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Modified distance regularized level set evolution for brain ventricles segmentation



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Abstract

Neurodegenerative disorders are commonly characterized by atrophy of the brain which is caused by neuronal loss. Ventricles are one of the prominent structures in the brain; their shape changes, due to their content, the cerebrospinal fluid. Analyzing the morphological changes of ventricles, aids in the diagnosis of atrophy, for which the region of interest needs to be separated from the background. This study presents a modified distance regularized level set evolution segmentation method, incorporating regional intensity information. The proposed method is implemented for segmenting ventricles from brain images for normal and atrophy subjects of magnetic resonance imaging and computed tomography images. Results of the proposed method were compared with ground truth images and produced sensitivity in the range of 65%–90%, specificity in the range of 98%–99%, and accuracy in the range of 95%–98%. Peak signal to noise ratio and structural similarity index were also used as performance measures for determining segmentation accuracy: 95% and 0.95, respectively. The parameters of level set formulation vary for different datasets. An optimization procedure was followed to fine tune parameters. The proposed method was found to be efficient and robust against noisy images. The proposed method is adaptive and multimodal.

Keywords: Ventricles, Atrophy, Segmentation, Level set, Diagnosis

Introduction

The brain gets affected by multiple factors; degenerative disorders are one of them. Degenerative disorders could be a result of environment changes, food habits, work depression, hereditary, traumatic injury or caused from the effect of other neurological disorders. Degenerative disorders in the brain render the subject unable to perform their activities of daily living such as communication, movement, balancing their body while walking and sometimes vital functions such as respiration and functioning of the heart. Atrophy in the brain or cerebral atrophy is a symptom or a form or stage which is caused by many diseases. In atrophy, the brain starts to shrink because of neuronal loss [1, 2]. Atrophy affects all structures of the brain, but in different ratios. Therefore, for the diagnosis of atrophy, certain regions or structures of

the brain are preferred by experts. Atrophy can be detected by several ways: mini mental state exam, electroencephalography, using imaging modalities such as computed tomography (CT), magnetic resonance imaging (MRI), positron emission tomography (PET) and single photon emission computed tomography (SPECT), measuring cerebrospinal fluid (CSF) pressure and biopsy of the brain tissues. Imaging modalities give an advantage of a clear view of the brain and helps in giving the location and structure which has more atrophy in terms of quantification [3, 4]. This will aid the physician in early detection.

The ventricular system is one of the prominent structures in the brain, located in the mid-regions of the brain, filled with CSF [5]. At different stages of degenerative disorders, atrophy in particular, the volume of the CSF tends to increase due the shrinkage or loss of neurons [5, 6]. As ventricles are filled with CSF, their shape and size also keep changing due to the changes occurring in the CSF and the subsequent progression of

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the disease [1, 4]. Thus, brain ventricles serve as a major landmark or biomarker for detecting atrophy. Anandh et al. [7] attempted a ventricular enlargement study to differentiate between mild cognitive impairment (MCI) and Alzheimer's disease. This group employed shape descriptors to study the shape differences of ventricles and used it for the classification purpose. Wang et al. [8] used the shape of the brain ventricles for improving the classification accuracy of Alzheimer's disease from normal subjects and other diseases using shape descriptors. Bader et al. [9] performed volume estimation of CSF and ventricles, to verify correlations between CSF oscillations and ventricular dilatation, and how their variations are indicated in brain disorders. An experimental based structural mapping was employed by Carmichael et al. [10] for distinguishing dementia and MCI in a large community.

For landmark based diagnostic systems such as morphometric and volumetric studies, the region of interest needs to be either highlighted or separated from the background [11–13]. Hence object segmentation is a pre-processing step in a diagnostic system. The manual segmentation process is labor intensive, time consuming, and could be biased, and its accuracy is based on the individual's training skills. Thus, computer-based segmentation procedures are preferred by experts to minimize manual errors. It is important that these computerized algorithms are accurate, rapid, cost effective, automatic, and robust [13].

There are different types of segmentation techniques that have been reported in the literature: thresholdingbased techniques, region growing and splitting algorithms, clustering techniques, atlas-based models, deformable models, and neural networks and classifiers [14]. Baghdadi et al. [15] came up with the Bayesian Generalized Fast Marching algorithm for segmenting brain structures including the ventricles. Mishra et al. [14] presented a hybrid segmentation procedure using the watershed algorithm and distance regularized level set evolution (DRLSE) algorithm for segmenting gray matter, white matter and CSF from MR images of the brain. Xia et al. [12] developed a knowledge-driven algorithm for automatic segmentation of the ventricles from MR images. Narr et al. [16] reported a 3D mapping of temporo-limbic regions and ventricles for schizophrenic subjects. Anandh et al. [17] incorporated Tukey's biweight edge indicator into a level set formulation to segment the ventricles in Alzheimer's subjects and the method was shown to be clinically significant. Angelini et al. [18] implemented a four phase, 3D active contour model on 3D images for partitioning brain MR images. This group demonstrated that, the proposed framework was efficient in segmenting the regions of interest and flexible with the set of features. Aloui and Naceur [19]

proposed an approach based on level set method for delineating tumor, followed by mesh simplification to separate them out.

