DefED-Net: Deformable Encoder-Decoder Network for Liver and Liver Tumor Segmentation

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Abstract-Deep convolutional neural networks have been widely used for medical image segmentation due to their superiority in feature learning. Although these networks are successful for simple object segmentation tasks, they suffer from two problems for liver and liver tumor segmentation in CT images. One is that convolutional kernels of fixed geometrical structure are unmatched with livers and liver tumors of irregular shapes. The other is that pooling and strided convolutional operations easily lead to the loss of spatial contextual information of images. To address these issues, we propose a deformable encoder-decoder network (DefED-Net) for liver and liver tumor segmentation. The proposed network makes two contributions. The first is that the deformable convolution is used to enhance the feature representation capability of DefED-Net, which can help the network to learn convolution kernels with adaptive spatial structuring information. The second is that we design a Ladder-atrous-spatial-pyramid-pooling module using multi-scale dilation rate (Ladder-ASPP) and apply the module to learn better context information than the atrous spatial pyramid pooling (ASPP) for CT image segmentation. The proposed DefED-Net is evaluated on two public benchmark datasets, the LiTS and the 3DIRCADb. Experiments demonstrate that the DefED-Net has better capability of feature representation as well as provides higher accuracy on liver and liver tumor segmentation than stateof-the art networks. The available code of DefED-Net we propose can be found from https://github.com/SUST-reynole/DefED-Net.

Index Terms—image segmentation, deep learning, U-Net, deformable convolution, Ladder-ASPP

I. INTRODUCTION

L IVER cancer is one of the most common and most lethal cancers in the world, which threatens life and health of humans seriously [1, 2]. In the clinical context a

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liver is a common site for both primary (e.g., hepatocellular carcinoma) or secondary (e.g., hepatic metastases due to colorectal cancer) tumor development. Accurate liver and liver tumor segmentation from enhanced abdominal CT images can help doctors to assess the function of livers and make a decision for disease diagnosis and treatment. However, as livers have a similar density with other neighboring organs and the liver tumors show very low contrast and serious intensity inhomogeneities in abdominal CT images, it is difficult to find accurate liver and liver tumor boundaries depending on human vision [3]. Manually labeling liver and liver tumor areas not only suffers from subjective judgment and limited accuracy, but also is tedious and inefficient. Therefore, semiautomatic or fully automatic approaches for liver and liver tumor segmentation have been a research goal in the field of medical image analysis to help in clinical applications [4].

Before the advent of deep learning techniques [5], liver and liver tumor segmentation were often semi-automatic and they mainly relied on image segmentation algorithms based on model-driven such as region growing [6], active contour models [7], graph cut [8], shape statistical models [9], etc. These approaches can be roughly categorized into three groups: pixel-based approaches, graph-based approaches, and contourbased approaches. The first type of approach mainly includes thresholding and region merging. The type of approach only achieves low segmentation accuracy for liver and liver tumor segmentation due to the employment of low-level features and limited capability of model representation. Graph-based approaches show clear superiority than pixel-based approaches, since they employ the max-flow/min-cut algorithm to find a minimum-cost closed set [10]. This kind of semi-automatic approach can achieve accurate liver segmentation by simply labeling the foreground and background, and it does not even require the iterative operation [11]. However, image segmentation results are easily influenced by labeling results, and graph cuts require high computational cost for high-resolution images since each pixel of images is viewed as a note [12]. Consequently, researchers often employ the combination of graph cuts and other algorithms, such as watershed [13], shape constrain [14], multi-scale registration [15], etc., to improve the segmentation accuracy and computational efficiency for liver and liver tumor segmentation.

Compared to the first two kinds of approaches, contourbased liver and liver tumor segmentation attracts more researchers' attention since they can provide better segmentation results using curve or shape evolution. Level-set [16] is one of the most popular algorithms in medical image segmentation, since the level-set utilizes energy optimization to evolve a given curve into the real boundaries of objects. A large number of improved level-set algorithms have been proposed by introducing partial differential equations into the evolution process to improve the convergence speed and segmentation accuracy [17-19]. The statistical shape model [20] is another popular algorithm for contour-based liver and liver tumor segmentation. Different from the level-set, this algorithm often constructs firstly a training set of liver and liver tumor shapes, and then employs machine learning algorithms, such as the random forest [21, 57, 58], support vector machine (SVM) [22] and adaboosted histogram [59] to learn an effective classifier. As a result, each liver and liver tumor shape can be represented by some corresponding patches from liver and liver tumor surface in the training set [23, 24]. The advantage of this kind of approach is that they can provide better segmentation results than unsupervised approaches, but the disadvantage is that segmentation results depend on the selection of training set and classifiers. Although numerous algorithms have been proposed for liver and liver tumor segmentation, they only provide good segmentation results for some slices with clear liver or tumor boundaries, and they are often unavailable for slices with blurred liver contour or intensity inhomogeneities tumors in practical applications. Ren et al. [54] proposed an automatic framework for atlasbased multiorgan segmentation in abdominal dynamic PET images with three different methods (4D-pair, 4D-PCA, and 3D), incorporating probabilistic atlas information into the segmentation as a spatial prior using maximum a posteriori (MAP) estimation. This provides a powerful and reliable region of interest (ROI) for dynamic abdominal PET multi-organ segmentation for better segmentation results. Although atlasbased segmentation can easily capture anatomical variation and thus offers higher segmentation accuracy, it suffers from a clear shortcoming of ravenous appetite for computational resources because analyzing, manipulating, and processing all atlases typically demands a substantial amount of memory and time. It is believed that this is one of the main reasons why atlas-based segmentation has not been widely used in clinical applications.

