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## TOPICAL REVIEW

# Developments in Brain Tumor Segmentation Using MRI: Deep Learning Insights and Future Perspectives

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**ABSTRACT** The human brain is an incredible and wonderful organ that governs all body actions. Due to its great importance, any defect in the shape of its regions should be reported quickly to reduce the death rate. The abnormal region segmentation helps to plan and monitor the treatment. The most critical procedure is isolating normal and abnormal tissues from each other. So far, remarkable imaging modalities are being used to diagnose abnormalities at their early stages, and magnetic resonance imaging (MRI) is renowned and noninvasive among those modalities. This paper investigates the current landscape of brain tumor segmentation (BTS) by exploring emerging deep learning (DL) methods for brain MRI analysis. The findings offer a comprehensive comparison of recent DL approaches, emphasizing their effectiveness in handling diverse tumor types while addressing limitations associated with data scarcity and robust validation. DL has shown a vital improvement for BTS, so our primary focus is to include significant DL robust models to analyze the brain MRI. However, DL outperforms traditional methods; still, there are several limitations, especially related to the diverse tumor types, lack of datasets, and weak validations. The future perspectives of DL-based BTS present significant potential for revolutionizing the diagnosis and treatment of brain tumors.

**INDEX TERMS** Brain tumor segmentation, deep learning, medical imaging, MRI.

## I. INTRODUCTION

Image segmentation is one of computer vision's basic and challenging tasks [1]. In recent decades, its research has been increasingly innovative and the development process continues to advance with the emergence of new technologies. Several high-level computer vision applications need positioning, recognition, detection, and segmentation functions.

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The processing of medical images needs to detect objects, and we often use the way of detecting specific semantic objects in digital images and locating them by their bounding boxes to obtain the results. Image segmentation aims to determine whether there are characteristic objects to be detected in the image. If so, the coordinate pixels and area size of each object detected can be attained [2].

With the continuous development and popularization of medical imaging, the tumor detection rate is increasing [3], [4]. Radiation and laser therapy greatly rely on accurate

tumor segmentation. Otherwise, it can be harmful to several sensitive nerves [5]. Medical treatment first requires doctors to give diagnostic results while observing the lesions. This process will cause a great burden on the clinical medical system for a large number of patient groups. Moreover, the subjective factors of finding between different doctors will cause trouble in clinical diagnosis and treatment. Therefore, a system that can comprehend automatic medical image segmentation will have great clinical significance. Glioma is the most frequent brain tumor in adult brain diseases [6]. It will cause irreversible damage to the brain by infiltrating the surrounding tissues. Accurately segmenting brain gliomas is very time-consuming in the process of clinical diagnosis and planning of treatment. If medical resources can be liberated through automatic segmentation, many medical resources will be saved for research work that needs more human intervention. Such resources promote the development of medical image processing-related technologies. Medical image processing is a necessary and technical means for diagnosing and planning by analyzing the lesions in the brain tumor focus area. BTS aims to classify the pixels of the lesion area in an image and obtain the detailed distribution of the lesion area, which can help doctors better understand the symptoms and diagnosis. Traditional image segmentation approaches can be divided into graph theory segmentation methods, watershed image segmentation algorithms, and so on [7]. These algorithms usually rely on the feature extraction of the image itself and its color, texture and other shallow features. Brain gliomas usually infiltrate the surrounding tissues. Their shallow features are not as obvious as those in natural images, which can affect the performance of traditional segmentation methods for BTS. It can affect the segmentation accuracy, and create other problems to prohibit the final segmented image to put into use, resulting in the decline of the significance of introducing image segmentation into clinical medical auxiliary diagnosis.

Various problems, such as environmental pollution, irregular life, and population aging, are becoming critical. Because of these factors, the incidence rate of brain diseases such as epilepsy, Parkinson's disease, brain tumors, and cerebrovascular diseases is increasing yearly [8]. It seriously threatens human health. Especially brain tumors and cerebrovascular diseases are becoming the first cause of death in China due to their high disability rate, high mortality, and high recurrence rate. The medical diagnosis of the brain, especially brain tumors and cerebrovascular diseases, often depends on medical imaging such as computed tomography (CT) [9], X-rays and magnetic resonance imaging (MRI) [10], [11]. In addition, these medical imaging techniques are also helpful for surgical planning and postoperative treatment effect evaluation. Therefore, an accurate computer-aided system can help doctors quickly locate lesions, assist in recognition, and improve doctors' efficiency and accuracy. With the development and large-scale application of machine learning, computer-aided medical image analysis based on machine

learning has become one of the research hotspots in medical image analysis and machine learning. Although the current level of medical technology has made significant progress compared with the past, there is still a lack of effective treatment for brain tumors. Patients can only prolong their lives through various conservative, comprehensive treatments and operations with high-risk coefficients. If the brain tumor can be found at the early stage of its growth then the patients can get the first opportunity for treatment, and the survival probability will be greatly improved. The technology based on medical image segmentation has been applied to early brain tumor detection, but the traditional manual segmentation method requires much professional labor. It needs doctors with rich clinical experience and much professional knowledge to divide manually. Due to the lack of medical resources, there is no doubt that this method cannot be popularized and applied on a large scale.

The core problem of BTS based on deep learning (DL) is to extract, fuse and classify the different levels of brain tumor image feature information through DL technology within acceptable consumption and to improve the final accuracy of the model.

Most research works mainly concentrate on traditional structures, with little creativity in creating new neural network architectures that are customized for the complexities of brain tumor segmentation (BTS). Closing this gap requires a move toward the creation of complex models that can efficiently utilize the complementing information present in multimodal data. Furthermore, the evaluation and comparison of current approaches frequently lack critical information, which makes it difficult to establish a benchmark for assessing the actual usefulness of various strategies. Many studies using image-guided surgery use a variety of datasets that display a range of imaging modalities, including MRI, CT, and intraoperative datasets [12]. Typically, these datasets include annotated tumor area examples, making algorithmic training and validation easier. This review paper tries to summarize the available literature on image-guided surgery, focusing on DL's critical role in improving precision. The most recent approaches are briefly reviewed and demonstrate advances in recurrent architectures, attention mechanisms, and convolutional neural networks (CNNs), demonstrating the ongoing development of DL methods for precise and effective BTS in the context of image-guided surgical interventions. This paper mainly focuses on the recent developments and challenges for BTS. The key points for this contribution are:

(a) The paper introduces and evaluates cutting-edge DL methods for BTS in MRI during 2016-2023, showcasing a significant leap forward in the methodology employed for accurate analysis.

(b) The findings offer a thorough comparison of recent DL approaches, emphasizing their effectiveness in handling diverse tumor types. The paper addresses data scarcity and validation robustness limitations, providing valuable insights for researchers.

(c) The open challenges for BTS based on DL and traditional approaches have been highlighted. The advanced DL impacts on BTS and future recommendations are comprehensively described.

This paper consists of seven sections. Section II describes the background of the study. Section III illustrates the basic theory of BTS. Sections IV and V present the traditional and DL-based BTS methods, respectively. Section VI comprises important types of image-guided surgery for brain tumors. Section VII covers discussion and future recommendations. Finally, section VIII concludes this review paper.

## II. BACKGROUND

The research fields of computer vision include remote sensing applications, vehicle and pedestrian monitoring, medical image segmentation and recognition, map satellite navigation, tire defect detection, object positioning and recognition [13], scene classification and segmentation [14], [15], etc. It has also been widely used in various fields of daily life, such as military territorial air defense, medical disease analysis, intelligent video monitoring [16], and remarkable success in specific objects such as intelligent home machine control, surveillance video recognition, and medical surgery object positioning. Many object detection achievements proposed by scientific researchers for a long time have gradually increased, mainly including traditional and DL-based object detection methods [17], [18].

DL is a new technology that has been rising recently, and it is a new branch under the machine learning sub-directory [19], [20]. Its concept comes from scientists' deeper research on artificial neural networks (ANNs) [21], such as multilayer perceptron (MLP) from the perspective of structure [22], which is a simple DL structure. The network contains hidden layers, which is a typical feature of the DL structure. The DL model mainly uses a mathematical means (convolution) to extract the deep hidden feature representation in the input data. Based on the extracted feature representation, the network model can fit the input data well without manual preset rules like other methods, which is the unique learning-fitting ability of the DL network. In recent years, the best DL method in medical image processing originated from LeCun, known as CNN [23]. Later generations have made many simple and effective improvements from different angles, further enhancing CNN's data representation ability. The CNN training is more stable and simpler, which is suitable for various task scenarios. Although CNN was proposed long ago, it is still promising for semantic segmentation and object detection. The models with extraordinary performance are basically improved based on CNN variations. The number of published papers during 2016-2023 are extracted from PubMed with the included expressions of "Brain Tumor", "Deep Learning", "Artificial Intelligence", "CNN" and excluded expressions "case study", "overall surgery", "Model training for information learning" and "Not accessible" as shown in Figure 1. This analysis shows that numerous articles have

been published on BTS during 2016-2023 which reveals the importance of this research.