Even though there are many techniques available, the technique chosen varies between organs, region of interest, depending on the disease, subject's age, imaging modality (X-ray, CT, MRI, PET, SPECT), method of visualization, features to be used from the segmented image [1, 2]. Hence adaptive and multi-modal segmentation algorithms need to be developed [19]. This study employs a modified DRLSE with optimized parameters for segmenting ventricles from brain images [20, 21]. The objective of this study was to segment the ventricles based on a modified level set technique to improve segmentation accuracy and diagnosis involving ventricles.

Methods

The methods section is divided into two parts: optimization of parameters and main segmentation framework.

Level set formulation

The level set method was first introduced by Stanley Osher and James Sethian in the 1980s [22]. It is based on the active contour model concept, where the curves are represented as zero level set, with a function of higher dimension, which is called as the level set function (LSF) [22, 23]. The curve evolves, depending on certain energy functions from the image, which are derived by partial differential equations. The energy functions give the speed at which the curve evolves and result in a smoothing process. The stopping point of the curve evolution is the boundary of the object to be segmented [3, 24]. The extraction of the curve is dependent on the initial parameters chosen by the user. For this reason, LSF's performance relies more on the initialization of the parameters. There have been different re-initialization procedures, regularization procedures reported in literature for making LSF efficient and accurate [25, 26]. It can represent shapes having complex topology [27, 28]. The LSF is defined as follows: parametric contour is given by $C_1(s,t): [0,t]\times [0,\infty] \rightarrow \Re^2$ which defines parameterization of a point in the contour [22, 23, 26]. The curve evolution is defined by the following equation

$$\frac{\partial C_1(s,t)}{\partial t} = \mathcal{F}N\tag{1}$$

 C_1 is curve, \mathcal{F} is speed function that controls the motion of the contour, N is inward normal vector to the contour. The level set surface needs to remain with the same smoothness level; hence initialization has to be given at regular intervals - this process is called as reinitialization [29, 30].

$$\phi_t = sign(\phi_0)(1 - |\nabla \phi_0|) \tag{2}$$

 ∇ is the gradient operator of the image and ϕ is the LSF that needs to be re-initialized.

Optimization of parameters

Even though the level set method has many advantages, it has a drawback in the form of re-initialization; which leads to increased complexity in terms of user intervention and execution time is more. Therefore, Li et al. [21, 22] proposed DRLSE method which eliminated the need for re-initialization process. The set of input parameters used in their study were not able to segment the brain ventricles correctly for the dataset used in this study. Therefore, a preliminary study of optimizing the parameters was implemented for input images. optimization procedure was followed for all the images (n = 80 subjects): 20 atrophy CT subjects, 20 atrophy MRI subjects, 20 normal CT subjects, and 20 normal MRI subjects. In selective optimization images were included randomly from the study groups (2 atrophy and 3 normal subjects from CT dataset, 3 atrophy and 2 normal subjects from MRI dataset). The parameters which were tuned were alpha (α), lambda (λ), epsilon (ϵ), time step (Δt), edge kernel (Gaussian, Average, Median), kernel size and sigma value; whose significance will be explained later. As given in the algorithm the value which produces highest accuracy value when compared with ground truth image is considered to be the tuned value for the respective parameter. This is explained in the results section with a sample table (Table 1). The tuned parameters are common for both the datasets used in this study; CT and MR images as the optimization procedure was carried out commonly for both the datasets (mingling images from CT and MR, making it into a single group and then carrying out optimization procedure).

There are many optimization techniques available: linear, nonlinear, grid search, Bayesian, random search, and gradient based to make this parameter tuning generalized. Generalization means, parameter optimization for the segmentation framework such that, it performs well on all the datasets - irrespective of the geographical region, age and gender of the subject, imaging properties and imaging modality (The optimization carried out in

this paper is specific for the datasets used for this study). However, this involves the appropriate selection of the optimization technique based on the data and dimension of the parameters, training and testing of the parameters, cross validation, etc. This will make the selection of the parameter computationally complex. The objective of this study was to improve the DRLSE method; the proposed modification was made to make it suitable for segmenting the input images. Moreover, the parameters used in the original DRLSE study for their synthetic data were found to be unsuitable for the input data used in this study, thereby leading to failure of the segmentation procedure or improper segmentation. Therefore, this study used a simple optimization procedure for selecting suitable values for the parameters as shown in Fig. 1. If we need to go for a generalization of the parameter selection that will make the entire algorithm complex, as the segmentation algorithm itself takes up to 2 min for execution. The algorithm is as follows (Fig. 1). The tuned parameter values are: alpha-2, lambda-1, epsilon-2, time step-1.5, edge kernel-Gaussian, sigma for Gaussian kernel-2.5, size of the kernel-15.