In recent years, with the rapid development of deep learning [25] in the field of computer vision, researchers prefer to use fully convolutional neural networks (FCN) [26] to achieve image semantic segmentation in an end-to-end way [27]. These networks usually adopt multi-level encoder-decoder structures, and the encoder and decoder are often composed of a large number of standard convolutional or deconvolutional layers. In addition, there is a residual or long-range connection between encoders and decoders. This kind of design can automatically remove insignificant features and maintain interesting features through the contraction and expansion paths; it can also achieve the fusion of low-level and high-level features. Compared with FCN, U-Net [28] proposed by Ronneberger et al. obtains great success for medical image segmentation, since the encoder and decoder of U-Net are perfectly symmetrical and upsampling gradually makes it possible to obtain finer segmentation results. Since then, researchers focused on the improvements of U-Net [55, 56]. The most common way

is to use the backbone of classic convolutional neural networks with pre-trained parameters such as VGG [29], ResNet [30], DenseNet [31] and GhostNet [50] etc. to replace the encoder achieving transfer learning [60]. The other popular way of improving U-Nets is to add attention mechanisms [32] between encoders and decoders to focus on interesting regions, such as attention U-Net [33] and RA-UNet [34]. To exploit further potentially useful information in feature maps, R2-UNet [35] introduces recurrent convolution that is able to extract features using the same layer many times. UNet++ [36] employs U-Nets with different depths instead of long-range connections to avoid the rough fusion of lowlevel and high-level features. Recently, mU-Net [37] believes that small targets may disappear after pooling since the skip connection in U-Net repeatedly processes low-resolution feature information. Therefore, mU-Net achieves better liver and liver tumor segmentation by adding a residual path with deconvolution and activation operations to the skip connection of U-Net. These improved 2D networks not only show better performance in medical image segmentation, but also achieve simpler design of data augmentation schemes while keeping the lower memory requirement than 3D networks. However, they cannot capture the spatial information along the z-axis due to the employment of 2D convolution kernels, which may degrade the performance in volumetric segmentations.

To extract the spatial information along the third dimension, Ji et al. [38] employed 3D convolution kernels to achieve 3D CNN, which makes it possible to process 3D volume data directly. Based on 3D CNN and U-Net, Cicek and Milletari et al. proposed 3D U-Net [39] and V-Net [40], respectively. The V-Net applies 3D convolutions together with residual connection to the feature encoder stage, and deepens the network depth to obtain better segmentation results than 3D-UNet. Furthermore, by introducing the strategy of depth supervision, both Med3D [41] and 3D DSN [42] achieve faster and more accurate segmentation of volumetric medical images. More application of 3D CNNs can be seen in [43]. Although these 3D networks can simultaneously explore the spatial information of interslice and inner-slice, these networks suffer from some new problems such as more parameters, much memory usage, and much narrow reception fields than 2D networks. To combine the advantages of 2D and 3D networks, researchers proposed H-DenseUNet [44]. This network firstly uses a 2D network to extract image features and perform segmentation tasks on a slice-by-slice basis. The pixel-wise probabilities produced by the 2D network are then concatenated with the original 3D volume and fed into a 3D network for a refinement. The H-DenseUNet finally achieves excellent liver and liver tumor segmentation. In addition, Vu et al. [53] applied the overlay of adjacent slices as input to the central slice prediction, and then fed the obtained 2D feature maps into a standard 2D network for model training. Although these pseudo-3D approaches can segment objects from 3D volume data, they only obtain limited accuracy improvement due to the utilization of local temporal information. Compared to pseudo-3D networks, hybrid cascading 2D and 3D networks are more popular for medical image segmentation.

Although the networks mentioned above can perform end-

to-end liver and liver tumor segmentation well, the use of vanilla convolution limits the further improvement of segmentation accuracy. Since standard convolution kernels have a regular sampling grid, they are unable to capture accurately liver and liver tumor features with variable shapes in different slices. Besides, some improved networks such as CE-Net [45] and MSB-Net [51] employ multi-scale feature fusion to enhance feature representation of networks, but many network branches lead to the requirement of more parameters. To address these issues, we propose a deformable encoder-decoder network (DefED-Net) to improve liver and liver tumor segmentation. The proposed DefED-Net includes following advantages:

(1) Feature extraction layers of the DefED-Net are constructed by using the deformable convolution with residual design. The design can more effectively extract the spatial context information of images while maintaining high-level features.

(2) The feature fusion module of the DefED-Net depends on a Ladder-atrous-spatial-pyramid-pooling (Ladder-ASPP) that employs multi-scale dilated convolution kernels using variable dilation rate to obtain better spatial context information.

