

# The Clique Potential of Markov Random Field in a Random Experiment for Estimation of Noise Levels in 2D Brain MRI

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**ABSTRACT:** Effective performance of many image processing and image analysis algorithms is strongly dependent on accurate estimation of noise level. We exploit the simplicity and similarity of statistics of human anatomy among different subjects to develop new noise level estimation algorithm for magnetic resonance images of brain. Objects of the experiment are noise-free 3D brain MRI of 422 subjects. There are 21 slices for each subject. For each slice, total clique potential (TCP) of Markov random field, computed from local clique potential, is indexed by 200 different levels of noise. The sample

space is the set of TCP-noise level data of each slice. The random variable is the set of indices of noise level of TCP in each element of sample space that is closest in numerical value to TCP measured from a test MRI slice. Noise level is estimated from the mean and variance of the random variable. We also report the formulation of a generalized mathematical model describing relationship between TCP and Rician noise level in brain MRI images. Our proposal can operate in the absence of signals in the background and significantly reduce modeling errors inherent in strong parametric assumptions adopted by some of the current algorithms. © 2013 Wiley Periodicals, Inc. *Int J Imaging Syst Technol* 23, 304–313, 2013; Published online in Wiley Online Library (wileyonlinelibrary.com). DOI: 10.1002/ima.22065

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Data used in the design of the two algorithms proposed in this article were obtained from the Alzheimer's Disease Neuroimaging Initiative (ADNI) database (adni.loni.ucla.edu). As such, the investigators within the ADNI provided data but did not participate in analysis or writing of this report. A complete listing of ADNI investigators can be found at: [http://adni.loni.ucla.edu/wp-content/uploads/how\\_to\\_apply/ADNI\\_Acknowledgement\\_List.pdf](http://adni.loni.ucla.edu/wp-content/uploads/how_to_apply/ADNI_Acknowledgement_List.pdf). Accessed on September 21, 2013.

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## 1. INTRODUCTION

Noise level estimate in magnitude MRI images is required as input parameter for algorithms developed for the task of noise removal (Nowak, 1999; Basu et al., 2006; Wang and Zhou, 2006), segmentation (Zhang et al., 2001; Akselrod-Ballin et al. 2006), registration (Rohde et al., 2005), quality assessment of functional magnetic resonance images (Sendur et al.; 2005), overall quality assessment of magnetic resonance imaging systems (McVeigh et al., 1985; Tapiovaara and Wagner, 1993), and performance evaluation of noise removal algorithms (Milindkumar and Deshmukh, 2011).

We reviewed seven contributions (Henkelman, 1986; Brummer et al., 1993; Murphy et al., 1993; Chang et al., 2005; Sijbers et al., 2007; Aja-Fernandez et al., 2008; Rajan et al., 2010) in the literature on estimation of noise levels in brain MRI. Based on chronology, we

classify them into two classes: the earlier and later groups. In the earliest, referred to as double acquisition method (Murphy et al., 1993), the same image is acquired twice, aligned, and subtracted. The noise level is estimated from the standard deviation of the subtracted image. Other early contributions (Henkelman, 1986; Brummer et al., 1993; Chang et al., 2005) estimate noise from the background region where the noise is described by Rayleigh distribution. The first later contribution (Sijbers et al., 2007) estimates noise level automatically from the maximum likelihood estimate of partial histogram of noise signal in the background region. The second in same category (Aja-Fernandez et al., 2008) estimates noise from local statistics in foreground where pixel intensities contaminated by noise are described by Gaussian distribution at high signal-to-noise-ratio and Rician at low signal-to-noise ratio. The third (Rajan et al., 2010) adopts maximum likelihood estimation principle to estimate the noise level from the local variance and local skewness of pixels.

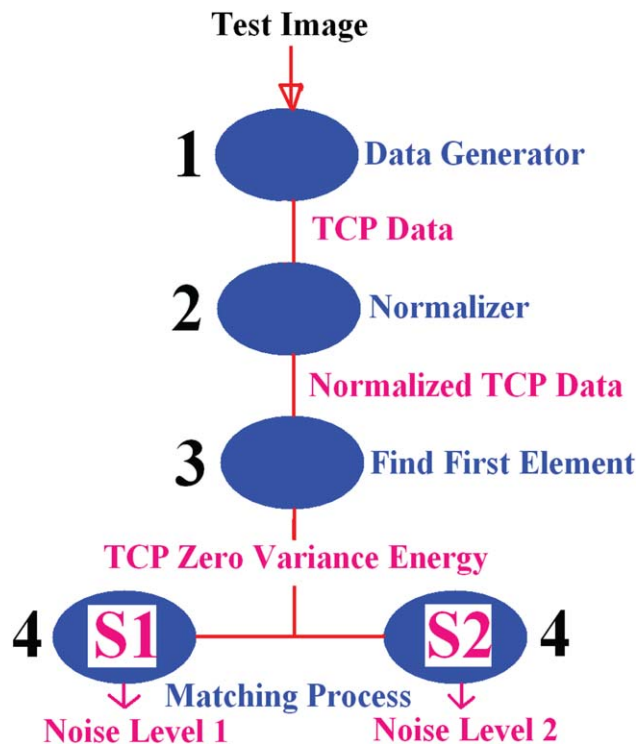
The human brain is statistically simple and geometrically similar (Zhang et al., 2001; Osadebey, 2009). It is composed of only three major brain structures: white matter (WM), grey matter (GM), and the ventricular system (VS), each can be distinguished by its clearly defined range of voxel intensity values (Ashburner and Friston, 2003). Magnetic resonance imaging system reveals strong spatial, structural, and voxel statistical similarities among corresponding inter subject MRI slices. These similarity features are exploited in the design of single-subject MRI-based brain atlases (Evans et al., 1994; Mazziotta et al., 2001), which are widely acknowledged powerful tools in the analysis of brain images (Doan et al., 2010). We exploit this principle further by inferring that, across subjects, profiles describing variation of MRI slice voxel intensities with different levels of noise degradation will exhibit similar characteristics.

