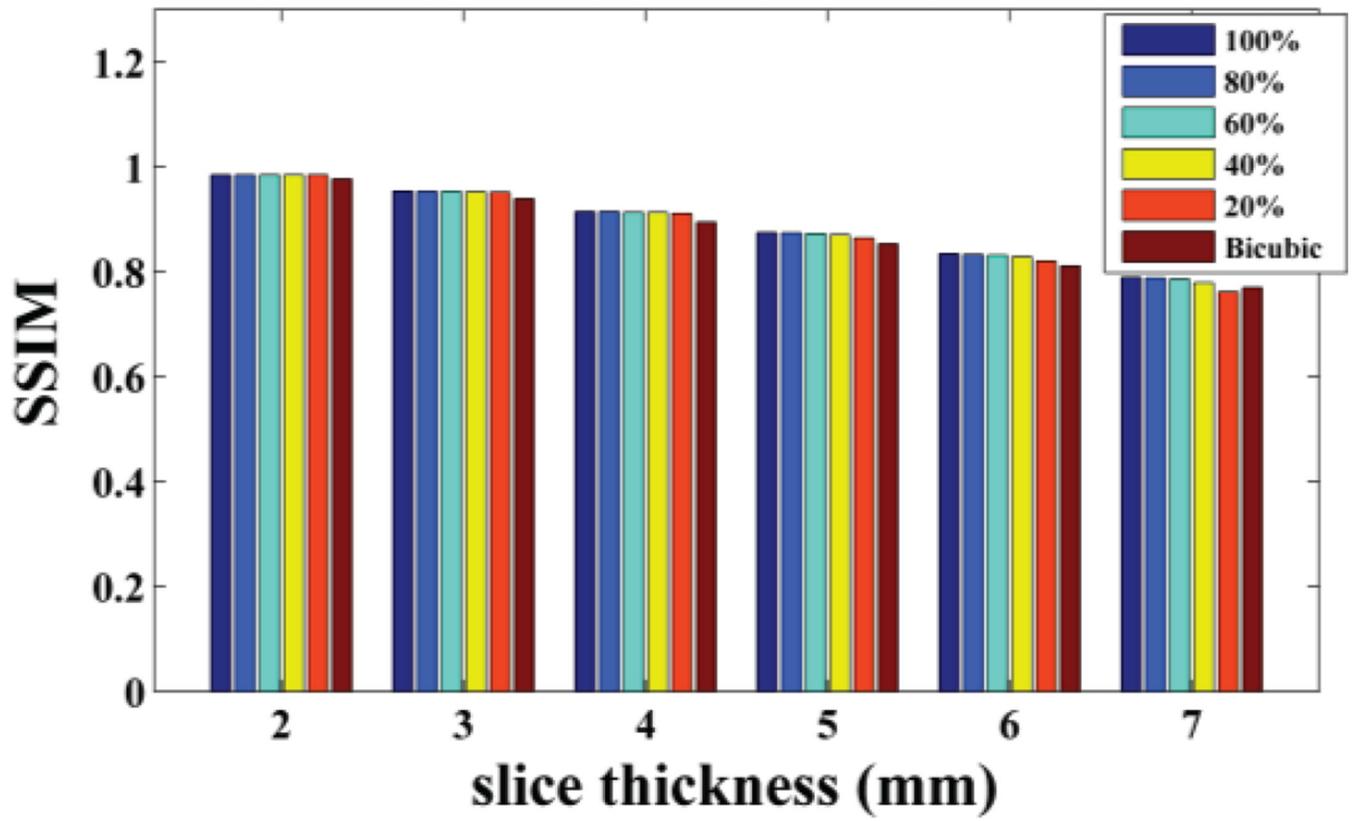


Fig. 6. SSIM value of the reconstruction results based on training set with different sizes