(3) The DefED-Net provides higher segmentation accuracy for liver and liver tumor than state-of-the-art approaches, and it requires smaller memory usage due to the employment of depth separable convolution.

The rest of the paper is organized as follows. In the next section, we detailedly introduced the design of network architecture and advantages of our proposed DefED-Net. To demonstrate the superiority of DefED-Net, we introduced our experimental environment and pretreatment, performed ablation studies and comparative experiments, and analyzed the experimental results in Section III, followed by the conclusion in Section IV.

II. METHOD

In this work, we propose a deformable encoder-decoder network (DefED-Net) and apply it to liver and liver tumor segmentation. Fig. 1 shows the architecture of the DefED-Net. As can be seen from Fig. 1 that the DefED-Net is an enhanced U-net and it is composed of three parts including an encoder, a middle processing module and a decoder. In contrast with the U-net, the DefED-Net employs the deformable convolution with residual structure to generate feature maps. Moreover, the original image is concatenated with outputs at different layers of the decoder to obtain better feature representation. Different from general pyramid pooling modules [45, 46], we design a better feature fusion module namely Ladder-ASPP and apply it to our DefED-Net. Although the Ladder-ASPP adopts the way of dense connection, it only requires smaller memory usage due to the utilization of the depth separable convolution. It is worth mentioning that the DefED-Net is designed in 2D domain.

A. Deformable Encoding

Although a large number of improved U-Nets have been proposed for medical image segmentation, they provide limited segmentation accuracy for livers and liver tumors in CT images. Here are two reasons that limit the performance of U-Nets. First, convolutional kernels with fixed geometric structures are employed by the U-Nets, which ignores the shape information of objects in an image. Secondly, the operation of polling and strided convolution leads to the loss of spatial context detail information.

To illustrate the first reason, we presented an example of image filtering as shown in Fig. 2. Fig. 2 shows that the morphological opening filter is able to smooth noise effectively by employing different structuring elements (SEs). However, these filtering results depend on the choice of SEs. Fig. 2(b) shows that a circular SE is useful for preserving the details of circular objects and Fig. 2(c) shows that a square SE is effective for square objects. Similarly, Fig. 2(d) shows that a linear SE can maintain the details of linear objects. Therefore, it is better to adopt multiple different SEs for an image including many different objects. In other words, we should consider adaptive filters that can obtain better filtering effect due to the consideration of geometrical shape information of objects. In addition, the design of convolution kernels also plays the same important role for convolutional neural networks. In practical applications, researchers often employ fixed-shape square convolutional kernels to perform feature learning such as U-Net, PSP-Net, CE-Net, etc. Since convolution kernels with fixed shape show weak ability for the extraction of image contextual information, these aforementioned networks only provide tolerable accuracy for liver and liver tumor segmentations. Instead, we use deformable convolution kernels to extract richer geometry information of the liver and liver tumor, which can better accommodate the irregular shape of liver and liver tumor and lead to better segmentation results. In Fig. 3, the deformable convolution shows better adaption for liver in a CT image than the vanilla convolution. In fact, Sun et al. [62] have started to explore the utilization of deformable convolution on automatic segmentation networks for gastric cancer, and their proposed network achieves better segmentation results than vanilla U-Net [28] and ResU-Net [61].

The deformable convolution is able to provide convolutional kernels with arbitrary shapes by learning offset locations, and thus adaptively decide scales of receptive field with fine localization. Therefore, the DefED-Net possesses better capability of modeling geometric transformation than common U-Nets due to the employment of deformation convolution. However, the implementation of deformable convolution is more complex than vanilla convolution since additional spatial offset locations are limited. Based on learned offset locations, the convolution kernels can achieve the deformation of different scales, shapes and orientations. Fig. 3 illustrates the principle of deformation convolution on liver segmentation.

In practical applications, a deformable convolution is composed of four layers: a convolutional layer, a convolutional offset layer, a batch normalization layer and an activation layer. The principle of deformable convolution is given as follows. Let x and y be the input and the output feature map, respectively. The L denotes a regular grid in 2D domain.

When performing the convolution operation on x using the



Fig. 1. The architecture of the proposed DefED-Net. Firstly, the feature encoder employs deformable convolution using the residual connection. Secondly, the Ladder-ASPP block is used to extract richer context information. Finally, both the skip connection and the dense connection of original images are used for the fusion of feature maps in decoder.



Fig. 2. Image filtering using a morphological opening filter with different structuring elements. (a) The original image. (b) The SE is a disk of size 20×20. (c) The SE is a square of size 20×20. (d) The SE is a line that the length is 20 and the orientation is $1/6\pi$.



Fig. 3. Comparison of vanilla convolution and deformable convolution for liver segmentation. In contrast with the standard convolution, the deformable convolution requires offset locations for each sampling location.