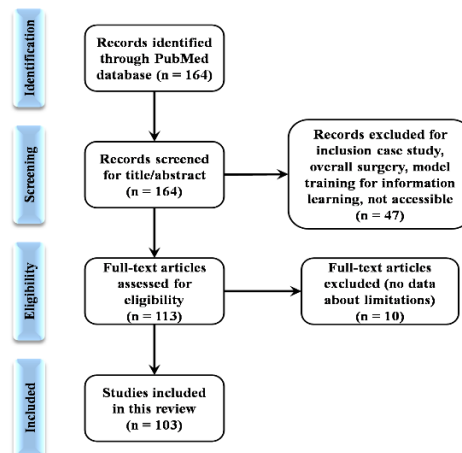


FIGURE 1. PRISMA diagram of systematic review of BTS from 2016 to 2023.

At present, partial resection of brain tumors through surgery is the best treatment method. Accurate BTS images can help doctors quickly view the lesions and implement treatment to reduce patients' pain. However, this segmentation task for brain tumors is not easy, not only because the size, shape and location of gliomas are quite different among patients, but also affected by many factors, which will lead to inaccurate segmentation, which greatly limits the availability of glioma segmentation information [24]. Big data analysis and preprocessing techniques such as cleansing, profiling, enrichment, and transformation play critical roles in early BTS [25].

In addition, the tumor mass effect will change the arrangement of surrounding normal tissues. All medical imaging modalities have some shortcomings and planar X-ray can be used to visualize the skull but has no diagnostic value for brain pathologies. For example, intensity heterogeneity or different intensity ranges between acquisition scanners [26]. The diversity of MRI acquisition parameters and sequences leads to great differences, increasing glioma variability between different patients. Therefore, it is very important to help patients find tumors as soon as possible and carry out relevant diagnosis and treatment [27]. At present, the tumor area is manually marked and segmented by radiologists. However, due to the changes caused by the appearance and shape of the glioma, the process is very time-consuming, and the consistency between evaluators is low. Therefore, automatic segmentation is attractive because it can describe relevant tumor parameters faster, more objectively, and accurately. However, due to the irregular nature of tumors, developing algorithms that can segment tumors efficiently and accurately is still challenging [28]. This inherent heterogeneity of glioma is also reflected in its imaging phenotype. Its sub-region is generated by multimodal MRI, which reflects

TABLE 1. Comparison of our review with existing reviews.

Ref. No.	Conventional Approaches	Deep Learning Approaches	Scope			Data/Repository Links
			Image Guided Surgery Techniques	Drawbacks/Limitations	Future Perspectives	
[24]	✓	✓		✓		
[26]		✓			✓	
[27]	✓	✓		✓	✓	
[28]			✓			
[29]	✓	✓		✓	✓	
[36]		✓		✓	✓	✓
[37]	✓	✓		✓	✓	
[38]		✓		✓	✓	✓
<b>Our paper</b>	✓	✓	✓	✓	✓	✓

different biological characteristics of tumors according to different intensity distributions [29].

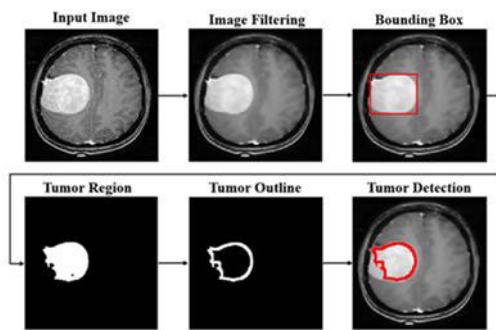


FIGURE 2. BTS and detection using MRI.

The object detection algorithm is also applied to the field of medical images. Tumor detection using MRI has been attained by implementing preprocessing approaches such as image filtering, thresholding, morphological operations, eroding, and subtraction, as shown in Figure 2. We have perceptibly described the contribution of this review compared to existing reviews in Table 1. Early tumor detection tasks are usually based on some traditional image algorithms [30]. Karkanis et al. proposed a method of tumor detection assisted by color wavelet features [31]. This method is based on a novel color feature extraction to depict different regions in the sequence and on the covariance of the second-order texture measure. A feature selection algorithm is proposed to determine the image regions along the video frame using linear discriminant analysis. Bercoff et al. proposed a tumor detection method using transient elastic imaging [32]. This method detects tumors by tracking the propagation of extremely low-frequency shear waves generated by the vibration system on the body surface in soft tissue. It has been developed to detect and quantify soft tissue and hard lesions. Wu et al. proposed a color-based tumor detection method using

K-means clustering technology to observe tumor objects in MRI [33]. Its core idea is to convert a given gray MR image into a color space image and then use K-means clustering. Iftekharuddin et al. proposed a fractal-based multimodal MRI tumor detection method, which integrates two novel texture features and intensity in multimodal MRI images [34]. The texture features involve the segmented triangular prism surface area (TPSA) technique for fractal features. Meanwhile, the other texture feature uses the Brownian motion method, which integrates wavelet and fractal analysis for fractal feature extraction. Mustaqeem et al. proposed a method for tumor detection based on watershed and threshold segmentation [35]. By improving the quality of scanned images, morphological operators are used to detect tumors in scanned images. For the traditional tumor detection methods, on the one hand, due to the variety of tumor shape, size and appearance, accurate measurement is challenging. The tumor can grow suddenly, leading to the defects of adjacent tissues, providing an overall abnormal structure for healthy tissues. On the other hand, applying a tumor detection algorithm is complicated due to the particularity of medical images.

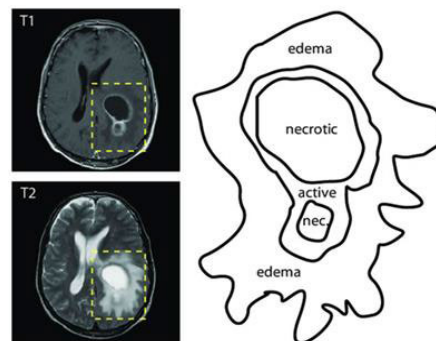
### III. THEORY OF BRAIN TUMOR SEGMENTATION

In the middle of the last century, Warren McCulloch et al. proposed the mathematical model of ANN, which laid the theoretical foundation of neural network structure in future generations [39]. Frank proposed a machine to simulate human perception and named it perceptron [40]. The structure of the perceptron is relatively simple. It is a single-layer network composed of threshold and linear units. Although single-layer perceptron can only classify linear tasks, its emergence represents the prototype of modern CNN structure. The brain tumor illustration has been presented in Figure 3, which describes the importance of different modalities. The recent developments and basic theory have been described in the following sections.

### A. ARTIFICIAL NEURAL NETWORK (ANN)

At the beginning of ANN, the computing power of the first generation of computers could not match it, which limited the role of neural networks, so it did not enter the stage of rapid development. The current computer can give full play to the characteristics of neural networks and promote the rapid development of artificial intelligence. In essence, ANN uses mathematical methods to imitate the complete process of biological visual perception [41]. In this bionic process, the data input into ANN can be regarded as the electrical stimulation signal obtained by visual cells. The network's data processing corresponds to the stimulation signal's transmission process in the biological visual perception system. Finally, the network simulates the performance of organisms for different stimuli and outputs different results. Artificial neurons play an absolute core role in the components of ANN. Multiple neurons can be stacked and connected in a certain way to form ANN.

The neurons in the same layer of ANN are not connected. The neurons between adjacent layers are connected through the weight matrix. The connection mode can be customized. Any neuron in the hidden layer is connected with all neurons in the adjacent layer. The network adopts the forward calculation method: the output of the previous layer is the input of the next layer, which is calculated layer by layer, and finally, the output is obtained. To learn the parameters in the network, Rumelhart et al. proposed an error backpropagation algorithm (BP) [42]. The algorithm first calculates the error between the network output and the true value, then backpropagates the error from back to front, adjusts the parameters of the network according to the error, and finally iterates continuously to make the network model converge to the global optimization or local optimization. From this intuitive process, we can understand the whole operation process of neurons. If a huge number of neurons are connected somehow, the ANN can disclose complex nonlinear tasks. Multilayer Perceptron (MLP) is a typical complex network that combines multiple neurons in a predetermined way. MLP is composed of layers of neurons and nodes. The layer closest to the input data is often called the input layer neuron, whose main function is to read the data into the shallow layer of the network model. The next series of neurons are called hidden layer neurons. In these layers, the network is mainly used to extract the multi-level features of input data, which is an important part of the network. The number of hidden layers is generally unlimited, and the appropriate number can be selected according to the task objectives. Developers need to make a trade-off between efficiency and accuracy. The last layer of neurons is called the output layer. After passing through the output layer, the network will provide the analyzed results of specific tasks. From this process, we can see that there is mutual propagation between multiple neurons. Based on this relationship, the characteristics of the input data move forward layer by layer in the neural network and finally form the output result of the network.