We regard each slice in a 3D MRI volume as similar “coin” or “dice” in a random experiment. Each “throw” or “roll” is the degradation of a noise-free MRI slice by 200 different levels of Rician noise. For each noise level, there is a corresponding outcome, the total clique potential of Markov random field. A slice is “thrown” 21 times (the number of slices in an individual subject’s 3D MRI volume) and the experiment is repeated 422 times (total number of subjects). From the random experiment, we derive two separate sample spaces. The first is 2D of size  $(422 \times 21 = 8862) \times 200$ . Each outcome of the sample space is TCP energy indexed by noise level for each slice. The second, derived from the mean of the first sample space, is 1D of size  $1 \times 200$ .

Our proposed algorithm is summarized in the schematic diagram displayed in Figure 1. Given a test image, TCP-noise level data is generated by degradation with increasing levels of noise, starting from zero noise level, up to  $\sigma=200$ . The data is normalized and the TCP zero variance energy is determined as the first element of the TCP-noise level data which corresponds to zero noise level. This prime energy value is numerically matched separately to each outcome in the two sample spaces resulting in two noise estimation results. The first noise estimate is determined from the mean and variance of the random variable generated from indices of each TCP-noise level data in the 2D sample space that is closest in numerical value to TCP-noise level data measured from the test image. The second noise level estimate is the unique index of MRF energy in the 1D sample space that is closest in numerical value to its counterpart in the test image data.

## II. MATERIALS AND METHODS

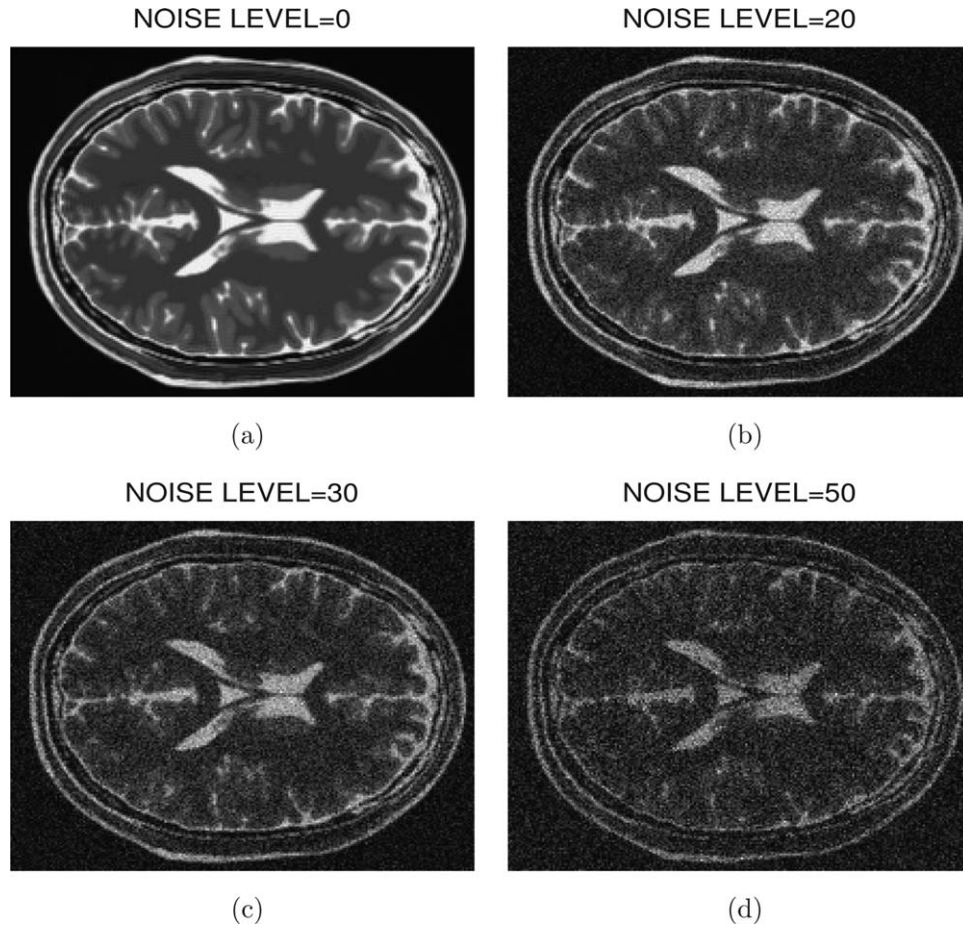
**A. Data for the Random Experiment.** The objects of the random experiment were original, noise-free, 3 mm  $T_2$  weighted axial 3D brain MRI sourced from the Alzheimer’s Disease Neuroimaging



**Figure 1.** The four steps in the estimation of noise level in 2D brain MRI. In step 1, TCP-noise level data of the test image is generated. The data is normalized in step 2 and the first element (TCP zero variance energy) of the normalized data is extracted in step 3. In the estimation of the noise (step 4), the TCP zero variance energy is separately matched to each of the 1D and 2D sample spaces, S1 and S2, respectively. [Color figure can be viewed in the online issue, which is available at [wileyonlinelibrary.com](http://wileyonlinelibrary.com).]

Initiative (ADNI) database ([www.adni.loni.ucla.edu](http://www.adni.loni.ucla.edu)). We chose as much as 422 subjects to satisfy the requirements of the central limit theorem (Leon-Garcia, 2008; Norman and Streiner, 2008; Walker and Shostak, 2010) and optimally extract similarities in the voxel statistics of inter subject brain MRI. The range, mean, and standard deviation of the subjects’ age were 71–87, 75, and 10, respectively.