L, the output is denoted by:

$$y(e_0) = \sum_{e_n \in L} w(e_n) \times x(e_0 + e_n),$$
 (1)

where w denotes the weight, e_0 denotes the location of a pixel and e_n denotes the location of neighboring pixels falling into L. If we perform the deformable convolution on x, the output can be represented by:

$$\widetilde{y}(e_0) = \sum_{e_n \in \widetilde{L}} w(e_n) \times x(e_0 + e_n + \Delta e_n), \qquad (2)$$

where \tilde{L} is the deformation result of L. Compared to L, \tilde{L} is an irregular grid including offset locations Δe_n .

The offset $\triangle e_n$ is usually a float number and the sampling position of the deformable convolution becomes irregular, so the bilinear interpolation is used to perform the process of determining the pixel value of the final sampling position. The pixel value x(e) at the final sampling position is defined as:

$$x(e) = B(w_i, q_j), \qquad (3)$$

where w_i denotes the corresponding weight, q_j denotes the four surrounding pixels involved in the computation at the irregular sampling position, B(,) is the bilinear interpolation kernel. Note that B is two dimensional and is defined as:

$$B(w_i, q_i) = w_1 q_1 + w_2 q_2 + w_3 q_3 + w_4 q_4.$$
(4)

For instance, if the coordinates we got from the sampling position is (2.2, 4.6), then its nearest pixel is (2, 4), (2, 5), (3, 4), (3, 5). Therefore, in the actual program calculation, we will use the bilinear interpolation of (2, 4), (2, 5), (3, 4), (3, 5) pixels for the sampled location (2.2, 4.6) pixels.

As shown in Fig. 3, the offset is obtained by applying a convolutional layer. Note that the convolutional layer to obtain the offset needs to have the same spatial resolution and dilation rate as the convolutional layer to extract the features in the offset feature map. For each layer of the deformable convolution, when the input of a convolutional layer is a feature map with N channels, the corresponding offset map includes 2N channels in this convolutional layer because each channel includes two offset maps in the x and y directions, separately. Note that the offset map of the output has the same spatial resolution as the input map in a convolutional layer. During training, the offset can be learned through the back propagation of Eq. (3) and Eq. (4). After the pixel values of all sampled positions are obtained, a new feature map will be generated. Although the deformable convolution is superior to vanilla convolution due to the employment of convolutional kernels with flexible shape, it can be further improved by using multi-scale convolutional kernels instead of single-scale kernels. For liver segmentation task, a large convolutional kernel is better than small ones for capturing coarse liver areas. However, a small convolutional kernel is more useful for obtaining accurate contour details. Therefore, here we use a large convolutional kernel 7×7 for the first deformable convolution layer while using small convolutional kernel 3×3 for subsequent layers. The proposed multi-scale deformable convolution is able to achieve better feature representation than single-scale deformable convolution, and thus leads to better liver and liver tumor segmentation results due to more accurate liver and liver tumor contours. In addition, the residual design is integrated in the proposed deformable encoder to avoid vanishing gradients and speeds up the convergence of networks.

B. Ladder-ASPP

Both pyramid pooling (PP) and atrous spatial pyramid pooling (ASPP) are two popular ways for encoding context information due to wider receptive fields than standard pooling. Since the PP directly performs pooling operation using multiscale pooling kernels, it often causes irreversible information loss leading to poor segmentation results for small objects such as liver tumor. However, the ASPP performs atrous convolution using multiple dilation convolutional kernels instead of multi-scale pooling kernels. Compared with PP, ASPP provides better context information since atrous convolution is superior to pooling operation for the preservation of detail information. However, ASPP still faces two challenges in practical applications. The first one is that the fixed dilation rate is used for ASPP, which causes gridding effect as shown in Figs. 4 (a-c); some pixels falling into receptive fields cannot take part in the convolutional operation. The second one is that ASPP ignores the global context information. To address these issues, we proposed a novel Ladder-ASPP as shown in Fig. 5.



Fig. 4. Comparison of standard atrous convolution and the atrous convolution with variable dilation rate. (a) Convolutional kernel: 3×3 , rate=2. (b) The cascade of two convolutional kernels: 3×3 , rate=2. (c) The cascade of three convolutional kernels: 3×3 , rate=2. (d) Convolutional kernel: 3×3 , rate=1. (e) The cascade of two convolutional kernels: 3×3 , rate=1, 2, and 3, rate=1, 2, and 4, rate at the convolution of the cascade of the convolutional kernels: 3×3 , rate=1, 2, and 3. Note that although (c) and (f) have similar receptive fields 13×13 , (c) has 70% pixel loss compared to (f).

The standard atrous convolution easily leads to the loss of spatial detail information. To overcome the drawback, we use variable dilation-rate instead of fixed dilation rate leads to better receptive fields. It is clear that each pixel in the receptive fields is covered as shown in Figs. 4 (d-f). Therefore, atrous convolution with variable dilation rate can overcome gridding effect caused by the standard atrous convolution.

Based on atrous convolution with variable dilation rate, we design a Ladder-ASPP to improve context encoding. Fig. 5 shows the architecture of the Ladder-ASPP.

First, the Ladder-ASPP employs variable dilation rate to achieve atrous convolution that was mentioned previously.

Secondly, the Ladder-ASPP uses densely ladder connection that is helpful for ASPP to achieve better feature fusion.