**FIGURE 3.** Example of a brain tumor showing the importance of the diverse modalities (T1 with contrast and T2) [43].

### B. CONVOLUTIONAL NEURAL NETWORK (CNN)

The most significant change brought by CNN to deep neural networks is the introduction of convolutional structures in the process of network construction. Applying this structure in deep neural networks can effectively reduce the memory and parameters occupied by the deep network calculation process in the training process, accelerate the training process, and make the network easier to converge. Compared with the traditional ANN, CNN reduces the training complexity of the network model by setting local connections in the model and sharing the weight value of each neuron in the calculation process. The number of parameters in the network is effectively reduced, and the possible overfitting phenomenon is alleviated. Another special feature of CNN is that it is very suitable for processing two-dimensional images. The convolution structure can effectively extract various features of the image (such as approximate shape, texture detail information, color information, etc.) in the network. These features help the network model understand the internal information in the image and improve the ability of the network to process image data. Because the CNN network has good translation invariance, it can still accurately extract and understand the image's content even if the image's shape changes. Because of these characteristics of CNN, there are more and more new ideas and methods in computer vision.

LeCun et al. [44] proposed CNN in handwritten character recognition and achieved great success. With the great success of AlexNet [45] in classifying ImageNet dataset, CNN has been widely used in image processing. The traditional ANN often adopts the way of full connection, and the network parameters increase exponentially with the increase of layers, which makes the network difficult to train. Even the shallow network is easy to overfit. In order to alleviate this problem, CNN introduced strategies such as local receptive field and weight sharing [46], which not only reduced the network parameters but also the weight sharing strategy can automatically extract image features from the layers. With the development of CNN, various excellent basic CNN models have emerged, such as VGG [47], GoogleNet [48], ResNet

[49], DenseNet [50], etc., but all networks are built on a series of functional layers.

#### IV. BRAIN TUMOR SEGMENTATION METHODS

Among the advanced technological developments, two main factors are limiting BTS; one is medical imaging technology, and the other is segmentation methods. MRI has become a key technology suitable for brain imaging with the continuous development of medical imaging technology. MRI completes the imaging process of complex brain structures based on the magnetic field and radio waves. MRI is a promising imaging method that does not damage the human body and can generate multimodal data. The tissue characteristics presented by different modal data are different, and the multimodal data complement each other to ameliorate the accuracy of tumor segmentation. After solving the imaging problem, the key point of developing BTS is the research of segmentation methods. In recent years, the development of deep neural networks has led to the segmentation accuracy of segmentation methods based on DL has far exceeded the traditional segmentation methods. The significant segmentation methods based on supervised learning are random forest, decision tree, particle swarm optimization (PSO), dynamic sparse field (DSF); and based on unsupervised methods are K-means, C-means, mean shift, self-organizing map (SOM), markov random field (MRF), gray-level cooccurrence matrix (GLCM), multi-context fuzzy clustering (MCFC), and extreme learning machine (ELM). Some BTS methods based on traditional and DL approaches are divided into groups, as shown in Figure 4.

In BTS, the region to be segmented is not an independent region but multiple overlapping regions, such as the complete tumor region, core tumor region, and enhanced tumor region. The complete tumor region includes all tumors, and the enhanced tumor will be included in the core tumor region. Due to this characteristic, some scholars proposed that the BTS task can be divided into multiple subtasks for training to reduce the complexity of directly segmenting all the regions. At the same time, optimizing the single-segmented region is easier, and improving the overall segmentation accuracy through the fusion of the final segmentation results. Based on this idea of sub-task segmentation, Wang et al. proposed a model including three cascaded segmentation networks for BTS [51]. The three segmentation networks segment different tumor regions, respectively. The overall segmentation process is divided into three steps, and each step is re-segmented based on the previous step. Zhou and others proposed a one-step multi-task segmentation model for learning shared features, which divides the brain tumor into different segmentation tasks, integrates these tasks into one model, strengthens the interaction between tasks through shared parameters in the training process, and can complete multi-task prediction in one step directly in the prediction stage [52]. Shen et al. proposed a multi-task segmentation network, which distinguishes the regions through the up-sampling operation of different paths, then uses the designed loss function for binary

classification, and finally fuses the different segmentation results to form a complete region segmentation [53].

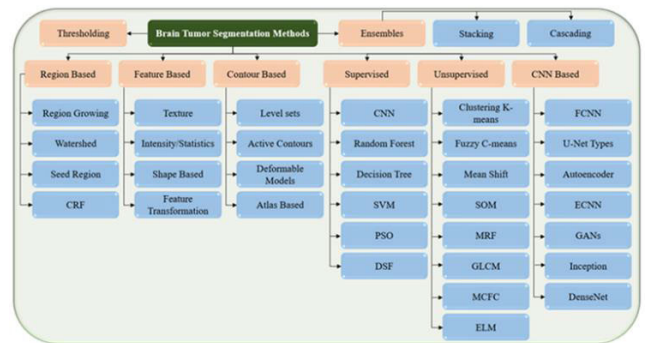


FIGURE 4. Classification of BTS methods.

##### A. REGION-BASED SEGMENTATION

The process of grouping pixels into larger regions is known as region growing which has been utilized in region-based image segmentation algorithms [54]. Initially, scan the adjacent pixels and evaluate the merging of adjacent pixels into the region. Each pixel in the same attribute set will be used to allocate pixels for the growth process of the region. The shape of the region grows according to the intensity to complete the morphological edge detection of the input image and reconstruct the input image based on expansion and corrosion to enhance the image. The center point selects the pixels within a certain gray range and the pixels are evenly placed in the divided area. The receiving area starts from the accurate position of the first pixel, and then the seed point grows according to the set rules. An example of region-based segmentation is shown in Figure 5.

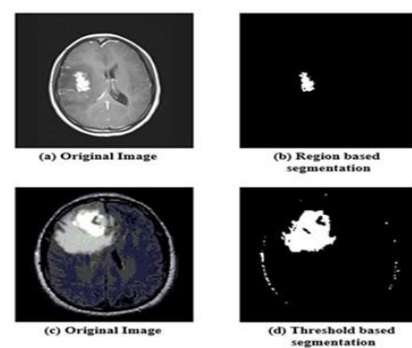
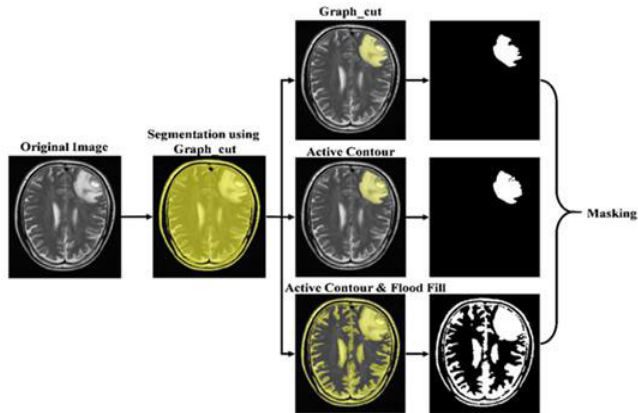


FIGURE 5. Results on MR brain image. (a, c) Original image (b) the result using the region growing model (d) threshold-based segmentation for brain MRI [55], [56].

##### B. THRESHOLD-BASED SEGMENTATION

Threshold-based segmentation is one of the simplest image segmentation methods because it consumes less computation and has high efficiency, so it plays an effective and significant role in image segmentation. The gray image is converted into

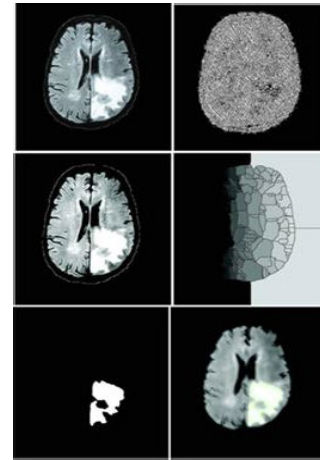
a binary image output corresponding to its region using a threshold. This method traverses all image pixels. If the pixel value of the image is higher than the determined threshold, the pixel is set to the maximum value of the proportion used; otherwise, it is the minimum value. Threshold selection techniques can be divided into two categories: two-level and multilevel. In the former case, only one threshold is needed to separate two objects of the image: one represents the object, and the other represents the background. When different objects are depicted in a given scene, multiple thresholds must be selected for correct segmentation, usually called multilevel threshold processing [57]. As seen in Figure 5, thresholding is used to process the brain MRI. Automatic image segmentation is proposed using cranium removal, morphological reconstruction, thresholding, and subtraction to get a binary mask for segmentation [58]. The segmentation results using active contour and graph cut theory have been shown in Figure 6.