DRLSE

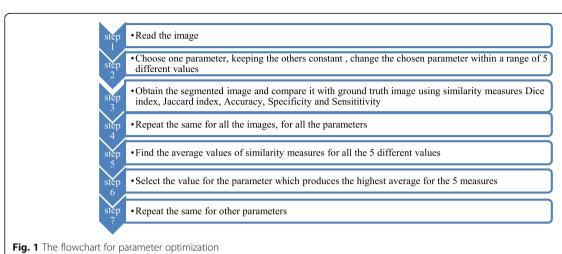
Li et al. [20, 21] proposed the DRLSE method to overcome the trouble of re-initialization. In DRLSE the surface regularity of the LSF is intrinsically maintained by a regularizing term. Hence with the surface fitting energy term, there will be a regularization term in the LSF [31]. The initial LSF, gradient operator and signed distance function are the same as the conventional level set method and have been adapted in DRLSE. Due to space constraint only; the relevant equations are given here. The remaining equations and their derivations can be referred from refs. [20, 21, 31]. The energy term is given as follows

$$\varepsilon_{\varepsilon}(\varnothing) = \mu \int_{\Omega} p \mid \nabla \varnothing \mid dx + \lambda \int_{\Omega} g \delta_{\varepsilon}(\varnothing) \mid \nabla \varnothing \mid dx + \alpha \int_{\Omega} g H_{\varepsilon}(-\varnothing) dx$$
(3)

Where, $\varepsilon_{\varepsilon}$ (\varnothing) is energy functional, ε is a constant, the first term is level set regularization term and $\mu > 0$ is a constant, p is potential function, $\lambda > 0$ coefficient of the area term: the second term in the equation, α is coefficient of weighted area term, it should be positive so that

Table 1 Average and SD values of similarity measures between ground truth and segmented results (n = 40)

DI	JI	Sensitivity	Specificity	Accuracy
0.55 ± 0.0148	0.38 ± 0.0112	87.8 ± 0.61	96.4 ± 0.72	96.2 ± 0.62
0.53 ± 0.0217	0.41 ± 0.0254	88.6 ± 0.45	96.5 ± 0.5	96.3 ± 0.62
0.52 ± 0.0356	0.39 ± 0.034	90.4 ± 0.78	96.5 ± 0.52	96.3 ± 0.72
0.59 ± 0.0294	0.46 ± 0.029	90.6 ± 0.86	96.59 ± 0.73	96.39 ± 0.54
0.56 ± 0.0378	0.39 ± 0.034	90 ± 0.6	96.5 ± 0.63	96.3 ± 0.42
	0.55 ± 0.0148 0.53 ± 0.0217 0.52 ± 0.0356 0.59 ± 0.0294	0.55 ± 0.0148 0.38 ± 0.0112 0.53 ± 0.0217 0.41 ± 0.0254 0.52 ± 0.0356 0.39 ± 0.034 0.59 ± 0.0294 0.46 ± 0.029	0.55 ± 0.0148 0.38 ± 0.0112 87.8 ± 0.61 0.53 ± 0.0217 0.41 ± 0.0254 88.6 ± 0.45 0.52 ± 0.0356 0.39 ± 0.034 90.4 ± 0.78 0.59 ± 0.0294 0.46 ± 0.029 90.6 ± 0.86	0.55 ± 0.0148 0.38 ± 0.0112 87.8 ± 0.61 96.4 ± 0.72 0.53 ± 0.0217 0.41 ± 0.0254 88.6 ± 0.45 96.5 ± 0.5 0.52 ± 0.0356 0.39 ± 0.034 90.4 ± 0.78 96.5 ± 0.52 0.59 ± 0.0294 0.46 ± 0.029 90.6 ± 0.86 96.59 ± 0.73



zero level contours can shrink in the level set evolution. The role of g in the energy term (area term) is to slow down the shrinking and expanding of the zero level contours when it arrives at object boundaries where g takes smaller values. The last term is the line integral, δ and H are Dirac delta function and Heaviside functions respectively [20, 24, 32]. Applications to image segmentation using edge information are obtained from the image

$$g \triangleq \frac{1}{1 + |\nabla G_{\sigma} * I|^{2}} \tag{4}$$

G stands for Gaussian kernel, I is the image and σ is the standard deviation (SD) of the kernel, which defines the blurring of Gaussian kernel. The convolution is used for smoothening the image to reduce noise.