Fig. 5. The architecture of the Ladder-ASPP. The output feature maps are concatenated by two parts. The first one is the output from global pooling and the second one is the densely connected feature fusion liking ladder.

However, the dense connection easily leads to the increase of the number of parameters and high memory requirement. To reduce the number of parameters to obtain a lightweight network, we introduce depthwise separable convolution (DSC) [52] to Ladder-ASPP. Compared to the standard convolution in which spatial features and channel features are often coupled together, the depthwise separable convolution can achieve the decoupling computation between spatial features and channel features leading to the requirement of fewer parameters.

It is well-known that the standard convolution requires parameters $D_K \times D_K \times M \times N$, where M is the dimension of input feature maps and N is the dimension of output feature maps, and D_K is the space-resolution of convolution kernels. In the depthwise separable convolution, the depthwise convolution only requires parameters $D_K \times D_K \times 1 \times M$ and the pointwise convolution only requires parameters $1 \times 1 \times M \times N$. Therefore, the number of parameters of depthwise separable convolution is $(1/N + 1/D_K^2)$ of the standard convolution. Here, the proposed Ladder-ASPP employs four kernels of size 3×3. Consequently, the Ladder-ASPP only requires 36% parameters compared to the one without using depthwise separable convolution.

Finally, to improve feature representation of ASPP, the global pooling is integrated into Ladder-ASPP since it can achieve the priority of channels including more important information. We can see from Fig. 5 that the information hidden in both space dimension and channel dimension is exploited simultaneously. The final feature maps fuse both the global and local information.

To illustrate the proposed Ladder-ASPP, let Y be the output feature map, y_1 be the global pooling result, and y_2 be the output from the module of ladder autrous convolution. It is clear that $Y = y_1 + y_2$. The y_1 is defined as:

$$y_1 = B[C_1[GP_S(x)]] \times x, \tag{5}$$

where x is the feature map obtained from the feature encoder, followed by global pooling denoted by $GP_S(x)$, and C_1 represents the weight of each feature channel through 1×1 convolution, B is the normalization of feature weight.

In our Ladder-ASPP, we adopt variable dilation rate, i.e., 1, 2, 5, and 7. Let $G_{K,D}$ be the output of densely connected

pyramid pooling, where K is the level of pyramid and D is dilation rate, we get:

$$y_2 = G_{1,1}(x^{(1)}) \oplus G_{2,2}(x^{(2)}) \oplus G_{3,5}(x^{(3)}) \oplus G_{4,7}(x^{(4)}),$$
(6)

where,

$$\begin{cases} x^{(1)} = x \\ x^{(2)} = x^{(1)} \oplus G_{1,1}(x^{(1)}) \\ x^{(3)} = x^{(2)} \oplus G_{2,2}(x^{(2)}) \\ x^{(4)} = x^{(3)} \oplus G_{3,5}(x^{(3)}) \end{cases}$$
(7)

where the \oplus denotes concatenation operation.

According to Eqs.(5-7), we can see that the output from Ladder-ASPP includes richer information than the original input. The Ladder-ASPP can help our DefED-Net to achieve better segmentation results due to the exploitation of significant spatial information.

C. Loss function

Our framework is an end-to-end deep learning system. As illustrated in Fig. 1, we need to train the proposed method to predict each pixel as foreground or background, which is a pixel-wise classification problem. The cross entropy is one of the most popular loss functions and it is defined as:

$$L_{cross} = -(plog(\hat{p}) + (1-p)log(1-\hat{p})), \qquad (8)$$

where the p and \hat{p} are the ground truth and predicted segmentation, respectively.

However, the tumor often occupies a small region in an image. The cross entropy loss is not optimal for such tasks. It is worth noting that the Dice loss [40] is suitable for uneven samples. This metric is essentially a measure of overlap between a segmentation result and corresponding ground truth. The Dice Loss is defined as:

$$L_{dice} = 1 - \frac{2 < p, \hat{p} >}{\|p\|_1 + \|\hat{p}\|_1},\tag{9}$$

where $p \in (0, 1)$ and $0 \le \hat{p} \le 1$. The p and \hat{p} are the ground truth and predicted segmentation, respectively, and $< p, \hat{p} >$ denotes dot product.

However, the use of the Dice loss easily influences the back propagation and leads to a training difficulty. Therefore, the final loss function is defined as a combination of both losses:

$$L_{loss} = L_{cross} + L_{dice}.$$
 (10)

D. Post Processing

Generally, the task of liver and liver tumor segmentation aims to obtain a binary image where the foreground denotes liver and liver tumor and the background denotes other areas. Based on section II.A and II.B, we can obtain a coarse segmentation result for livers and liver tumors. However, the segmented image often includes a lot of small and isolated areas or some holes. In practical applications, binary image filtering is often used to remove false liver areas or fill holes within livers. For binary image filtering, morphological filters are very popular for the removal of small segmentation areas. Although both classic morphological opening and closing operations can effectively improve binary segmentation results, they often smooth the boundaries of main objects as well. It is difficult to remove false objects while maintaining the boundary accuracy of real objects. For this problem, morphological reconstruction is an excellent tool and it has been widely used for object extraction [48]. Morphological reconstruction is able to achieve binary image filtering while maintaining the large objects unchanged. The operation requires to set the parameter of SEs. If the parameter is large, more small areas would be removed. On the contrary, fewer areas are removed in the case of small value of parameters. To address the issue, we propose an adaptive morphological reconstruction to optimize liver and liver segmentation results from the DefED-Net.