**FIGURE 6.** Examples of segmentation using graph cut, active contour, and flood fill algorithms.

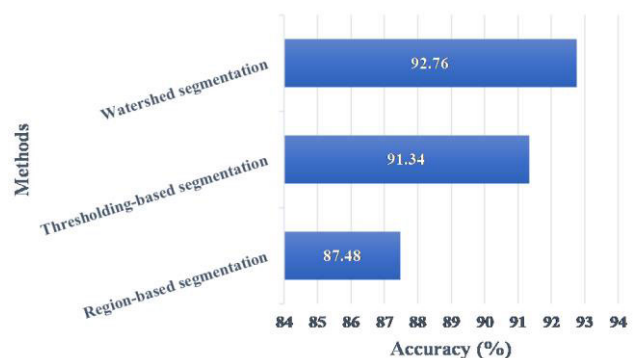
### C. WATERSHED SEGMENTATION

A watershed algorithm is a mathematical morphology segmentation algorithm based on topology theory. It was introduced and improved by Lantuéjoul and Beucher [61], [62]. Subsequently, in terms of definition and implementation, several algorithms of watershed transformation with different variants are developed to reduce the computational complexity [63], [64]. The watershed algorithm realizes segmentation by constructing regions. The image to be segmented is regarded as a topographic map in geography. The gray value of pixels is used to represent the height of the terrain [65]. Mountains and valleys correspond to the gray value, respectively. When we inject a drop of water into the valley, the water will eventually converge to the local lowest point due to the action of gravity. There may be a minimum plane in which all points are minimum points. Pierce the valley, the bottom begins to fill with water, and the water level rises at a constant rate. As the water level rises, the water becomes more and more, and finally covers the whole surface. The



**FIGURE 7.** Result of watershed segmentation for brain MRI from top to bottom (a) artifacts removal (b) watershed transform and (c) edge smoothing [59].

watershed segmentation and accuracy assessments of conventional methods for BTS are shown in Figure 7 and 8 respectively.



**FIGURE 8.** Accuracy assessment of the traditional segmentation methods [60].

### D. CLUSTERING-BASED SEGMENTATION

The collection of similar data into clusters depends on homogeneity, known as clustering. Among clustering algorithms, K-means clustering [66] is the base for image segmentation. The similar components in the dataset belong to the same cluster in this algorithm. On the other hand, fuzzy C-means (FCM) clustering resolves the multi-grouping issue by assigning each pixel to a separate class. It means one pixel could associate with more clusters, although each pixel reveals a distinct similarity value for every cluster. The optimization function of the C-means algorithm affects the accuracy of results. Among clustering algorithms, K-means improved K-means [67], and C-means [68] are most commonly used in segmentation [69]. There are several improved methods based on the FCM fundamental approach, such as kernelized fuzzy C-means (KFCM) [70], generalized fuzzy C-means (GFCM) [71], fast generalized FCM algorithm

(FGFCM) [72], enhanced FCM (EnFCM) (..) [73] and so on. Similarly, the K-means algorithm has been widely used for image segmentation [74], [75], [76], [77], [78]. The brain image segmentation using K-means clustering at different K levels is shown in Figure 9.

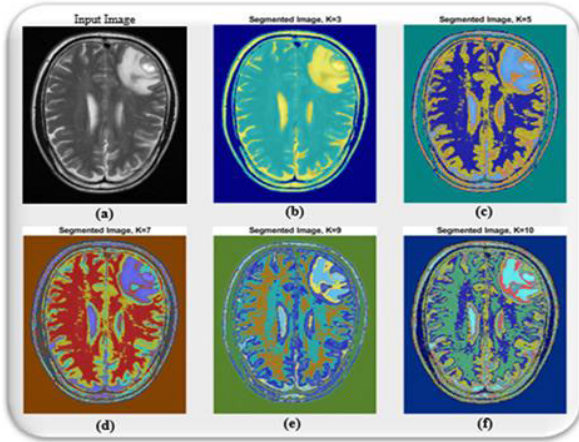


FIGURE 9. Brain image segmentation using K-means clustering algorithm at different K values.

### V. NEURAL NETWORK-BASED BRAIN TUMOR SEGMENTATION

In the basic theory of BTS, we will introduce the related basis of neural networks. Firstly, the related principle of ANN, the structure and application of common and advanced CNNs are introduced. Then several common segmentation frameworks based on CNN will be discussed in the following sections. Image segmentation is one of the widely used directions of the CNN model. According to the input and output, image segmentation methods based on CNN can be divided into two categories: image block-based segmentation and end-to-end segmentation.

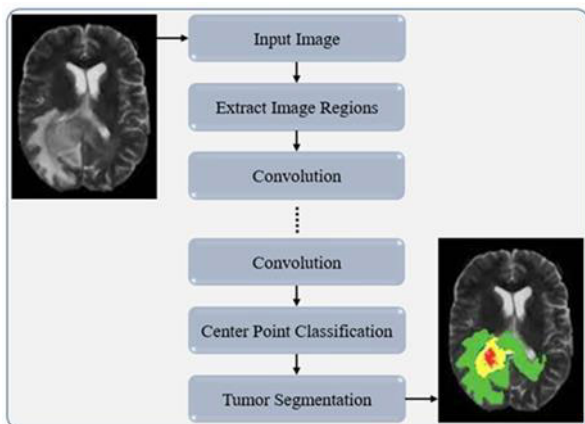


FIGURE 10. BTS using region extraction and CNN.

#### A. IMAGE REGION-BASED SEGMENTATION METHODS

The success of AlexNet in image classification promotes the application of CNN in image segmentation. The traditional

CNN model includes three functional layers: convolutional layer, pooling layer, and fully connected layer. It can automatically extract the features of the image and classify the image by using the image features. Therefore, the segmentation method based on image block maps a fixed-size image block in the image to the category using CNN’s powerful fitting ability. Figure 10 shows the segmentation method framework based on regions. Firstly, the regions are extracted. Since the center points of the regions are classified, it is necessary to use the sliding window method to obtain the fixed-size blocks in the image. Then, CNN is used to extract features and classify the center points of regions. Finally, the classification of all pixels is the segmentation result of the whole image.

Because the image segmentation task is transformed into the classification of region center points, the commonly used CNN classification models, such as AlexNet, GoogleNet, ResNet, etc., can be used for image segmentation. However, unlike image classification, the size of regions often greatly impacts the network’s results. Another point is that different types of regions can be selected through certain strategies in network training. Havaei et al. developed a CNN-based tumor segmentation method by exploiting global and local features simultaneously, exhibiting promising results [79].

#### B. END-TO-END SEGMENTATION METHODS

The resolution of the feature image extracted by the traditional CNN model decreases with the increase of the number of network layers, so the classification center point is used to evaluate image segmentation. However, with the upsampling and deconvolution layers, the low-resolution feature map can be generated into a high-resolution feature map to analyze the end-to-end segmentation method. Common end-to-end segmentation frameworks include FCN [80], SegNet [81], U-Net [82], DeepLab [83], [84], [85], PSPNET [86], etc. the following will introduce three classic segmentation networks: FCN, U-Net, and DeepLab, of which FCN and U-Net are commonly used in medical image segmentation. Nevertheless, the overall review of end-to-end tumor segmentation methods describes the tumor tissues, which include necrosis, active tumor, and peritumoral edema [26]. It also clearly discloses brain anatomy, end-to-end method scheme, classifications and comparisons.

#### 1) GENERATIVE ADVERSARIAL NETWORKS (GANS)

The GAN model is a generalized basic unsupervised learning model, and G and D are used as generative and discriminative models. The success of the GAN model in unsupervised image generation has driven the application of the model in other image processing fields, including unsupervised image domain conversion [87], [88], text to image [89], image fusion [90], etc. In addition, the GAN model is also widely used in medical image processing, such as image super-resolution reconstruction [91], region of interest location [92], image segmentation [93], etc. To obtain image features, DCGAN uses a deconvolution network as



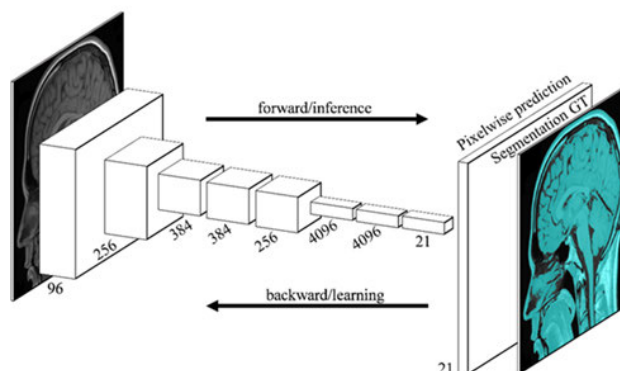
the generation network and a convolution network as the discrimination network, which enhances the expression ability of the network and can obtain high-quality generated images [94]. Similarly, LapGAN generates a series of images with different resolutions by sampling images and then uses multiple DCGANs to generate high-quality images from low resolution to high resolution [95]. However, LapGAN needs too many networks and is difficult to train. ProGAN generates low-resolution images to high-resolution images by iteratively increasing the network depth [96]. Compared with LapGAN, it generates high-quality images with many parameters reduced.