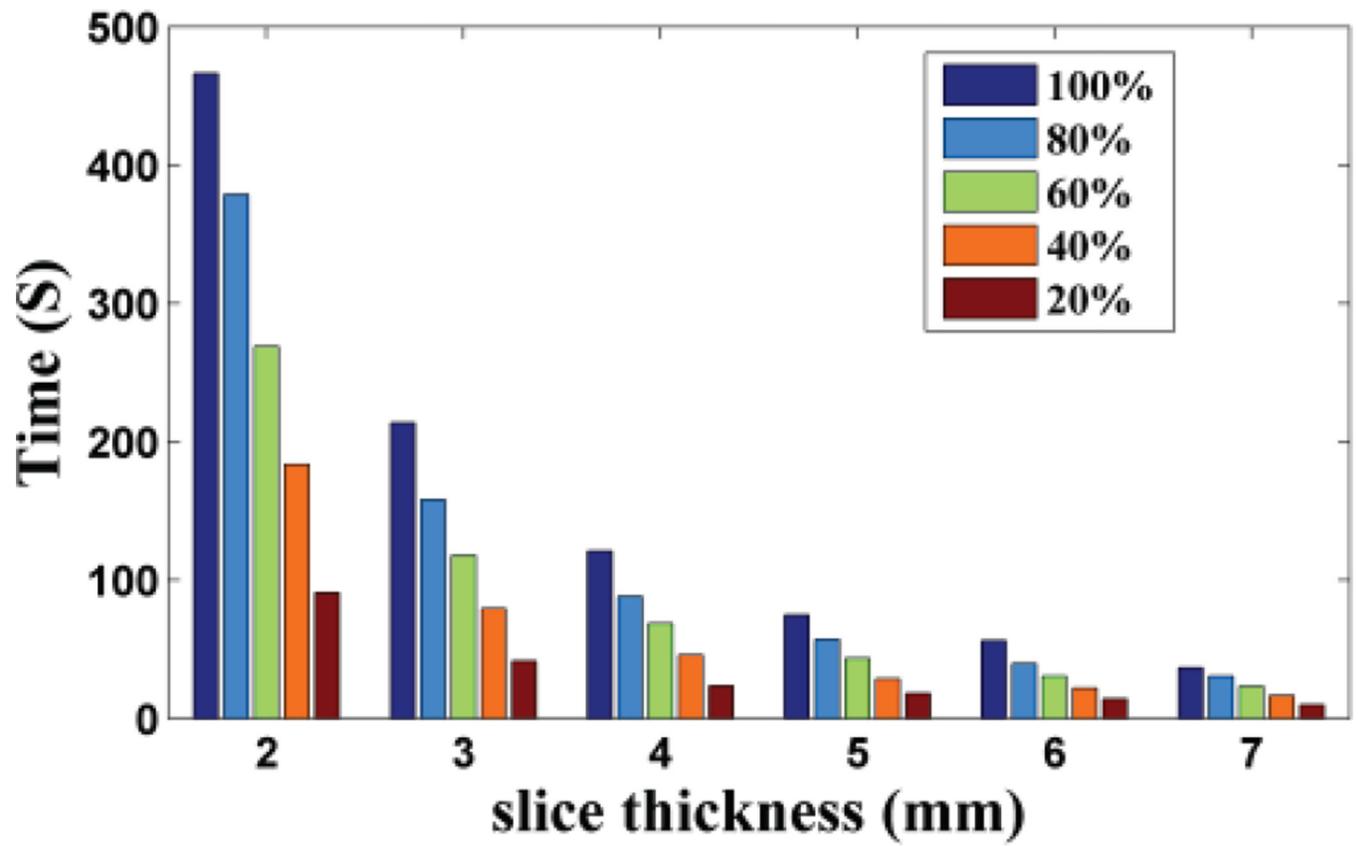


Fig. 7.
Training time of training set with different sizes.

Table I

Reconstruction results using the proposed method over different patch sizes

Patch Size	3×3	5×5	7×7	9×9
PSNR(dB)	31.5719	31.2039	30.8332	30.6010
SSIM	0.9845	0.9830	0.9802	0.9784
Time(s)	466.2860	142.3267	81.7516	57.4088

Author Manuscript

Author Manuscript

Author Manuscript

Author Manuscript

Table II

Reconstruction results using the proposed method based on 3×3 patch over different overlap pixels

overlap	0	1	2
PSNR(dB)	31.1907	31.5719	31.7358
SSIM	0.9830	0.9845	0.9850
Time(s)	192.8939	466.2860	1701.6480

Author Manuscript

Author Manuscript

Author Manuscript

Author Manuscript

Reconstruction results using classical interpolation method and the proposed method over non-noisy MR images