Fig. 6. The post-processing results using adaptive morphological reconstruction. Top: segmentation results from the DefED-Net. Bottom: post-processing results.

We first compute the proportion of the maximal connected component in an image to the total area of the image. If the value is large, then a large SE will be adopted. Conversely, a small SE will be adopted when the value is small. Here, the SE is a disk and its radius is denoted by r:

$$r = 30 \times round(R/(H \times W)) + 1, \tag{11}$$

where R denotes the area of the maximal connected component in the segmentation result, H and W denote the height and width of the input image, respectively. Fig. 6 shows postprocessing results using the proposed adaptive morphological reconstruction. Note that it is unnecessary to make postprocessing for liver tumor segmentation since the area of liver tumors is generally small.

III. RESULTS AND DISCUSSION

A. Dataset and Pre-processing

Two public contrast-enhanced CT scans datasets: Liver Tumor Segmentation Challenge (LiTS-ISBI2017) and the 3D Image Reconstruction for Comparison of Algorithm and DataBase (3Dircadb) datasets are considered as experimental data. The LiTS dataset is a large dataset that contains 130 3D abdominal CT scans, where the image size is 512×512, slice thickness varied from 0.55 mm to 6 mm, pixel spacing varied from 0.55 mm to 1 mm. The 3DIRCADb is a small dataset that contains 22 3D data, where the image size is 512×512, slice thickness varied from 1 mm to 4 mm, pixel spacing varied from 0.56 mm to 0.86 mm, and slice number varied from 184



Fig. 7. Comparison of feature maps generated by U-Net, U-Net+DC and U-Net+RDC, respectively.

to 260. We constructed the training set and validation set using 90 patients (total 43,219 axial slices) and 10 patients (total 1,500 axial slices), respectively. Then the other 30 patients (total 15,419 axial slices) are considered as the test set. For the 3DIRCADb, it was split into 17 patients for training and 5 patients for test.

Medical CT axial slices are different from normal axial slices, the former is able to obtain wider range of values from -1000 to 3000 than the latter from 0 to 255. To remove interferences and enhance liver areas, we truncated the image intensity values of all scans of [-200, 250] HU and performed the normalization on these scans. In our experiments, the given models are independently and separately performed for liver and liver tumor segmentation.

B. Experimental Setup and Evaluation Metrics

All algorithms were implemented on a desktop PC with double NVIDIA GeForce RTX 2080 Ti with 11GBVRAM. The convolutional neural networks were performed and trained using the framework of Pytorch 1.3.0.

On the model training, we set the initial learning rate (lr) to 0.001, and define the decay strategy for learning rate during training as:

$$lr = lr \times (1 - i/t_i)^{0.9},$$
(12)

where *i* denotes the number of iterations of this training, t_i denotes the total number of iterations Note that the deformable convolution requires two learning rates compared to one for vanilla convolution. We set $lr_2 = lr \times 0.01$ for offset convolutional layers used for deformable convolution networks, and

used the Adam gradient descend with momentum to optimize the model.

Five popular evaluation metrics are used to measure the accuracy of segmentation results such as dice score (DICE) [49], volumetric overlap error (VOE), relative volume difference (RVD), average symmetric surface distance (ASD) and root mean square symmetric surface distance (RMSD). The tumor burden of the liver is a measure of the fraction of the liver afflicted by cancer. In particular, as a metric, we measure the root mean square error (RMSE) in tumor burden estimates from lesion predictions. The value of DICE ranges from 0 to 1, and a perfect segmentation yields a DICE value of 1. In fact, the DICE is one of the most important metrics in image segmentation evaluation. The VOE is the complement of the Jaccard coefficient, and thus a perfect segmentation yields a VOE value of 0. The RVD is an asymmetric metric, and a smaller value of RVD means a better segmentation result. Both ASD and RMSD are used to measure the surface distance between segmentation results and ground truths, the former is used to compute the average distance but the latter is used to compute the maximal distance. Consequently, a better segmentation result corresponds to high values of DICE but low values of VOE, RVD, ASD and RMSD. Note that we evaluate segmentation results based on 3D volumes.

C. Ablation Study

This paper focuses on liver and liver tumor segmentation. Two contributions are highlighted, one is that the deformable convolution is used to instead of the vanilla convolution; the other is that Ladder-ASPP is integrated into the proposed DefED-Net to improve the context information. To demonstrate the two contributions and the effectiveness of the DefED-Net, we conducted comprehensive experiments on both LiTS liver and liver tumor datasets.