The proposed direct PET image reconstruction network by Wasserstein GAN (WGAN) model increases the image quality. Experiments were performed on mouse and patient data using a different method, and the results show that DPIR-NET enhances the reconstructed PET image quality more than DeepPET [97]. Generally, the image quality decreases due to a decrease in the radiotracer dose of positron emission tomography (PET) imaging. To enable the GAN model to generate images according to the given category, the conditional GAN (cGAN) model adds category conditions to the network when generating the network and determining the network input [98], [99]. To enable the discriminant network to learn the category features better, the proposed auxiliary classifier GAN (ACGAN) removed the conditions in the input of the discriminant network and added an output in the image category [100]. InfoGAN uses information entropy to comprehend unsupervised feature extraction of different types of images [101]. KL divergence is often used in GAN to measure the distance between the real data distribution and the generated data distribution, which leads to the discontinuity of the loss function at some points and the problem of mode collapse. To alleviate this problem, Arjovsky et al. proposed WGAN, which uses the Earth Mover distance instead of KL divergence to measure the difference between the two distributions [102]. Gulrajani et al. adopted a gradient penalty term instead of weight clipping to achieve Lipschitz restriction to solve the problem of parameter centralization caused by weight clipping of WGAN [103]. In addition, Mao et al. adopted the least square loss function, and WGAN can be considered a special case [104]. Lim et al. used hinge loss (HL) to measure the distance between two distributions [105]. HL is an extension of EM distance, which can make the GAN model training more stable.

## 2) FULLY CONVOLUTION NEURAL NETWORK (FCN)

In recent years, there have been more and more excellent models in the field of image segmentation. Long et al. proposed a fully convolution neural network (FCN), which is undoubtedly a classic network [80]. Unlike the traditional CNN, which uses the full connection layer to obtain the probability of a one-dimensional vector corresponding to many categories, FCN replaces the full connection layer with the convolution layer, outputs an image, and realizes end-to-end

training. Moreover, FCN can input images of any size, and the size of input and output images is consistent through the upsampling operation. Using a jump connection structure, the low-level and high-level feature information is added and fused point by point to ensure that more features can be learned.



**FIGURE 11.** FCN framework for pixel-wise prediction and semantic segmentation.

Based on the upsampling technology and multi-scale fusion method, the middle layer can be divided into FCN-32s, FCN-16s, and FCN-8s. FCN-32s directly outputs an image with the same size as the original image through the upsampling operation with a step size of 32, but loses a lot of detail information; FCN-16s performs two upsampling operations. First, the last layer is upsampled, and then a convolution operation of  $1 \times 1$  is added to the fourth pool layer. Then, the convolution results are fused with the upsampling results, and an upsampling operation is performed to restore the original image size. The network retains more detailed features; FCN-8s also samples up the output of the last layer, then samples up the fourth pool layer, and then integrates the results of the two upsampling with the third pool layer before upsampling, to obtain better results. However, FCN still has poor extraction of image detail information, and the processing of pixels is independent and lacks spatial consistency [106]. The authors have proposed a feature extraction and concatenation method, which would help the early detection of BT [107]. DensNet201 and Inception-v3 models are used to evaluate brain tumor detection and classification, resulting in 99.34% and 99.51% accuracies and achieving the highest performance in brain tumor detection. Figure 11 is the structural diagram of FCN.

## 3) SEGNET

SegNet is an improved model based on FCN, including encoding and decoding structures [81]. Compared with the pooling layer in other network structures, the advantage of SegNet is that the pooling layer has an index function, which is utilized to record the position of the max pooling result associated with the pooling core. The pool index connects the pooled layer output to the corresponding upper sampling layer. Since the network exploits the symmetrical structure,

the first pooling layer links to the last upsampling. In the encoding phase, each pooling operation will save the relative position of the weight selected by maximizing the  $2 \times 2$  filter. Then, in the decoding phase, the pooling operation is used to process these stored indexes and their corresponding feature maps. The decoding process uses zero filling volume, mainly used to fill the feature map information sampled on the sampling without generating parameters to be learned; it saves memory space and generates a sparse feature map. These feature maps are then convoluted to produce dense feature maps. Finally, the classifier outputs the maximum values of different categories through softmax to obtain the final segmentation graph. Salma et al. proposed FCNN SegNet to automatically segment brain tumors and parts of tumors by applying four different imaging modalities of MRI (i.e., T1, T1ce, T2, and Flair) [108]. The basic architecture of SegNet is shown in Figure 12.

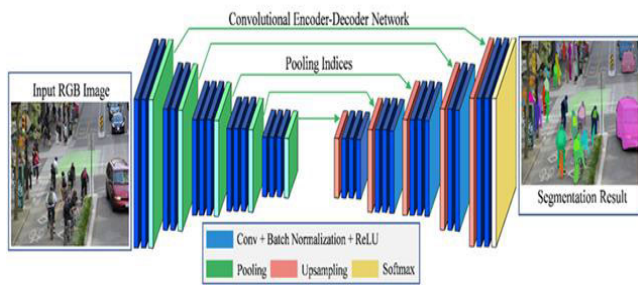


FIGURE 12. SegNet architecture for pixelwise segmentation [81].

4) U-NET

Unlike natural images, for medical images such as brain tumor imaging, even small marginal segmentation errors will significantly reduce the experience of DL automatic segmentation. The main reason for limiting the model to accurately segment this part of the difficult areas such as irregular contour, discontinuous organization, and fine-grained organization of the target is the ability of the model to make full use of shallow information and capture more details. To solve this problem, 3D U-net introduced a dual channel attention mechanism and applied it to the channel dimension and spatial dimension of the feature map through the channel attention and spatial attention branch, respectively, and embedded it into the layer hopping connection to alleviate the semantic gap caused by the direct integration of low-level features and high-level features, also to utilize the efficiency of low-level features.

The U-Net model adopts the encoding-decoding method, as shown in Figure 13 [82]. The encoding method is to continuously obtain the characteristics of the input image through the convolution layer and the lower sampling layer. With the increase in the number of layers, the image features extracted by the coding layer are more and more abstract, and the feature resolution is decreasing. Unlike FCN, which uses these features for prediction results, U-Net uses decoding

to generate high-resolution (fine-grained) features from low-resolution (coarse-grained) features. Finally, U-Net classifies high-resolution features to obtain segmentation results with the same input resolution. As shown in Figure 13, U-Net adopts a multilayer downsampling layer during encoding. However, the downsampling layer will lose image information, which leads to the lack of image information for the features after upsampling during decoding. In order to resolve this problem, u-net uses a horizontal connection to connect the encoded features with the decoded features. In U-Net, the upper sampling layer is used to upgrade the dimension of features. It can be seen that the deconvolution layer also has the same ability and is more flexible than the upper sampling layer. Therefore, the deconvolution layer can replace the upper sampling layer.

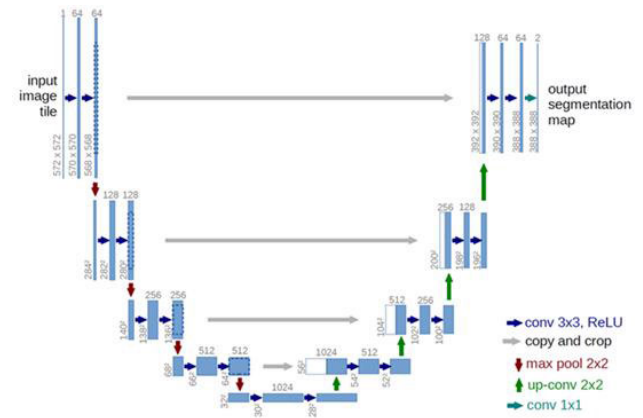


FIGURE 13. U-Net architecture [82].

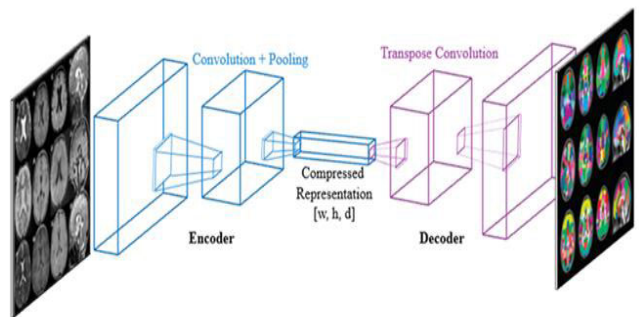


FIGURE 14. DeepLab architecture for image segmentation.

5) DEEPLAB MODELS

The DeepLab model is also an encoding-decoding method. Its coding method also adopts convolutional down sampling. DeepLab also adds a residual layer to better extract features and build a deeper network. In the network, too many downsampling layers will lead to image information loss. However, the downsampling layer can increase the receptive field of the network and can obtain global features. In order to ensure that the network has a larger receptive field without

reducing the feature resolution, DeepLab cancels the last two lower sampling layers and uses the Atrous Spatial Pyramid Pooling (ASPP) module to segment targets at different scales. The ASPP module uses cavity convolution to obtain larger receptive fields. At the same time, it uses cavity convolution with different expansion rates to obtain receptive fields of different sizes and then combines these features to obtain pyramid features. The decoding method of the DeepLab model also adopts the upsampling plus convolution method and introduces the horizontal connection.

However, unlike the U-Net method, DeepLab finds that the 4-fold upsampling is better [109]. Later on, the authors proposed DeepLab v1, which introduced the variation in pooling stride and padding size, and this model was based on the VGG network [83]. In DeepLab v1, conditional random fields (CRFs) are introduced to enhance the segmentation accuracy [110]. Consequently, DeepLab v2 was proposed to resolve the segmentation problem caused by the same object scale variations in the same image [109]. DeepLab v3 utilized ResNet-101 with a cascaded Atrous module to get promising image segmentation at multi-scales [84]. The authors have utilized the cascaded network of two DeepLab v3+ [85] with ResNet-50 to get improved segmentation results [111]. The DeepLab architecture for image segmentation is shown in Figure 14.