ADNI was launched in 2003 by the National Institute on Aging (NIA), the National Institute of Biomedical Imaging and Bioengineering (NIBIB), the Food and Drug Administration (FDA), private pharmaceutical companies, and non-profit organizations, as 60 million dollars, 5-year public private partnership. The primary goal of ADNI has been to test whether serial magnetic resonance imaging (MRI), positron emission tomography (PET), other biological markers, and clinical and neuropsychological assessment can be combined to measure the progression of mild cognitive impairment (MCI) and early Alzheimer’s disease (AD). Determination of sensitive and specific markers of very early AD progression is intended to aid researchers and clinicians to develop new treatments and monitor their effectiveness, as well as lessen the time and cost of clinical trials. The principal investigator of this initiative is Michael W. Weiner, MD, VA Medical Center and University of California in San Francisco. ADNI is the result of efforts of many co-investigators from a broad range of academic institutions and private corporations, and subjects have been recruited from over 50 sites across the US and Canada. The initial goal of ADNI was to recruit 800 adults (ages from 55 to 90) to participate in the research, approximately 200 cognitively normal older individuals to be followed for 3 years, 400



**Figure 2.** An MRI slice image at various levels of degradation by Rician noise (a)  $\sigma=0$ , (b)  $\sigma=20$ , (c)  $\sigma=30$ , and (d)  $\sigma=50$ .

people with MCI to be followed for 3 years, and 200 people with early AD to be followed for 2 years. For up-to-date information, the reader is referred to [www.adni-info.org](http://www.adni-info.org).

**B. Single-Layered Markov Random Field.** An image  $I$  is modeled as a single-layered Markov random field. Unlike the classical Markov random field theory (Geman and Geman, 1984), the observed image is the only physical system under consideration and there is no reference to a prior model. The Markov random field energy  $U$  is dependent on voxel configuration  $f$ . The voxel configuration is quantized into local cliques  $c$  in a clique system  $S$  that describes spatial coherence or clusters of similar voxels, so that the single-layered Markov random field energy is the sum of local clique potentials  $V_c$ :

$$U(f) = \sum_{c \in S} V_c(f) \quad (1)$$

For the clique system, we adopt second order neighborhood of size two with neighboring voxels indexed as  $(i, i')$ . In this system, the Markov random field energy is the sum of potential function contributions from single site and pair-site cliques (Li, 2009):

$$\hat{U}(f) = \sum_{i \in S} \alpha_1 V_1(f_i) + \sum_{i \in S} \sum_{i' \in N} \alpha_2 V_2(f_i, f_{i'}) \equiv E_a + E_b \quad (2)$$

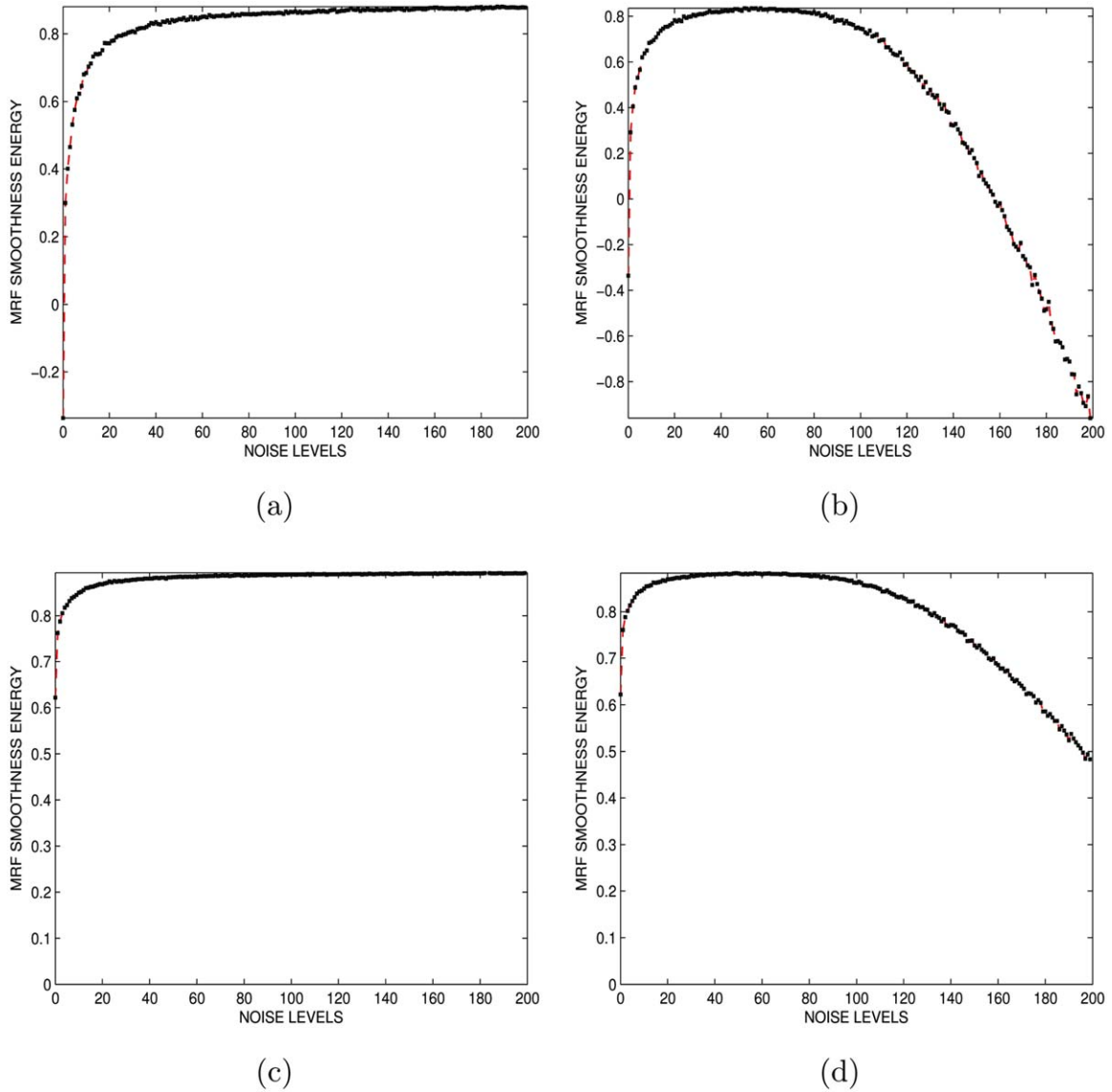
where  $\alpha_1, \alpha_2$  are interaction coefficients. The first term  $E_a$  referred to as the data term is the contribution from comparing each voxel

intensity level to itself. The second term  $E_b$ , the smoothness term, is contribution from comparing each voxel intensity to its neighbors. At each local clique, the contribution of each neighboring voxel to the clique potential energy is determined according to the expression:

$$V_c(I(x, y)) = \begin{cases} \xi_r & \text{if } I(i) = I(i') \\ \xi_p & \text{otherwise} \end{cases} \quad (3)$$

where  $\xi_r$  is the reward and  $\xi_p$  the penalty for conformity ( $f_i = f_{i'}$ ) and violations ( $f_i \neq f_{i'}$ ) of the smoothness constraints, respectively.