Region-scalable fitting model

The region-scalable fitting (RSF) model uses the region based active contour model to provide information on the low-intensity pixels. A fitting energy is obtained from the region of interest in the image based on pixel intensities [33]. This fitting energy is divided into two functions: one defines the intensity information inside the boundary and the other approximates the region outside the boundary [34–36]. Due to space constraint only, the relevant equations are given here as it will be a repetition of what is already present in ref. [35]. The remaining equations and their derivations can be referred from ref. [35]. For an image I(x, y), the energy functional is

$$\mathcal{F}^{MS}(u,C) = \int_{\Omega} (u-I)^2 dx + \mu \int_{\Omega/c} |\nabla u|^2 dx + \nu |C| \qquad (5)$$

|C| is the length of contour C, I is the input image, u approximates the image I, μ is a constant, Ω is the image

domain and ν is a constant. The energy function is derived by minimizing Mumford –Shah function for a curve. It can be rewritten as follows for a given point $x \in \Omega$

$$\varepsilon^{fit}{}_{x}[C, f_{1}(x), f_{2}(x)] = \sum_{i=1}^{2} \lambda_{i} \int_{\Omega} K_{\sigma}(x - y) |I(y) - f_{i}(x)|^{2} *M_{i}[\mathcal{O}(y)] dy$$
(6)

 e^{fit}_{x} = fitting energy, λ_1 and λ_2 are positive constants, $f_1(x)$ and $f_2(x)$ are two values that approximate image intensities in domains Ω_1 and Ω_2 . I(y) = image intensities involved in the above fitting energy. K_σ is non negative kernel function.

Modified DRLSE

DRLSE has many advantages compared to the conventional level set method, in terms of initialization. However, it lacks in one feature that it uses only the edge information from the input image. This leads to incorrect segmentation of the ventricles, as it has both sharp and weak boundaries present at different regions of the ventricles as shown in Fig. 2. Even with the optimized parameter, it fails to converge sometime as shown in Fig. 3. Wherever a weak boundary is present with small change in the intensities, it was not well detected by the edge indicator as it considers only gradient changes, eventually leading to incorrect segmentation. DRLSE needs to have a term that accounts for pixel wise intensity information from the images which will detect very small changes in the intensity such as weak boundaries. Therefore, this study proposes the incorporation of the RSF concept into DRLSE for better segmentation.

RSF-based segmentation works well in case of weak boundaries as it accounts for regional (pixel wise information in a particular region) intensity information but

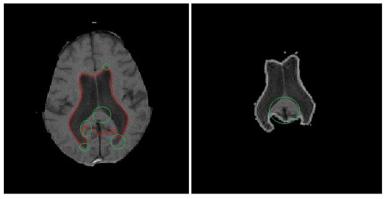


Fig. 2 Resultant images of segmentation using the conventional DRLSE method. The incorrect segmented regions are marked in green color circles

suffers from re-initialization [34–36]. RSF does not account for any edge information; sometimes it fails to converge and suffers from re-initialization. A study has been reported in the literature where DRLSE was applied first on the image, followed by RSF; which merely increases the computational complexity and time [37].

Therefore, this study proposes to incorporate regional intensity approximations of two regions from the RSF to the DRLSE framework which already has edge indicator and the regularization term. This way, the LSF will have region based and boundary based fitting energy term and a regularization term to avoid the complex re-initialization process. This reduces the computational complexity as there is only intensity information incorporated into the DRLSE equation instead of merely adding it to the framework, less computational time and efficient segmentation. This proposal is made to overcome the limitations of the DRLSE method. The proposed framework is given as follows (only the final equations combining both the frameworks are presented here as other equations and terms have been presented in the previous sections): Energy Functional equation is obtained by combining DRLSE and RSF. The RSF term is

$$\varepsilon(\varnothing) = \lambda_1 \int_{\Omega} K_{\sigma}(x - y) |I(y) - f_1(x)|^2 * M_1[\varnothing(y)] dy + \lambda_2 \int_{\Omega} K_{\sigma}(x - y) |I(y) - f_2(x)|^2 * M_2[\varnothing(y)] dy$$
(7)