## VI. IMAGE GUIDED SURGERY FOR BRAIN TUMORS

### A. STEREOTACTIC SURGERY

In the form of image-guided surgery (IGS), stereotactic surgery uses several imaging modalities, including CT and MRI, to produce a precise map of the brain [142], [143]. The surgeon then used this map to precisely direct a surgical tool, such as a drill or biopsy needle, to the tumor's position. This method seeks to precisely target the tumor while causing the least amount of harm possible to nearby healthy brain tissue. In order to keep the patient's head motionless throughout the treatment, the patient is first placed in a unique head-holding equipment [144], [145]. Next, a CT or MRI scan of the patient's head can be performed to get precise images of the patient's brain. A 3-dimensional map of the brain can be made using MRI scans which helps the surgery. A frameless stereotactic device used by the surgeon throughout the process, confers to the head-holding device and guides the surgical tools to the exact position of the tumor. The surgeon can sample or remove the tumor using the instruments. Diverse types of brain tumors, including benign and malignant tumors, can be treated with IGS. It is particularly beneficial for brain tumors that are located in hard-to-reach regions, like deep-seated.

### B. FLUORESCENCE-GUIDED SURGERY

A specific dye can be placed into the patient's blood circulation during fluorescence-guided surgery (FGS), which uses IGS to target the tumor only [28], [146]. The fluorescence can be detected by specific cameras, which enable the surgeon to

see the tumor and surrounding structures during the surgery. The risk of harming the surrounding healthy brain tissue can be reduced using this method though increasing the precision and accuracy of brain tumor surgery. The process initiates with controlling a specific dye to the patient. Tumor cells favorably absorb this dye, which causes them to glow when struck by light. An exclusive camera detects the fluorescence scans and develops detailed brain images. The surgical procedure can be planned using these images. The specialist uses the fluorescence images to lead the surgical instruments to the precise spot of the tumor. This aids in reducing the chance of destruction of healthy brain tissue by extending the specialist's capability.

Gliomas, a form of brain tumor known to invade surrounding brain tissue, have been found to respond particularly well to FGS [147]. These tumors can be challenging to remove with conventional surgical methods. The probability of leaving any tumor cells behind after surgery can be reduced by FGS. It has been discovered that the method is efficient and safe, with a high degree of precision.

### C. INTRAOPERATIVE MRI (iMRI)

Intraoperative MRI (iMRI) is a type of IGS that uses an MRI machine in the operating room to give the physician real-time brain imaging while executing surgery [148]. This technique intends to improve the accuracy and safety of brain tumor surgery by giving the physician instant access to the location and size of the tumor. Using MRI, detailed brain scans can be taken prior to the treatment. These images guide the surgical tools according to the procedure. The physician can confirm the surgery's position and extent using real-time images.

Cancers located in difficult-to-reach parts of the brain, such as deep-seated tumors or tumors close to critical functioning areas, have been found to benefit particularly from iMRI [149]. This systematic procedure has proven to be quite accurate and rarely problematic. It also enables the physician to confirm that the tumor has been completely removed. The primary benefit of iMRI is that it provides the physician with real-time imaging during the procedure, to enhance the accuracy and safety. iMRI is not yet widely used because it is comparatively a new technique.

### D. IMAGE-GUIDED RADIATION THERAPY (IGRT)

Using imaging tools like CT or MRI, image-guided radiation treatment correctly targets and carries radiation to the brain tumor while limiting exposure to the surrounding tissues [150], [151]. This method intends to enhance the precision and efficacy of radiation therapy for brain malignancies. Initially, the patient undergoes imaging, such as CT or MRI scan, to get accurate brain images. During therapy, the patient is positioned on the treatment table, and the position of their body is tracked by specialized imaging tools. A linear accelerator generates high-energy x-rays or particles and can be utilized to guide the radiation beams at the tumor. Imaging equipment can be employed to improve the accuracy.

TABLE 2. Comparison of DL methods for BTS.

DL Method	Model Architecture	Segmentation Task	Advantages	Disadvantages
Attention-Gated Network (AGN) [112][113][114][115]	U-Net with attention gates	Whole tumor, tumor core, enhancing tumor	High accuracy, robust to noise	Requires large amounts of data
Hybrid Attention Network (HAN) [116][117][118][119]	U-Net with hybrid attention mechanism	Whole tumor, tumor core, enhancing tumor	Efficient training handles complex tumor shapes	Limited ability to generalize to other datasets
Multi-Scale Attention U-Net (MSAU-Net) [120][121][122][123]	U-Net with multi-scale attention modules	Whole tumor, tumor core, enhancing tumor	High accuracy, captures long-range dependencies	Increased computational complexity
Conditional Random Field (CRF) with U-Net [124][125][126][127][10]	U-Net with CRF post-processing	Whole tumor, tumor core, enhancing tumor	Improved spatial consistency, reduces segmentation errors	Requires additional training for the CRF
DeepLabV3+ and ASPP [85][128][129][130]	DeepLabV3+ and atrous spatial pyramid pooling	Whole tumor, tumor core, enhancing tumor	Efficient for large datasets, handles different tumor sizes	Less accurate than some U-Net based methods
Swin Transformer [131][132][133][134][135]	Swin Transformer with multi-level attention	Whole tumor, tumor core, enhancing tumor	State-of-the-art performance handles complex tumor shapes and pathologies	High computational cost, requires large amounts of data
Generative Adversarial Network (GAN) [136][137][138][139][140]	GAN-based model for generating synthetic brain tumor images	Whole tumor, tumor core, enhancing tumor	Improved data augmentation, generates realistic synthetic data	Requires careful training to avoid artifacts, limited control over specific tumor characteristics
SegNet [81][108][141]	SegNet with encoders and decoders	Whole tumor, tumor core, enhancing tumor	Efficient architecture, easy to implement	Lower performance compared to newer methods

It has been discovered that IGRT is particularly helpful for brain cancers which are located in hard-to-reach regions of the brain. This method is highly accurate and it enables the tumor to get a larger radiation dose while reducing exposure to the surrounding healthy regions. The benefit of IGRT is that it enables the radiation oncologist to have an eye on the tumor's location and change the radiation beams as necessary to ensure that the tumor receives the right amount of radiation while minimizing exposure to the surrounding tissues [152]. Radiation oncologists, technologists, and radiologists have been trained in the tools and image interpretations are needed as part of a specialized team.

### E. AUGMENTED REALITY-BASED SURGERY

Augmented reality (AR) is being employed more and more in the field of neurosurgery to remove brain tumors [153], [154]. Real-time, 3-dimensional (3D) scans of a patient's anatomy can be sent to the physician using AR. As the brain is a fragile, complex organ that can be exciting to explore, AR can be exclusively helpful. The physician's ability to view the tumor and surrounding anatomy in 3D can increase surgical precision and accuracy. Since it can be difficult for the physician to see the tumor and surrounding tissues with normal 2D imaging techniques.

AR can be used to aid surgical planning and reduce human errors. For instance, the physician can utilize AR to construct a virtual map of a patient's brain [155]. It may be beneficial to

reduce the risk of destruction and improve surgical accuracy. AR can help to lessen the need for numerous surgical passes, which can shorten the surgery's duration. There are already a variety of systems available for AR-based brain tumor surgery, each with exceptional high-tech capabilities [156]. AR is progressively being used in neurosurgery, and the advantages of employing AR in brain tumor surgery include increased precision and accuracy, better surgical planning, and a shorter anesthetic stay. AR will play a bigger role in the treatment of brain tumors and other neurological diseases as technology advances.

### F. ULTRASOUND-GUIDED SURGERY

Brain tumors can be found and removed with the help of ultrasound-guided surgery (UGS) [157]. High-frequency sound waves are used in this procedure. They are sent to the tumor and then returned to a computer screen. The physician can use this information to lead the surgical tools precisely to the tumor's location. Real-time brain imaging is the primary benefit of UGS. Previously, physicians were forced to rely on preoperative imaging in a standard tumor surgery, such as MRI or CT scans, which could be less accurate and could fail to pinpoint the tumor's specific location. The benefit of UGS being less intrusive than conventional brain surgery. UGS is more precise and with less harm to the surrounding tissues. This phenomenon may result in fewer difficulties and quicker recovery times. Transcranial ultrasonography,

or TCUS, is the ultrasound technology used in this procedure [158]. With the use of this technology, deep-seated lesions like brain tumors can be seen in real-time. A computer analyzes and compiles the echoes formed by the tumor.

UGS is continuously being studied as a relatively new procedure [159]. However, early research proposes it may help treat brain malignancies. In one study, researchers discovered that when compared to conventional brain surgery, UGS resulted in a very high rate of total tumor removal [160]. Patients who underwent surgery with UGS also experienced a shorter hospital stay. Moreover, there are several restrictions with UGS. For larger tumors or cancers found in specific regions of the brain, for instance, it might not be suitable.