Noise changes the pattern of arrangement of pixels in the image as demonstrated in Figure 2. There is gradual change in pixel configurations for noise levels increasing from  $\sigma=0$  in Figure 2a to  $\sigma=20$ ,  $\sigma=30$ , and  $\sigma=50$  in Figures 2b–2d, respectively. The different noise levels result in change in the strength of the clusters within the image. Thus, different noise levels  $\sigma$  give rise to different pixel configurations and their corresponding levels of energy  $\hat{U}_\sigma(f)$ . The data term is set to zero because it is a constant for different levels of noise. This renders the choice of values assigned to  $\alpha_1$  and  $\alpha_2$  in Eq. (2) irrelevant, hence for computational convenience, we set  $\alpha_1$  and  $\alpha_2$  to an arbitrary value of 1. For different image dimensions, we define total clique potential per pixel  $U$ :



**Figure 3.** TCP energy-noise level curves associated with different image formats and combinations of conformity  $\xi_r$  and violations  $\xi_p$  of the smoothness constraints, (a) double precision and  $(\xi_r = -8, \xi_p = 1)$ , (b) 8-bit integer and  $(\xi_r = -8, \xi_p = 1)$ , (c) double precision and  $(\xi_r = -1, \xi_p = 1)$ , and (d) 8-bit integer and  $(\xi_r = -1, \xi_p = 1)$ . [Color figure can be viewed in the online issue, which is available at [wileyonlinelibrary.com](http://wileyonlinelibrary.com).]

$$U(f) = \frac{\hat{U}}{D} = \sum_{i \in S} \sum_{i' \in N} V_2(f_i, f_{i'}) \quad (4)$$

normalized by  $D$ , the product of row and column dimensions of the image.

It is well known that, in the presence of noise, the pixel intensities in the foreground of magnitude MRI follow a Rician distribution (Gravel et al., 2004):

$$P(M) = \frac{M}{\sigma^2} e^{-\frac{M^2 + A^2}{2\sigma^2}} I_0\left(\frac{AM}{\sigma^2}\right) \varepsilon(M) \quad (5)$$

where  $A$  is the image pixel intensity in the absence of noise,  $M$  is the measured pixel intensity,  $I_0$  is the modified zeroth order Bessel function of the first kind, and  $\sigma$  denotes the standard deviation of the Gaussian noise in the real and the imaginary images. In the

background, where there is no image signal, the measured noise intensity is described by Rayleigh distribution.

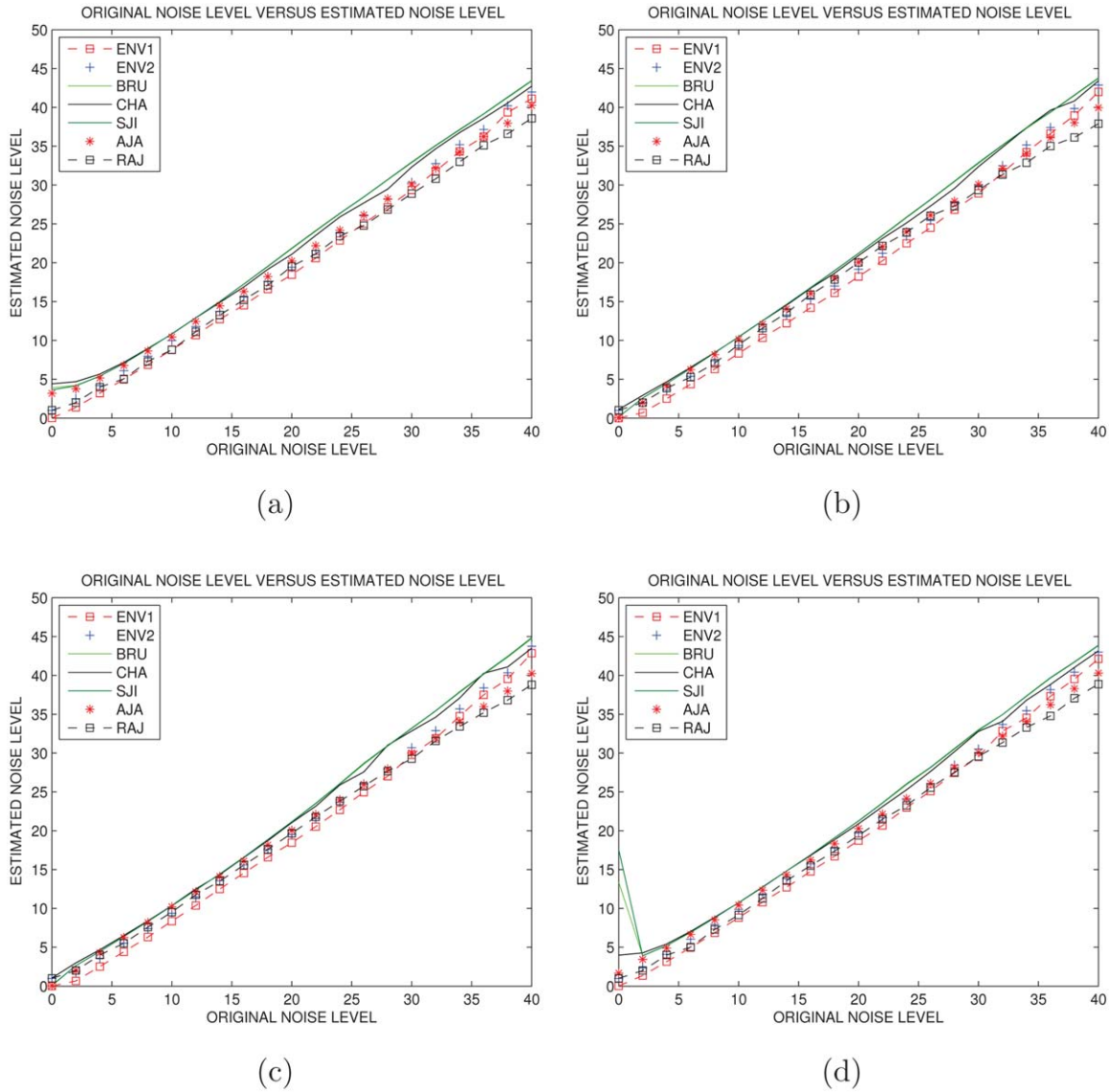
$$P(M) = \frac{M}{\sigma^2} e^{-\frac{M^2}{2\sigma^2}} \quad (6)$$