Table III

Slice Thickness	2mm		3mm		4mm		5mm		6mm		7mm	
	PSNR (dB)	SSIM										
Nearest Neighbor	27.105	0.9634	23.886	0.9178	22.098	0.8719	20.939	0.8305	20.142	0.7963	19.373	0.7576
Bilinear	28.254	0.9673	25.004	0.9287	22.859	0.8794	21.600	0.8385	20.639	0.7981	19.793	0.7575
Bicubic	29.613	0.9767	25.619	0.9388	23.418	0.8944	22.050	0.8527	21.033	0.8114	20.133	0.7700
B-Spline	30.008	0.9780	25.713	0.9389	23.460	0.8933	22.089	0.8509	21.027	0.8064	20.129	0.7638
Proposed Method	31.555	0.9849	26.791	0.9544	24.355	0.9174	22.803	0.8793	21.635	0.8382	20.635	0.7950

Reconstruction results using classical interpolation method and the proposed method over noisy MR images

Table IV

Slice Thickness	0%		1%		3%		5%		7%		9%	
	PSNR (dB)	SSIM										
Nearest Neighbor	27.257	0.9636	27.371	0.9531	27.436	0.9099	27.164	0.8719	26.658	0.8436	26.033	0.8217
Bilinear	28.422	0.9674	28.491	0.9554	28.281	0.9028	27.674	0.8541	26.893	0.8167	26.064	0.7871
Bicubic	29.794	0.9768	29.821	0.9651	29.371	0.9156	28.498	0.8708	27.520	0.8367	26.561	0.8098
B-Spline	30.200	0.9782	30.201	0.9660	29.619	0.9150	28.609	0.8691	27.539	0.8345	26.522	0.8071
Proposed Method	31.602	0.9846	31.537	0.9729	30.582	0.9251	29.271	0.8837	28.046	0.8535	26.941	0.8295

Reconstruction results using classical interpolation method and the proposed method over pathologic MR images

Table V

	Nearest Neighbor	Bilinear	Bicubic	B-Spline	Proposed Method
PSNR(dB)	28.344	29.612	31.022	31.441	32.696
SSIM	0.9644	0.9687	0.9780	0.9793	0.9842

Table VI

Reconstruction results using the proposed method and NLM over non-noisy MR images

Slice Thickness	2 mm		3 mm	
	PSNR	SSIM	PSNR	SSIM
NLM	33.2143	0.9899	27.7150	0.9629
Proposed Method	31.5550	0.9849	26.7908	0.9544
Slice Thickness	4 mm		5 mm	
	PSNR	SSIM	PSNR	SSIM
NLM	24.8521	0.9239	22.9371	0.8768
Proposed Method	24.3551	0.9174	22.8025	0.8793
Slice Thickness	6 mm		7 mm	
	PSNR	SSIM	PSNR	SSIM
NLM	21.7449	0.8333	20.6463	0.7852
Proposed Method	21.6345	0.8382	20.6349	0.7950

Table VII

Reconstruction results using the proposed method and NLM over noisy MR images

Noise Percentage	0%		1%	
	PSNR	SSIM	PSNR	SSIM
NLM	33.4947	0.9901	33.2515	0.9776
Proposed Method	31.6016	0.9846	31.5370	0.9729
Noise Percentage	3%		5%	
	PSNR	SSIM	PSNR	SSIM
NLM	31.6812	0.9312	29.8899	0.8880
Proposed Method	30.5816	0.9251	29.2708	0.8837
Noise Percentage	7%		9%	
	PSNR	SSIM	PSNR	SSIM
NLM	28.3897	0.8562	27.1382	0.8312
Proposed Method	28.0463	0.8535	26.9405	0.8295

Table VIII

Computation time and peak memory of NLM and the proposed method

	Axial		Coronal		Sagittal		Average	
	T ^a (s)	P.M ^b (Mb)						
NLM	345	179	352	179	394	377	364	245
Proposed Method	124^c +41^d =165	99	118^c +37^d =155	99	151^c +47^d =198	124	131^c +42^d =173	107

^aT means running time;

^bP.M means peak memory is used;

^cTraining time;

^dreconstruction time.