## VII. DISCUSSION

The multimodal MRI brain tumor image segmentation is to segment the whole tumor region, core tumor region, and enhanced tumor region from normal brain tissue, using the image data of different imaging modalities. Accurate segmentation of multimodal MRI brain tumor images with AI has great significance in clinical diagnosis. It can reduce much time for doctors to manually divide brain tumor areas, make doctors pay more attention to treating patients and saving people, and improve doctors' medical intellectual level. Therefore, it has always been an important topic in medical image processing. With the continuous development of modern medical imaging, the traditional manually labeled MRI image segmentation is gradually replaced by computer-aided diagnosis. Accurate and rapid diagnosis not only helps doctors make judgments in a short time but also reduces the error of manual markings. As a common malignant tumor, early non-invasive diagnosis is very helpful for treating patients. However, due to the differences in the appearance and shape of brain tumors among different patients, evaluating tumor areas is very time-consuming. Therefore, the automatic segmentation based on DL is very attractive in this case because it can describe the relevant tumor parameters more objectively and accurately. Databases with complete details provide easy and quick access to specific and significant studies [29], [36]. Similarly, the pros and cons of overall categorized methods into thresholding, traditional machine learning, region-based, DL variants, and hybrid approaches can be found in [37], [161], [162], and [163]. Suppose the multi-scale information helps the network segment some easily confused areas at the edge of brain tumors. In that case, two branches are added to the original single-scale confrontation network to form a parallel multi-scale segmentation network. Stacked residual blocks and attention mechanisms are introduced to improve the segmentation accuracy of the model. The residual module can alleviate the problems of gradient dispersion and network failure in the process of deepening the network depth, accelerate the convergence process of the network, and allow the network to extract the characteristic information of data at different scales simultaneously in combination with the multi-path structure. The feature information is used by multi-scale fusion. It focuses

limited attention on the details of the lesion area, which helps to improve the sensitivity of the model and improve the accuracy of fine segmentation of brain tumor areas.

The progress of semantic segmentation—from traditional approaches to sophisticated DL techniques—is highlighted, with a particular emphasis on pixel-level classification [164]. The authors cover both supervised and unsupervised learning algorithms, emphasizing the vital significance of DL in tackling important issues. It also offers an extensive examination of surveys devoted to semantic segmentation in the context of DL. BTS has made rapid developments with the development of DL. The major advantage of CNN is that its hidden layer automatically learns image features and iterates features through the connection between layers. However, CNN has the problem of learning blindness. The design of network depth and structure is very important for the speed and accuracy of feature extraction. Several DL methods have been discussed, which are the most significant and outperformers compared to all earlier models, as shown in Table 2. The dice scores are calculated in the literature based on three categories: tumor core, whole tumor, and enhancing tumor, and we have quoted the mean values of the whole tumor from test data. A couple of articles have provided accuracy (i.e., given in %) instead of dice scores, which can be observed in Table 2. Their associated dice scores cannot predict the performance of the given methods because these values vary with respect to test and training data, and it is recommended that the complete article is read for a better understanding.

### A. LIMITATIONS AND FUTURE PERSPECTIVES

For BTS, DL methods have been frequently used for the last decade, which involve multiple layers and many steps in the computer vision algorithms to comprehend the intensity and symmetry-related information. All these properties are combinedly and used for the classification of different regions of tumors such as necrosis, oedemic, gliomas, enhancing or non-enhancing tumors, etc. Frequent use of AI still needs many validations, specifically clinical and biological validations [165]. Firstly, there are some limitations to getting such validations in which the data is a key limitation. In several computer vision scenarios, we are improving the application outcomes by utilizing our models and with the help of non-related datasets. Therefore, various techniques have been developed to overcome this limitation, such as data augmentation, transfer learning, etc. Logically, data augmentation techniques should be avoided for sensitive and deadly scenarios. As we know, several types of tumors are different in shape, location, and size. Boundaries of tumors are irregular, discontinuous, and unclear, so it is better to develop a cumbersome dataset exclusively for one specific disease, which can better assist computer vision approaches. Nevertheless, BraTS development has lessened this limitation dramatically by introducing multiple imaging for BTS [166]. Secondly, biological validation reveals that images cannot interpret the biological structures because tumor lesions can

TABLE 3. Description of DL models with their dice scores and training data.

Sr. No.	Model Name/Backbone	Papers Description (code link)	Dice Score	Extra Training Data	Year
1	CNN [161]	Automated segmentation based on CNN that explores small 3×3 kernels. The small kernels let scheming the deeper network, in addition to prohibiting overfitting, and adapt to the smaller amount of weights. Intensity normalization is utilized at the pre-processing stage. (NA)	0.88	No	2016
2	Cascaded CNN [79]	Automatic BTS is conferred by DNN and utilizes local and global features. ( <a href="https://github.com/jadevaibhav/Brain-Tumor-Segmentation-using-Deep-Neural-networks">https://github.com/jadevaibhav/Brain-Tumor-Segmentation-using-Deep-Neural-networks</a> )	0.88	No	2017
3	3D CNN + CRF [167]	It is based on 11-layers deep structure, multi-scale, 3D CNN architecture and a robust training process. ( <a href="https://github.com/deepmedic/deepmedic">https://github.com/deepmedic/deepmedic</a> )	0.85	No	2017
4	CNN + 3D filters [168]	A CNN with 3D filters is used for brain MRI and also applied to hand bones. ( <a href="https://github.com/BRML/CNNbasedMedicalSegmentation">https://github.com/BRML/CNNbasedMedicalSegmentation</a> )	0.85	Yes	2017
5	Cascaded Anisotropic CNNs [51]	A cascade of FCN is developed and introduces a hierarchy of subregions (i.e. tumor categories). ( <a href="https://github.com/hellopipu/brats17">https://github.com/hellopipu/brats17</a> )	0.87	No	2017
6	U-Net + more filters + data augmentation + dice-loss [169]	A U-Net based model and a dice loss function are employed to improve the performance and overcome overfitting. ( <a href="https://github.com/shalabh147/Brain-Tumor-Segmentation-and-Survival-Prediction-using-Deep-Neural-Networks">https://github.com/shalabh147/Brain-Tumor-Segmentation-and-Survival-Prediction-using-Deep-Neural-Networks</a> )	0.85	No	2018
7	AFN-6 [170]	An autofocus convolutional layer with the capability of multi-scale processing generates strong features that work adaptively for semantic segmentation. ( <a href="https://github.com/yaq007/Autofocus-Layer">https://github.com/yaq007/Autofocus-Layer</a> )	0.84	No	2018
8	Att U-Net [113]	The attention gate model is proposed which observes the varying shapes. It eliminates the modules for organ localization in cascaded CNN. ( <a href="https://github.com/sfczekalski/attention_unet">https://github.com/sfczekalski/attention_unet</a> )	0.84	No	2018
9	NVDLMED [171]	An encoder-decoder-based semantic segmentation network is proposed for subregion segmentation in 3D MRI. ( <a href="https://github.com/IAmSuyogJadhav/3d-mri-brain-tumor-segmentation-using-autoencoder-regularization">https://github.com/IAmSuyogJadhav/3d-mri-brain-tumor-segmentation-using-autoencoder-regularization</a> )	0.87	No	2018
10	ModelGenesis [172]	This model named Models Genesis is created using self-supervision, without labeling and application-specific models for segmentation and classification problems. ( <a href="https://github.com/MrGiovanni/ModelsGenesis">https://github.com/MrGiovanni/ModelsGenesis</a> )	0.925	No	2019
11	KMFCM [173]	A hybrid energy-efficient uses active contours K-means and fuzzy C-means to segment tumors. (NA)	0.82	Yes	2019
12	ECNN [174]	An enhanced CNN model with an optimized loss function is developed for automatic segmentation. (NA)	0.90	No	2019
13	DCNN [175]	It uses DL-based 3D CNN and multimodal MRIs for BTS. The potent features are selected from segmented regions using a cross-validation decision tree for survival prediction. ( <a href="https://github.com/shalabh147/Brain-Tumor-Segmentation-and-Survival-Prediction-using-Deep-Neural-Networks">https://github.com/shalabh147/Brain-Tumor-Segmentation-and-Survival-Prediction-using-Deep-Neural-Networks</a> )	0.90	No	2019
14	Bag of tricks [176]	It categorizes tumor segmentation tricks into semi-supervised, patch-size training, and data sampling. It also elaborates on optimization methods. (NA)	0.88	No	2019
15	OM-Net + CGAP [177]	The authors proposed a light weight deep network to resolve class imbalance compared to cascaded models. It integrates joint and discriminative features into one model. ( <a href="https://github.com/chenhong-zhou/OM-Net">https://github.com/chenhong-zhou/OM-Net</a> )	0.92	No	2020
16	MS-Dual-Guided [178]	This model overcomes the drawbacks of the encoder-decoder network by utilizing contextual dependencies and self-attention. ( <a href="https://github.com/sinAshish/Multi-Scale-Attention">https://github.com/sinAshish/Multi-Scale-Attention</a> )	0.80	No	2020
17	Semantic Genesis [179]	To semantically enrich information, self-classification, self-discovery, and self-restoration are introduced to improve segmentation. ( <a href="https://github.com/fhaghi/SemanticGenesis">https://github.com/fhaghi/SemanticGenesis</a> )	68.8%	No	2020
18	Feature Concatenation [107]	It introduces extraction concatenation of multi-level features to diagnose tumors at an early stage. The DensNet201 and Inception-v3 pretrained models are used for validation. (NA)	99.51%	No	2020

TABLE 3. (Continued.) Description of DL models with their dice scores and training data.