The DRLSE equation for LSF

$$\varepsilon_{\varepsilon}(\emptyset) = \mu \int_{\Omega} p \mid \nabla \emptyset \mid dx + \lambda \int_{\Omega} g \delta_{\varepsilon}(\emptyset) \mid \nabla \emptyset$$
$$\mid dx + \alpha \int_{\Omega} g H_{\varepsilon}(-\emptyset) dx \tag{8}$$

Approximations for line and area term in the DRLSE framework

$$\lambda_{1} \int_{\Omega} K_{\sigma}(x-y) |I(y) - f_{1}(x)|^{2} * M_{1}[\mathcal{O}(y)] dy \cong \lambda \int_{\Omega} g \delta_{\varepsilon}(\mathcal{O}) |\nabla \mathcal{O}| dx$$
(9)

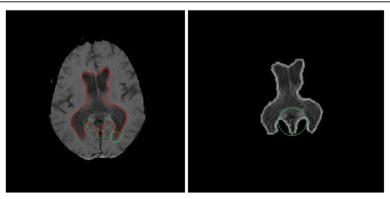


Fig. 3 Resultant images of segmentation using the conventional DRLSE method with optimized parameters. Without the regional information, the DRLSE framework fails to converge to the actual boundary

$$\lambda_2 \int_{\Omega} K_{\sigma}(x-y) |I(y) - f_2(x)|^2 * M_2[\mathcal{O}(y)] dy \cong \alpha \int_{\Omega} g H_{\varepsilon}(-\mathcal{O}) dx \tag{10}$$

The final equation by combining (8), (9) and (10).

$$\varepsilon(\emptyset) = \mu R_p(\emptyset)
+ \lambda \int_{\Omega} g |I(y) - f_1(x)|^2 * M_1[\emptyset(y)] dy
+ \alpha \int_{\Omega} g |I(y) - f_2(x)|^2 * M_2[\emptyset(y)] dy$$
(11)

Where M_1 and M_2 are the equivalents of Heaviside and Dirac delta functions presented in the second and third terms of Eq. (8), which were applied in the final computation. In Eq. (11), the edge indicator term g is from the DRLSE framework, $g|I(y)-f_1(x)|^2*M_1[\varnothing(y)]dy$ is the regional intensity approximation from the RSF framework from domain 1 (inside the boundary), and $g|I(y)-f_2(x)|^2*M_2[\varnothing(y)]dy$ is the regional intensity approximation from the RSF framework from domain 2 (outside the boundary) along with the edge indicator. The edge indicator of the DRLSE model is used here (g = edge indicator). Figure 4 shows the segmented ventricles using the proposed method. Step by step implementation of the proposed algorithm is shown in flowchart in Fig. 5.

$$K_{\sigma}(x - y) = g \tag{12}$$

Results

Twenty CT images and twenty MRI T1 weighted images of healthy subjects were considered. Twenty CT images and twenty MRI T1 weighted images of atrophy subjects were included in the dataset. The CT images chosen were of axial slices with slice thickness 5 mm and size of 512×512 pixels. The MRI images used were of axial slices with slice thickness 5 mm and size of 256×256

pixels. The images were collected from Calcutta Medical College, Kolkata, West Bengal, in digital imaging and communications in medicine format. The atrophy subjects were chosen by the expert radiologist and were confirmed by a clinical diagnostic process. Each brain volume was skull stripped using an algorithm which involves binarization, labeling, finding region properties and morphological operations. The segmented results were validated with ground truth images, made with the help of an expert radiologist. The brain images of respective slices were edited using image editing software; the borders of the ventricles were traced manually. After drawing the borders manually, the images were shown to the radiologist to verify the accuracy. Then, the regions inside the borders were extracted using MATLAB coding. These were used as ground truth images for comparison with the segmented images. Then the modified DRLSE method was applied as shown in Fig. 5.

Results of optimization of parameters

Table 1 shows the results of the optimized parameter for the input images. The segmented results were compared with the ground truth images using sensitivity, specificity, accuracy, Dice index, and Jaccard index. The average values of sensitivity, specificity, accuracy, Dice index, and Jaccard index for the input images were obtained and that producing higher values was chosen to be the optimized value of that parameter for segmentation. The Jaccard similarity index between two images A and B was defined as

$$JI = \frac{|A \cap B|}{|A \cup B|} \tag{13}$$

The Dice similarity coefficient between two images A and B was defined as

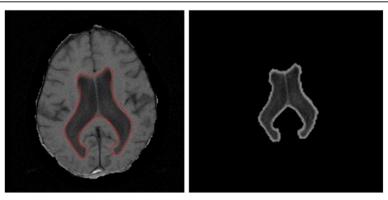
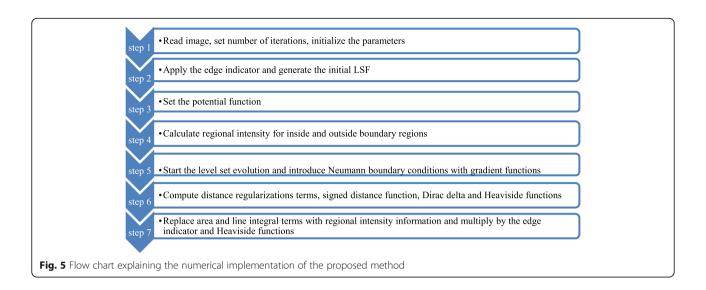