With the exception of Murphy et al. (1993), most of current algorithms estimate noise from the probability distributions expressed in Eqs. (5) and (6). In our approach, we relate the potential function  $U$  expressed in Eq. (2) to the signal-to-noise ratio  $\left(\frac{A}{\sigma^2}\right)$ , a variable on the right hand side of Eq. (5).

$$\sum_{i \in S} V_{1,\sigma}(f_i) + \sum_{i \in S} \sum_{i' \in N} V_{2,\sigma}(f_i, f_{i'}) \equiv \left(\frac{A}{\sigma_n^2}\right) \quad (7)$$

It is noteworthy that the choice of  $\xi$  assigned to each clique is a strong determinant of the relationship between  $U$  and  $\sigma$  in Eq. (7). In





**Figure 5.** Comparative performance evaluation. The plots of the estimated noise levels  $\hat{\sigma}$  versus original noise levels  $\sigma$  for four different MRI acquisitions: (a) T2, (b) T1, (c) PD, and (d) FLAIR. [Color figure can be viewed in the online issue, which is available at [wileyonlinelibrary.com](http://wileyonlinelibrary.com).]

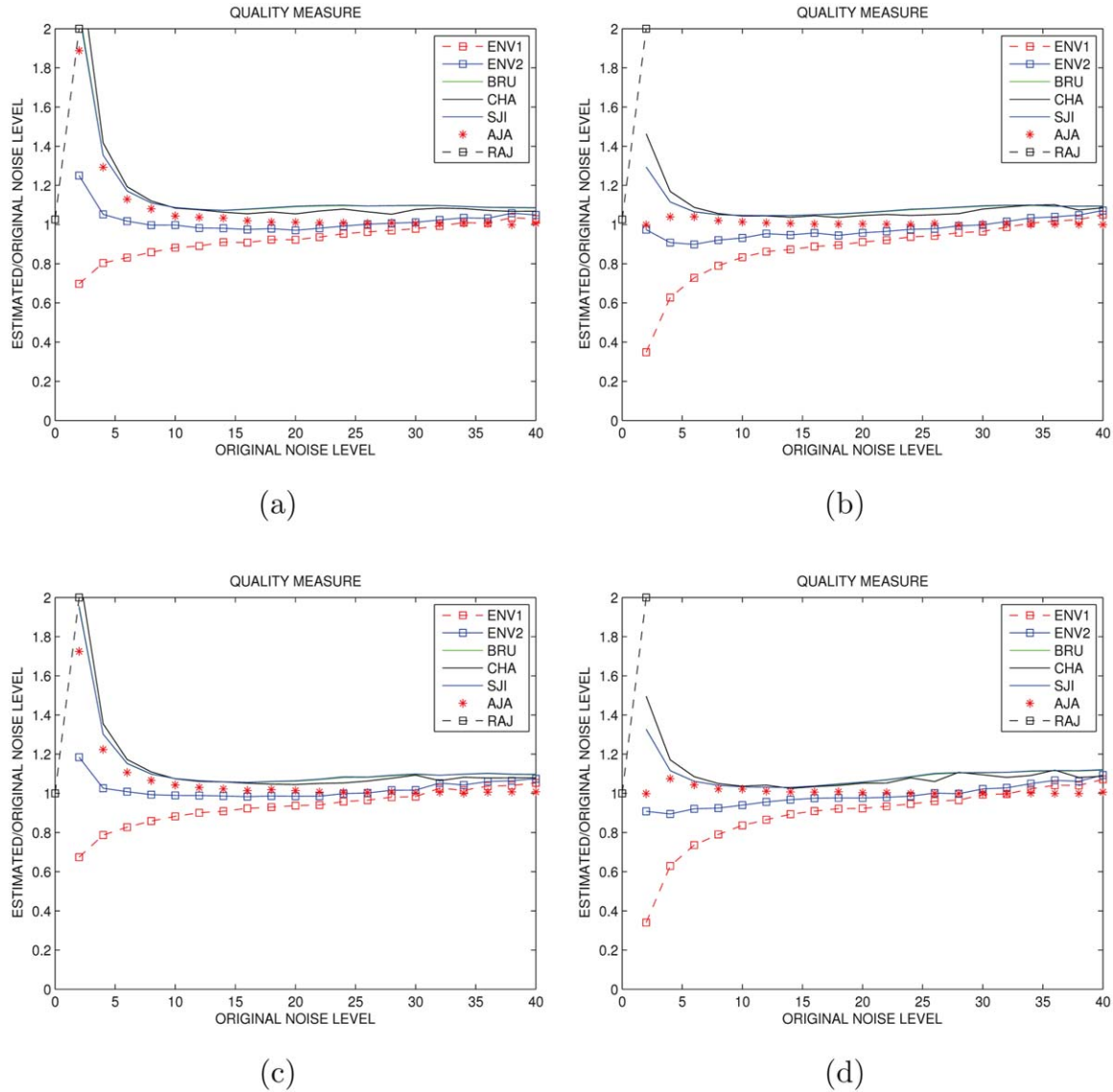
**E. Random Variable.** A random variable  $X$  is defined as a function that assigns a real number  $X(\omega)$ ,  $\omega \in \Omega$ , to each outcome  $\omega$  in the sample space  $\Omega$  of a random experiment (Leon-Garcia, 2008). Different random variables can be defined based on this random experiment. For example, there are a total of  $T = ZL$  MRF energies indexed by  $\{L : L < Z\}$  levels of noise. If  $Z$  is large enough such that  $Z \gg L$ , then  $T$  MRF energies will be randomly distributed into each noise level. Thus, the set of all possible MRF energies  $\{\hat{U}_{1,l}, \hat{U}_{2,l}, \hat{U}_{3,l}, \dots, \hat{U}_{Z,l}\}$ , consisting of  $Z$  number of elements, that are indexed by a particular noise level  $l$  is a random variable. The random variable of interest to our proposal is defined as the set containing the indices  $X_z(\omega) = \{l_1, l_2, l_3, \dots, l_Z\}$ ,  $l \in [1, L]$  of TCP energy  $U_{z,l}$  in each outcome  $\omega_z = \{U_{z,l} : 1 \leq z \leq Z, 1 \leq l \leq L\}$  of the sample space that is closest in value to the TCP energy measured from a test TCP image:

$$\{X_z(\omega)\}_{X_z \in [1, L]} = \arg \min_{l \in [1, L]} : \{l | \forall z : \|E_l - \{U_{z,l}\}\| \} \quad (17)$$

**F. Estimation of Noise Level.** The test image is corrupted with increasing level  $\{l : 0 \leq l \leq 200\}$  of Rician noise, starting from the first noise level  $\sigma=0$ . For each noise level, the TCP energy  $E_{t,l}$  is computed according to Eq. (4) to obtain a  $1 \times 200$  TCP energy-noise level data. The data is normalized by dividing each element with the maximum TCP energy. The normalized TCP energy  $\hat{E}_{t,l}$  measured from the test image is the first element  $l = 1$  of the normalized TCP energy-noise level data.

The normalized TCP energy measured from the test data is matched separately to each TCP energy-noise level data of an MRI slice in the 1D and 2D sample spaces. In both cases, we find the indices of the TCP energy in each TCP energy-noise level data that is closest in value to the normalized TCP energy measured from the test data.

The first matching generates random variable consisting of sequence of indices  $X_z(\omega) = \{l_1, l_2, l_3, \dots, l_Z\}$  of same number of elements  $Z$  as the MRI slice images in the sample space giving a total of  $Z$  number of  $Q$  unique discrete indices  $\theta_z \in [0, L]$  each with



**Figure 6.** Comparative performance evaluation. The plots of the quality measures for four different MRI acquisitions: (a) T2, (b) T1, (c) PD, and (d) FLAIR. [Color figure can be viewed in the online issue, which is available at [wileyonlinelibrary.com](http://wileyonlinelibrary.com).]

frequency of occurrence  $\lambda_q$ . The estimated level of noise  $ENV1$  is the mean of the probability distribution.

$$ENV1 = \theta_1 \left( \frac{\lambda_1}{Z} \right) + \theta_2 \left( \frac{\lambda_2}{Z} \right) + \theta_3 \left( \frac{\lambda_3}{Z} \right) \dots + \theta_Z \left( \frac{\lambda_Q}{D} \right) - 1 \quad (18)$$

The deviation from the mean is:

$$ENV1_{std} = \sqrt{\sum_{q=1}^Q (\theta_q - ENV1)^2 \left( \frac{\lambda_q}{Z} \right)} \quad (19)$$

so that the estimate for the upper and lower limits of the noise level is  $ENV1 + ENV1_{std}$  and  $ENV1 - ENV1_{std}$ , respectively.

The second matching generates only a single and unique index number  $ENV2$  because the matching is with a single MRI energy-noise level data.

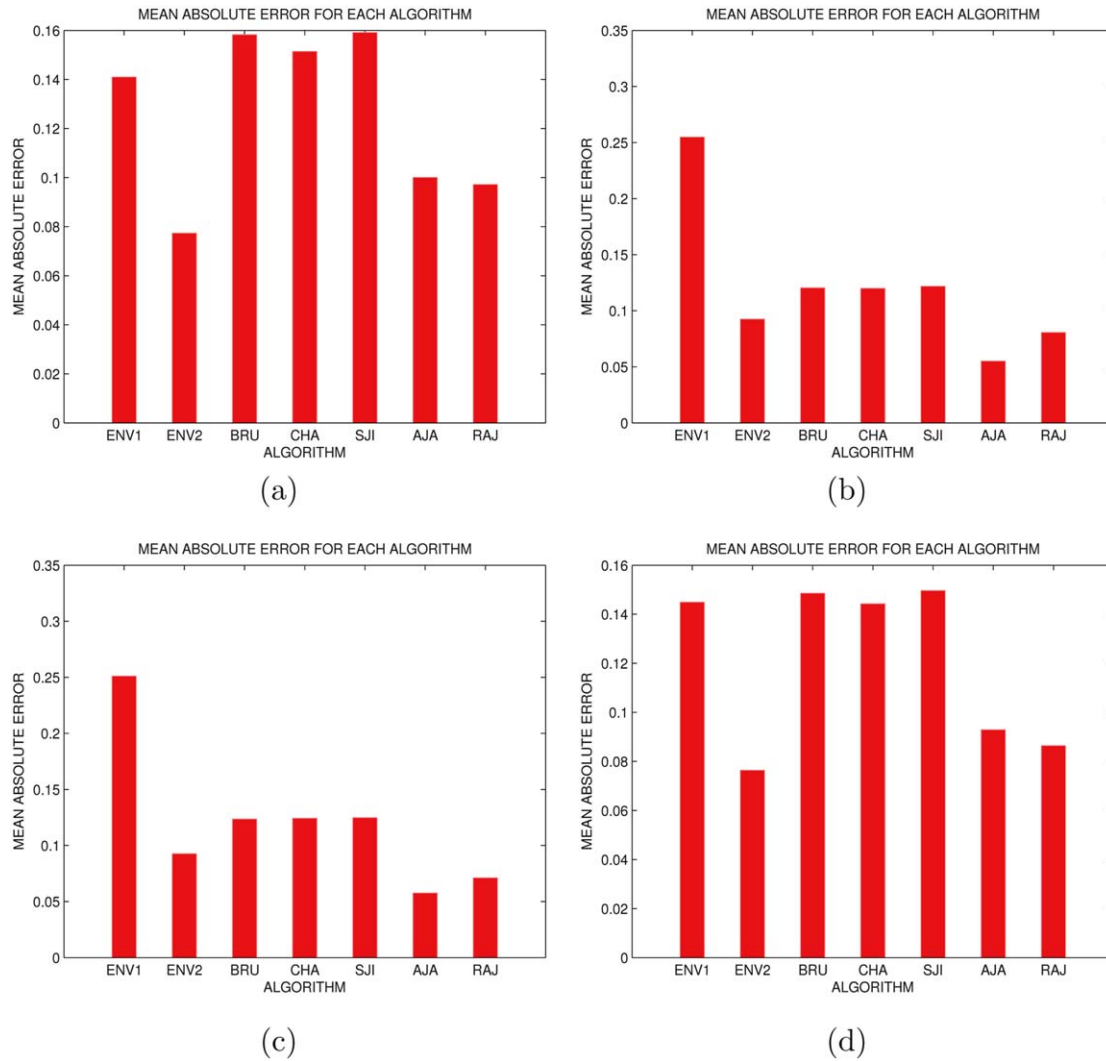
$$ENV2 = \min_{l \in [1, L]} \left| \hat{E}_{t,1} - \kappa_l \right| - 1 \quad (20)$$