19	ELM and RELM-LOO [180]	The images have been enhanced by fuzzy median filters which help to segment accurately using fuzzy set approach. (NA)	0.96, 0.87	No	2020
20	DeepSeg [181]	A decoupling framework is developed in an encoder-decoder way. A CNN provides spatial information in the encoder part. ( <a href="https://github.com/razeineldin/DeepSeg">https://github.com/razeineldin/DeepSeg</a> )	0.84	No	2020
21	U-Net and CNN based Model [182]	The authors employed U-Net to segment tumors; pretrained Vgg16 using transfer learning and a classifier are used to grade tumors. (NA)	0.84	Yes	2020
22	nnU-Net [183]	Several modifications are introduced in BraTS to improve segmentation such as postprocessing, data augmentation, region-based training and minor variations to nnU-Net. ( <a href="https://github.com/MIC-DKFZ/nnUNet">https://github.com/MIC-DKFZ/nnUNet</a> )	0.89	No	2020
23	Extension of nnU-Net [184]	3D U-Net based model is trained by utilizing patch strategies and normalization for BTS. ( <a href="https://github.com/woodywff/brats">https://github.com/woodywff/brats</a> )	0.89	No	2020
24	Segtran (i3d) [185]	A transformer-based model is developed to overcome the limitation of receptive field which is not addressed in the earlier models. The squeezed attention can regularize the self-attention and the expansion block can learn diversified representations. ( <a href="https://github.com/askerlee/segtran">https://github.com/askerlee/segtran</a> )	0.82	No	2021
25	TransBTS [186]	The transformer is utilized in 3D CNN to segment brain tumors. This framework is analogous to the encoder-decoder network. ( <a href="https://github.com/Wenxuan-1119/TransBTS">https://github.com/Wenxuan-1119/TransBTS</a> )	0.90	No	2021
26	DL and attention mechanism [187]	The region-oriented preprocessing is proposed in this paper. It ends overfitting and decreases computational time. (NA)	0.92	No	2021
27	CMFT and CMFF [188]	The cross-modality feature transition and fusion are introduced in the DL framework to enhance the feature representation. (NA)	0.90	No	2021
28	MRA-UNet [126]	Multiscale residual attention-UNet preserves sequential information. It exploits cascaded type multiscale learning to employ adaptive ROIs to segment tumor regions precisely. (NA)	0.90	No	2022
29	DCNN [189]	A DCNN based model is proposed which introduces a dynamic loss function to significantly consider tumor sub-regions. Morphological relationship and symmetric attention are utilized to lessen boundary constraints and highlight locations of tumors, respectively. (NA)	0.916	No	2022
30	MFF-DNet [190]	A dual path framework based on multimodal feature fusion is proposed which uses kernel multiplexing to comprehend perceptual domain and nonlinear mapping features (i.e. to enhance coherence). (NA)	0.92	No	2022
31	SwinBTS [191]	SwinBTS blends edge and semantic data to produce precise segmentation. The method, which uses multimodal MRI scans to segment brain tumors, makes use of 3D context and performs at the cutting-edge level on benchmark datasets. (NA)	0.89	No	2022
32	TBraTS [192]	A trusted BTS network that offers trustworthy uncertainty estimations and strong segmentation outcomes. It does so without placing an undue strain on the backbone network's computing resources. ( <a href="https://github.com/Cocofeat/TBraTS">https://github.com/Cocofeat/TBraTS</a> )	--	No	2022
33	D <sup>2</sup> -Net [193]	D2-Net with missing modalities consists of a modality disentanglement stage (MD-Stage) and a tumor-region disentanglement stage (TD-Stage). The MD-Stage utilizes a spatial-frequency joint modality contrastive learning scheme to separate modality-specific information from MRI data. ( <a href="https://github.com/CityU-AIM-Group/D2Net">https://github.com/CityU-AIM-Group/D2Net</a> )	0.72	No	2022
34	Transformer [194]	It consists of an edge detection module using CNNs, an edge spatial attention block (ESAB) for feature improvement, and a semantic segmentation module using the Swin Transformer and a shifted patch tokenization technique. ( <a href="https://github.com/HXY-99/brats">https://github.com/HXY-99/brats</a> )	0.91	No	2023
35	DPAFNet [195]	Dual path convolution is used by DPAFNet to increase network scale and residual connections are included to prevent degradation. At the channel level, it introduces an attention fusion module that integrates global and local information. DPAFNet improves the semantic information in the collected features by combining feature maps of various scales. (NA)	0.90	No	2023

**TABLE 3.** (Continued.) Description of DL models with their dice scores and training data.

36	U-Net [196]	U-Net-based network, redundant filters are found using a genetic approach. Filter pruning is viewed as a multiobjective optimization issue that concurrently seeks to optimize performance and inference time. The compressed network is then used for the segmentation of brain tumors. ( <a href="https://github.com/FBehrad/Evo_conv">https://github.com/FBehrad/Evo_conv</a> )	0.87	No	2023
37	CKD-TransBTS [197]	Based on MRI imaging principles, it divides input modalities into two categories, utilizing radiologists' experience in identifying brain cancers from a variety of modalities. The model combines the advantages of the Transformer and CNN to pinpoint precise lesion borders and collect long-range features in 3D volumetric images. (NA)	0.93	No	2023
38	Federated Learning [198]	Federated learning-based model permits collaborative learning by training the model with distributed data. Due to the scalability of this proposed model, it can be applicable to multiple medical imaging scenarios.	0.89	No	2023
39	Handcrafted features + CNN [199]	The proposed approach utilizes handcrafted features and CNN network to improve BTS performance. The features are based on intensity, shape, and texture from MRI scans.	0.91	No	2023

grow differently after removing them, and tumor invasions cannot be observed by AI or imaging techniques. Some limitations in the field of BTS are listed below:

- Lack of training datasets for DL methods
- Sophisticated segmentation techniques are mandatory when applying for annotations or changing structured labels
- 3D segmentation models have been implemented using 2D segmentation [200].
- The training of the DL system on BraTS dataset needs consideration to handle uncertainty and noise.
- The small receptive field for a large dataset, then deep models are not worthwhile. Training a network has various constraints, such as limited memory, GPU, and bandwidth.
- Fixed-size for a kernel for image slicing may damage some valuable information.
- Data augmentation (i.e., scaling and rotation) and normalization approaches have been used to develop new lesions of brain tumors, which may generate class imbalance [161].

In medical image segmentation, especially the end-to-end image segmentation algorithm, there is a critical problem of data imbalance. Although scholars have proposed many methods to alleviate the data imbalance problem, such as using data enhancement, reducing image size, changing the loss function of network training, etc., these methods cannot solve the problem. In the future, we need to explore effective methods to make it easier for the network to learn the characteristics of a small number of samples in brain medical image segmentation and improve the segmentation accuracy of the network. Currently, the main algorithm of medical image segmentation is still supervised. Still, the supervised algorithm has high requirements for input data, and the acquisition of label data requires a lot of human resources, which leads to high-cost requirements for this algorithm and makes it unsuitable for the current situation in

the medical field. Using weakly supervised or unsupervised algorithms to realize medical image analysis is an important direction in the future. The commonly used brain medical image segmentation algorithms based on neural networks are derived from natural image processing algorithms. However, brain medical images differ from natural images, and there is a large number of prior knowledge of medical anatomy in brain medical images. This prior knowledge helps to enhance the segmentation performance of the network, but the existing brain medical image segmentation based on the neural network often ignores this point. Therefore, in the future, it is necessary to combine neural networks with medical anatomy knowledge to enhance the segmentation performance of the network.

## VIII. CONCLUSION

This paper delineates the significant contributions of recent studies for BTS. The key role of this article is to provide an understanding of selecting robust methods for competent segmentation and future studies, which will help doctors get concrete disease identification. However, we have also grouped the BTS methods with respect to their characteristics. Several limitations related to datasets, algorithms, state of the arts, etc., have been discussed briefly to fascinate researchers for deep studies. Furthermore, we have presented various optimal methods with their dice scores, training data, and published time with references. Neurologists persist in discussing that the datasets are limited for several diseases individually, which can be helpful for computer-based diagnosis systems, and that can be overcome by centralizing the systems of hospitals. Several hospitals around the globe might have limited datasets individually, but we can devise and assist in benchmarking the datasets in the medical field, which can be a good prospect. DL approaches outperform conventional methods due to their limitations. The recent models can be further improved with ensemble and data augmentation procedures.



## DATA AVAILABILITY

The data are available from the corresponding author upon reasonable request.

## CONFLICT OF INTEREST

The authors declare that there are no conflicts of interest regarding the publication of this paper.

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