Fig. 4 Resultant images of segmentation using the proposed method with optimized parameters. With the regional information, the proposed method evolves to the actual boundary of the object



$$DI = 2*\frac{|A \cap B|}{|A| + |B|} \tag{14}$$

In case of the parameter epsilon which is used in the Heaviside function, epsilon-2 produced better results than other values for all the five measures (0.5 to 2.5) as highlighted in the table.

Segmentation results

Statistical analysis

Sensitivity, specificity and accuracy, peak signal to noise ratio (PSNR) and structural similarity index measure (SSIM) were used as performance measures. The formulae for these measures are as follows

sensitivity =
$$\frac{TP}{TP + FP}$$
 (15)

specificity =
$$\frac{TP}{TN + FP}$$
 (16)

$$accuracy = \frac{TP + TN}{TP + FN + TN + FP}$$
 (17)

$$PSNR = 10 \log_{10} \left(\frac{peakvaluve^2}{meansquarederror} \right)$$
 (18)

True negative (TN) is pixels correctly detected as background, true positive (TP) is pixels correctly segmented as foreground, false negative (FN) is pixels falsely detected as background, and false positive (FP) is pixels falsely segmented as foreground. Figure 6 shows the resultant images of segmentation using the proposed method. Figure 6a shows the initial bounding box around the ventricles in the input image.

Figure 6b shows the LSF for the initial bounding box. Figure 6c shows the final zero level contour of the object of interest after 236 iterations. Figure 6d shows the final LSF for the segmented object from the image after 236 iterations. This figure was included in the paper to show the steps involved in the segmentation process in terms of figures for the image and LSF.

Figures 7 and 8 show the measures used for assessing segmentation performance metrics such as sensitivity, specificity and accuracy for normal and atrophy subjects of CT and MR images respectively. It could be seen that the segmentation specificity (98%–99%) and accuracy (95%–98%) values were found to be higher for CT and MR images of normal and atrophy subjects. The image segmentation sensitivity values were found to be 90% for normal and atrophy subjects as shown in the Figs. 7 and 8. Further, it was observed that the lowest range of sensitivity for image segmentation was found to be low (46%).

Figure 9 shows the results of PSNR estimated by comparing the segmented images with the ground truth images. The mean and SD for normal and atrophy groups of CT and MR images are shown in Fig. 9. Both MRI and CT images for normal and atrophy subjects resulted in PSNR mean value of 95%.

Figure 10 shows the results of the SSIM calculated between the segmented images and the ground truth images. The similarity index is measured between 0 and 1. If the SSIM is 0 there is no similarity and if it is 1 there is 100% similarity between the images. Mean and SD for normal and atrophy groups of CT and MR datasets are shown in Fig. 10. Both normal and atrophy groups of CT and MRI datasets yielded SSIM values of 0.95 with varying SDs.

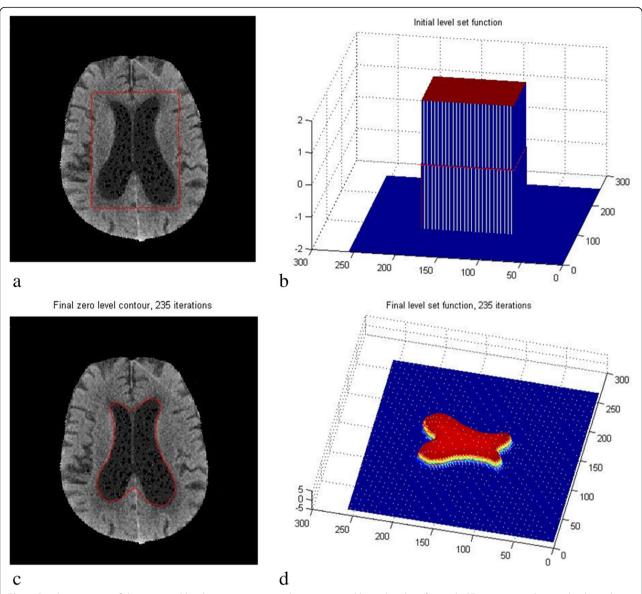
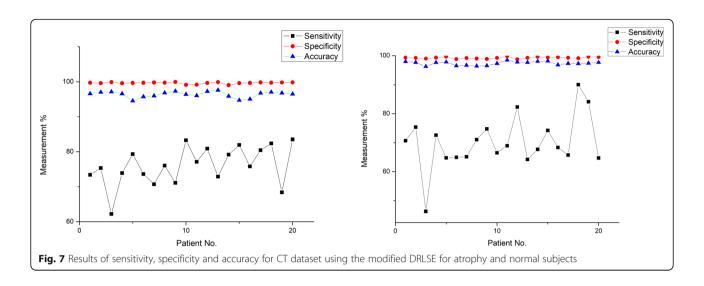