The subtraction of unity in both Eqs. (18) and (20) is rescaling of the indices to account for index of noise level associated with Rician noise of zero variance.

The accuracy of the estimated noise level is strongly dependent on  $\hat{E}_t$ . To reduce error our algorithm monitors the smoothness of the TCP energy-noise level data generated from the test image by computing its coefficient of variation  $\{C_v | C_v \in \mathbf{Z}\}$ , defined as the ratio of the standard deviation  $\sigma$  to its mean value  $\mu$ , rounded to the nearest integer (Rosner, 2006).

$$C_v = \left( \frac{\sigma}{\mu} \right) \quad (21)$$

Based on trials of many test images, we set threshold value of  $T_v = 23$  for  $C_v$ . Below this threshold, the noise is estimated using the



**Figure 7.** Comparative performance evaluation. The plots of the mean absolute error for four different MRI acquisitions: (a) T2, (b) T1, (c) PD, and (d) FLAIR. [Color figure can be viewed in the online issue, which is available at [wileyonlinelibrary.com](http://wileyonlinelibrary.com).]

first element  $\hat{E}_{t,1}$  of the normalized test data as input parameter, and above it, the data is first fitted with a power model followed by updating  $\hat{E}_t$  according to the following operation

$$E_{t,1} \rightarrow E_{t,N}, \quad N = (T_v - C_v) \Delta \quad (22)$$

where  $N$ , an integer, is the index of the fitted model that is now assumed as the updated MRF energy computed from the test image and  $\Delta$  is a user-defined correction factor, in the range  $(0 \leq \Delta \leq 1)$ , that takes into consideration the complexity of the features in the test image. In our case, we set  $\Delta=1$  because each MRI slices of individual subjects in the database were free of noise and had undergone intensity correction.