Effectiveness of the deformable convolution: We analyzed the performance of deformable convolution (DC) and residual deformable convolution (RDC), respectively. Fig. 7 shows the comparison of U-Net, U-Net+DC and U-Net+RDC on liver segmentation. It is clear that both DC and RDC can help U-Net to focus on the interesting regions and remove irrelevant background information, but RDC can help the network converge faster and obtain more accurate edge predictions. In the third column of Fig. 7, the feature maps provided by U-Net+RDC include less information that is unrelated with the liver. Consequently, U-Net obtains more fake liver regions than U-Net+RDC and U-Net+DC in the fifth column of Fig. 7. Table I demonstrates the effectiveness of the first contribution. We can see that the utilization of DC effectively raises the segmentation accuracy of U-Net. The residual design not only speeds up the convergence of U-Net, but also further improves segmentation accuracy.



Fig. 8. Difference of prediction results and ground truths. Top: Input images, Middle: U-Net results, and Bottom: U-Net+Ladder-ASPP results.

 TABLE I

 Comparison of Ablation study on LITS test datasets. The best values are in bold.

Method	Liver	Tumor	
Wellou	DICE (%)	DICE (%)	
U-Net [28]	93.99±1.18	82.16±6.26	
U-Net+post-processing	94.25±1.06	82.16±6.26	
U-Net+DC	95.23±1.13	84.57±6.21	
U-Net+RDC	95.80±1.12	85.68±6.18	
U-Net+ASPP	94.30±1.16	85.32±6.08	
U-Net+DenseASPP	95.32±1.14	85.43±6.02	
U-Net+Ladder-ASPP	95.50±1.09	86.72±5.87	
DefED-Net(without post-processing)	96.02±1.04	87.52±5.32	
DefED-Net	96.30 ±1.01	87.52±5.32	
			_

Effectiveness of Ladder-ASPP: Both U-Net+ASPP and U-Net+Ladder-ASPP use the idea of context encoding to improve feature representation of networks. The difference is that

Ladder-ASPP uses atrous convolution with variable dilation rate and dense connection to obtain better context information than ASPP. Experimental results in Table I consistently demonstrate that both ASPP and Ladder-ASPP can help U-Net to improve segmentation accuracy of livers, and the latter is superior to the former. Fig. 8 shows the difference of segmentation results between prediction results and ground truths, where the foreground is the difference and the background is the same. It is clear that the prediction result obtained by U-Net+Ladder-ASPP is closer to the ground truth than U-Net. Furthermore, U-Net+ASPP only improves the representation capability of models on the capture of spatial context information, which is unavailable for the optimization of channel dimension. Therefore, U-Net+Ladder-ASPP provides higher DICE than U-Net+ASPP as shown in Table I.



Fig. 9. Comparison of segmentation boundaries using different approaches. The green denotes ground truth, the white denotes the result provided by U-Net, the purple denotes the result provided by CE-Net and the red denotes the result provided by DefED-Net.

In addition, post-processing is also useful for improving segmentation accuracy. Table I shows that RDC plays a more important role than ASPP and post-processing for improving segmentation accuracy. The results further demonstrate that location information is more important than feature fusion for image segmentation. Fig. 9 shows the comparison of segmentation boundaries, which further illustrates the ablation study. All these comparison results demonstrate the effectiveness of the deformable convolution, Ladder-ASPP and post-processing on liver and liver tumor segmentation.

D. Experimental Comparison on Test Datasets

To validate the superiority of the proposed DefED-Net, six state-of-the-art networks used for liver and liver tumor segmentation are considered as comparative approaches. These networks can be grouped into three categories: 2D networks, 3D networks, and hybrid networks with 2D and 3D, where 2D networks include U-Net, U-Net++ and CE-Net, 3D networks include 3D U-Net and V-Net, hybrid networks include H-DenseUNet. Note that we do not give experimental results obtained by 3D U-Net and V-Net in Tables IV-V due to high risk of over-fitting on the 3DIRCADb dataset.

It is known that 3D networks can provide better segmentation results than 2D networks due to their exploitation of information between slices. Table II and Table III demonstrate



Fig. 10. Liver and Liver tumor segmentation results using different approaches.

that both 3D U-Net and V-Net provide higher segmentation accuracy than U-Net. However, CE-Net is superior to U-Net since it employs SPP to achieve feature fusion. In contrast with those networks mentioned above, H-DenseUNet provides better segmentation accuracy since it balances the advantages of both 2D networks and 3D networks. The proposed DefED-Net provides the best quantitative scores (DICE, VOE and ASD) than comparative approaches. As the DefED-Net belongs to 2D networks, it obtains lower values of RMSD than 3D networks such as 3D U-Net and V-Net. Since the 3DIRCADb is a small dataset, 3D networks including a mountain of parameters easily lead to over-fitting for the dataset. Therefore, we only show the comparison results of U-Net, U-Net++, CE-Net, H- DenseUNet and DefED-Net in Table IV and Table V, which demonstrates the DefED-Net outperforms those comparative networks on the 3DIRCADb dataset. DICE, VOE, and RVD are all overlap measures while ASD and RMSD are surface distance measures. The former focuses more on the interior of the segmentation target, while the latter focuses more on the shape similarity of the segmentation target. It is important to note that the shape and size of liver tumors vary greatly among patients as well as in the same patient at different times compared to the liver, which make it more difficult to achieve fully automatic segmentation of liver tumors. Therefore, as recorded in Tables III and V, the ASD and RMSD values for liver tumors are obviously larger than the values for livers.