Fig. 6 Resultant images of the proposed level set segmentation showing **a** initial bounding box, **b** initial LSF, **c** segmented ventricles; boundary marked in red color after iterations and **d** LSF of the segmented ventricles

Table 2 presents the comparison of results with recent papers, based on statistical analysis measures such as sensitivity, specificity and accuracy. In this study, few segmentation methods based on level set formulation and few based on other segmentation frameworks such as fuzzy based method, hybrid algorithm, and self-organizing map-based methods were included for comparison. The conventional DRLSE method was also implemented for the images and the results are presented in the Table 2.

It is evident from Table 2 that, the proposed DRLSE with RSF framework has yielded better results than other methods (overall based on all the three

measures). Moreover, the proposed method is multimodal; performs well on CT and MR images. Accuracy (95%–98%) and specificity (98%–99%) are higher than all the other methods compared in this study. The sensitivity also falls in close range with other studies. The proposed modification was made in the external energy part that is derived from the image information (edge and regional intensity). Since the regularization term is not modified there will be no changes in the stability and re-initialization of the level set framework as DRLSE does not require reinitialization. All the methods shown in the table were applied on MR images.



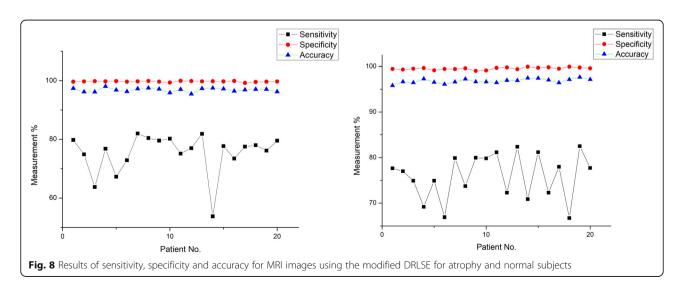
Discussion

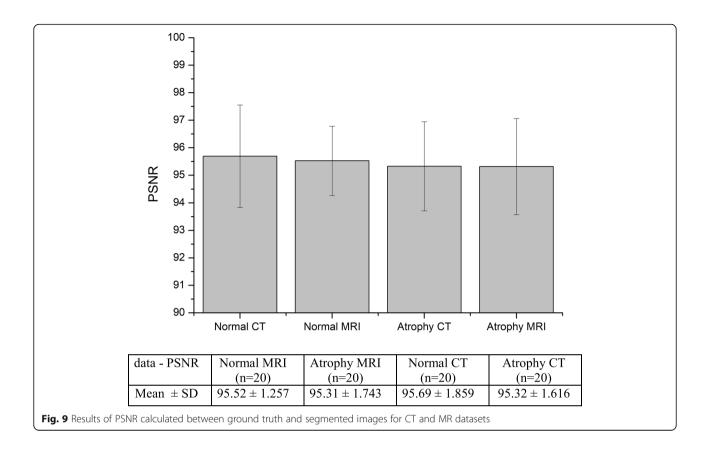
Computer-aided diagnostic systems are required to perform quantitative characterization of medical images. This needs enhanced visualization which could be used for differential diagnosis. It is a common procedure to first extract the object of interest from these images to get a better visualization of the region of interest.

This paper presents a segmentation framework based on the modified DRLSE formulation for segmenting the ventricles from brain images of normal and atrophy subjects. Optimization was performed for the parameters that are involved in the segmentation algorithm for the given dataset. Figures 7 and 8 illustrate the segmentation results for CT and MRI datasets of normal and atrophy groups. Consistent results were obtained for both MRI and CT images containing normal and atrophy subjects. The specificity had

the most consistent results across normal and atrophy subjects for CT and MRI dataset, with a SD of 0.1% to 0.3% while accuracy was the second most consistent with a SD in the scale of 0.4% to 0.9%. The sensitivity yielded the lowest values with a wide variation of 5% to 9%.