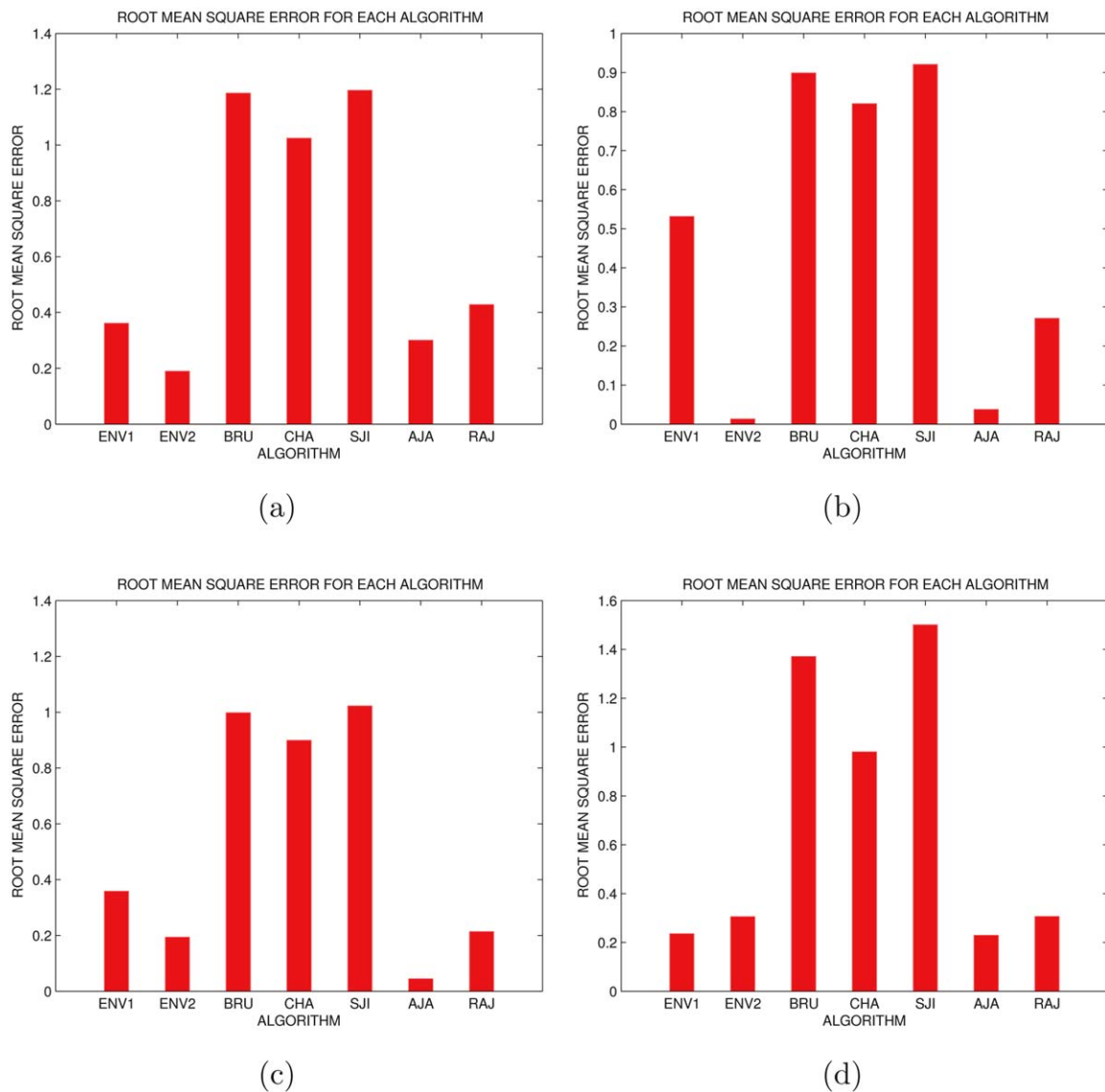
### III. RESULTS

**A. Data for Evaluation.** The test data are four different acquisition modes of real MRI data made available by NeuroRx research Inc., a Montreal based clinical research organization. They are  $T_1$  relaxation time,  $T_2$  relaxation time, and proton density (PD) as well as fluid attenuated inversion recovery (FLAIR). Each acquisition consists of thirty eight 3 mm axial MRI slices.

**B. Comparative Performance Evaluation Results.** The two methods of estimating noise in our proposed algorithm were denoted **ENV1** and **ENV2**. They were compared alongside five current algorithms. Brummer et al. (1993) **BRU**, Chang et al. (2005) **CHA**, Sijbers et al. 2007 **SJI**, Aja-Fernandez et al. (2008) **AJA**, and Rajan et al. (2010) **RAJ**. Each slice in the real MRI data was modified to follow Rician noise distribution ranging from  $(0 \leq \sigma \leq 40)$ . The artificially induced noise level was based on assumption that the magnitude MRI was acquired from real and imaginary data having equal standard deviations  $\sigma$ . For each original noise level  $\sigma$ , the estimate level of noise  $\hat{\sigma}$  from an algorithm is determined from the mean value of noise level estimated for all individual slices in the volume data. The algorithms were compared based on four evaluation parameters. The plot of estimated noise level versus original noise level is displayed in Figure 5. The corresponding plots for quality measure, mean absolute error, and root mean square error are displayed in Figures 6–8, respectively.

### IV. DISCUSSION

The algorithms were evaluated on images with both background and foreground signals. Three algorithms **BRU**, **CHA**, and **SJI**



**Figure 8.** Comparative performance evaluation. The plots of the root mean square error for four different MRI acquisitions: (a) T2, (b) T1, (c) PD, and (d) FLAIR. [Color figure can be viewed in the online issue, which is available at [wileyonlinelibrary.com](http://wileyonlinelibrary.com).]

are designed to operate on images for which the background signal is clearly segmented from the foreground, and are therefore not optimal in performance in the presence of only foreground or limited background signals that are typical outputs of modern MRI scanners and MRI images with small field of view Rajan et al. (2010). Our proposal can be ranked alongside **AJA** and **RAJ** which can operate with and without background signals. Another state-of-the-art algorithm which can operate in the presence or absence of background was proposed by Coupe et al. (2010), but it is designed only for 3D MRI volume data. Our proposal can be adapted to measure noise level in 3D volume data by measuring noise level in each slice and averaging by the total number of slices in the volume. The time it takes to process an MRI slice is dependent on the mode adopted for estimation of noise level. On a computer having 3GB RAM and 1.65 GHz processor, the second approach **ENV2** can estimate noise level in an MRI slice in less than 10 s compared to **ENV1** that takes less than 30 s because it makes reference to a database.

The data points in the plots of estimated noise levels versus original noise levels in Figure 5 and quality measure in Figure 6 are strongly clustered at significant levels of noise. This indicates that all the algorithms under consideration are comparable to each other and that our proposal exhibit state-of-the-art characteristics of current algorithms. However, the distinction in their performances can be deduced from the plots of mean absolute errors and root mean square errors displayed in Figures 7 and 8, respectively.

The second approach in our proposal **ENV2** demonstrates superior performance in terms of mean absolute error ( $\approx 0.08$ ) on T2 (Fig. 7a) and FLAIR (Fig. 7d) MRI images. In the same evaluation, parameter **AJA** was the best algorithm with ( $\approx 0.07$ ) for T1 (Fig. 7b) and PD (Fig. 7c) images. It is closely followed by **RAJ** and our proposal **ENV2**.

Again our **ENV2** stands out clearly as the best algorithm based on root mean square error for T1 (Fig. 8b), ( $\approx 0.02$ ) and T2 (Fig. 8a), ( $\approx 0.2$ ) images. Closely behind is our first proposal **ENV1**, **AJA**, and **RAJ**. The algorithm **AJA** is the best for PD images

( $\approx 0.2$ ) closely followed by our two proposals **ENV1** and **ENV2** and **RAJ**. As can be seen in (Fig. 8d) for FLAIR images, the first approach in our proposal **ENV1** and **AJA** show the same level of best performance ( $\approx 0.2$ ) and closely followed by our second proposal and **RAJ**.

## V. CONCLUSION

We exploit the simplicity and similarity of statistics of human anatomy for different subjects in a random experiment and hereby propose a two-in-one application-specific algorithm that adopts the total clique potential of Markov random field energy as metric for estimation of noise variance in brain MRI. In the first approach, MRF energy is a variable in a random experiment. The second formulates the MRF energy-noise level relationship as a mathematical model. Both approaches are fast, accurate, and efficient in the estimation of noise variance. It is invariant to the presence or absence of background features in an image and is potentially immune to the modeling errors inherent in some of the current state-of-the-art algorithms.

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