In addition to sensitivity, specificity, and accuracy measures, PSNR and SSIM were also measured to estimate the efficiency of the proposed method. PSNR values were in the range of –95% with SDs ranging from 1%–2%. SSIM also produced similar results in the range of 0.95 as shown in Figs. 9 and 10. It was observed that, the segmentation algorithm produced similar results for both groups. The proposed method was shown to perform better than other segmentation algorithms reported in the literature as shown in Table 2. The execution time for the proposed method takes up to 2 min for both CT and MR images.





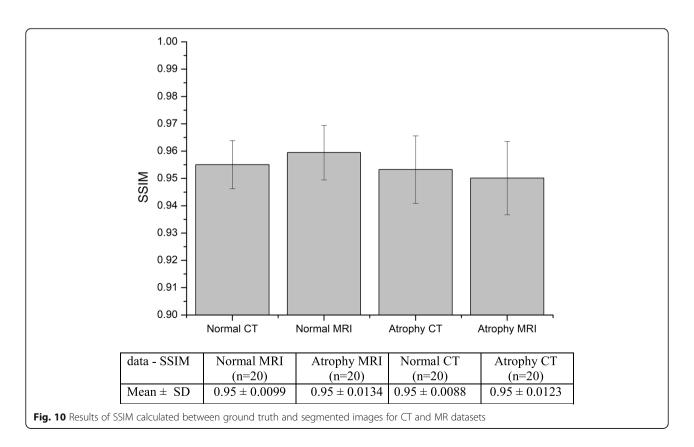


Table 2 Results of the comparison of the proposed method with other segmentation methods from the literature

Methods	Sensitivity	Specificity	Accuracy
Proposed method (atrophy CT, $n = 20$; normal CT, $n = 20$; atrophy MRI, $n = 20$; normal MRI, $n = 20$)	65%–90%	98%–99%	95%–98%
DRLSE (implemented)	74%-79%	91%-93%	92%-94%
Region based level set (MRI, $n = 124$) [38]	84%-89%	96%-97%	-
DRLSE and deep structure (MRI, $n = 45$) [39]	-	_	84%-95%
Fuzzy and evidential reasoning (MRI, $n = 17$) [40]	55%-90%	_	60%-80%
Hybrid genetic and EM algorithm (MRI, $n = 10$) [41]	-	_	45%-85%
Bayesian morphometry (MRI, $n = 95$) [42]	60%-80%	30%-60%	-
EGS-SOM (MRI, $n = 57$) [43]	76%-80%	95%–96%	-
HFS-SOM (MRI, $n = 57$) [43]	66%-77%	80%-85%	=

Conclusions

This study presents a level set based segmentation technique for segmenting ventricles from brain images for atrophy and normal subjects in CT and MRI datasets. The proposed method is based on DRLSE and RSF. The DRLSE method does not require reinitialization process, but it lacks the information on the regional intensity of the object being segmented. RSF derives its surface energy from the regional intensity information. The present study proposed a framework, combining the regional information from RSF with the conventional DRLSE method to improve its efficiency. The proposed method produced sensitivity in the range of 65%-90%, specificity in the range of 98%-99% and accuracy in the range of 95%-98%. PSNR and structural similarity index were also used to measure the segmentation accuracy and produced good results: 95% and 0.95 respectively. In addition to segmentation, an optimization procedure was carried out to fine tune parameters of DRLSE formulation for the input image dataset used. Presented method was found to be efficient and fast. The proposed method is adaptive for normal and atrophy subjects and is multi-modal. Future studies should involve analyzing shape differences of ventricles of normal and abnormal subjects for diagnosis and forming a generalized optimization technique.

Abbreviations

CT: Computed tomography; MRI: Magnetic resonance imaging; PET: Positron emission tomography; SPECT: Single photon emission computed tomography; CSF: Cerebrospinal fluid; MCI: Mild cognitive impairment; DRLSE: Distance regularized level set evolution; LSF: Level set function; RSF: Region-scalable fitting; SD: Standard deviation; PSNR: Peak signal to noise ratio; SSIM: Structural similarity index measure; TN: True negative; TP: True positive; FN: False negative; FP: False positive

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Authors' contributions

AS analyzed and interpreted the patient data regarding the atrophy condition and normal subjects. SR performed the optimization process and helped in generating the ground truth images. TJ carried out literature

survey, analyzed and processed the images and was a major contributor in writing the manuscript. MM and PKD supervised the data collection, image processing and manuscript structure. All authors read and approved the final manuscript.

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Availability of data and materials

The datasets generated and/or analyzed during the current study are not publicly available due to ethical approval and policy but are available from the corresponding author on reasonable request.

Competing interests

The authors declare that they have no competing interests.

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