TABLE II

QUANTITATIVE SCORES OF THE LIVER SEGMENTATION RESULTS USING DIFFERENT APPROACHES ON THE LITS DATASET. THE BEST VALUES ARE IN BOLD.

Method					
Wiethou	DICE (%)	VOE (%)	RVD (%)	ASD (mm)	RMSD (mm)
U-Net [28]	93.99±1.23	11.13±2.47	3.22 ± 0.20	5.79±0.53	123.57±6.28
U-Net++ [36]	94.01±1.18	11.12±2.37	2.36±0.15	5.23±0.45	120.36±5.03
CE-Net [45]	94.04±1.15	11.03±2.31	6.19±0.16	4.11±0.51	115.40 ± 5.82
3D U-Net [39]	94.10±1.06	11.13±2.23	1.42±0.13	2.61±0.45	36.43 ±5.38
V-Net [40]	94.25±1.03	10.65 ± 2.17	1.92±0.11	2.48±0.38	38.28±5.05
H-DenseUNet [44]	96.10±1.02	7.02±2.00	1.53±0.12	1.56±0.28	37.26±3.64
DefED-Net	96.30±1.01	6.88±2.10	1.46 ± 0.12	1.37±0.23	77.60±4.26

TABLE III QUANTITATIVE SCORES OF THE LIVER TUMOR SEGMENTATION RESULTS USING DIFFERENT APPROACHES ON THE LITS DATASET. THE BEST VALUES ARE IN BOLD.

Method	LITS-Tumor					Tumor Burden
Wethod	DICE (%)	VOE (%)	RVD (%)	ASD (mm)	RMSD (mm)	RMSE
U-Net [28]	82.16±6.26	26.85±16.21	3.54 ± 0.18	22.26±0.30	155.15±5.62	0.020
U-Net++ [36]	83.23±6.36	26.03±15.39	2.16±0.17	21.36±0.25	112.36±4.89	0.017
CE-Net [45]	84.02±6.15	25.62±15.21	1.59 ± 0.17	20.79 ± 0.28	100.29 ± 5.23	0.018
3D U-Net [39]	85.13±5.87	25.13 ± 15.02	1.23 ± 0.14	20.32±0.27	62.36±5.16	0.018
V-Net [40]	85.87±5.42	24.52±14.86	1.09±0.15	19.23±0.25	68.32±4.52	0.017
H-DenseUNet [44]	86.23± 5.13	24.46±13.25	0.53 ± 0.13	18.83±0.22	54.32±4.32	0.015
DefED-Net	87.52±5.32	23.85±14.62	0.52 ± 0.10	17.41±0.28	64.25±4.87	0.016

TABLE IV QUANTITATIVE SCORES OF THE LIVER SEGMENTATION RESULTS USING DIFFERENT APPROACHES ON THE 3DIRCADB DATASET. THE BEST VALUES ARE IN BOLD.

Method					
	DICE (%)	VOE (%)	RVD (%)	ASD (mm)	RMSD (mm)
U-Net [28]	92.30±1.27	11.78±3.62	-2.83±0.38	4.25±1.56	75.67±5.68
U-Net++ [36]	93.60±1.29	10.36±3.90	1.21±0.23	3.87±1.36	56.39±4.76
CE-Net [45]	94.28±1.22	10.02 ± 3.53	-1.80 ± 0.24	3.52±1.25	30.29±4.82
H-DenseUNet [44]	95.72±1.14	9.88±2.91	0.39±0.12	2.85±0.89	9.63 ±3.95
DefED-Net	96.60±1.08	5.65±2.81	0.23±0.11	2.61±0.84	12.76± 3.43

Fig. 10 shows the segmentation results from different approaches. First, from the segmentation results obtained by 2D networks, both U-Net and U-Net++ fail to identify large liver tumors but CE-Net is successful in the first column of results. U-Net++ obtains poorer segmentation results than U-Net and CE-Net for small liver tumor as shown in the second

TABLE V QUANTITATIVE SCORES OF THE LIVER TUMOR SEGMENTATION RESULTS USING DIFFERENT APPROACHES ON THE 3DIRCADB DATASET. THE BEST VALUES ARE IN BOLD.

Mathad	3DIRCADb-Tumor					Tumor Burden
Method	DICE (%)	VOE (%)	RVD (%)	ASD (mm)	RMSD (mm)	RMSE
U-Net [28]	51.25±8.28	50.75±18.26	-1.11±0.48	16.72±0.92	130.54±5.64	0.023
U-Net++ [36]	60.36±7.36	46.72±17.64	1.36±0.78	14.76±0.63	118.23±5.64	0.018
CE-Net [45]	60.25±7.18	40.36±17.26	0.93±0.32	12.42±0.79	110.67±4.92	0.020
H-DenseUNet [44]	65.47± 6.54	36.74±12.86	-0.74±0.18	12.21±0.51	32.52±3.28	0.015
DefED-Net	66.25 ±6.62	34.28±13.43	0.81±0.20	11.21±0.63	70.05±3.10	0.018

column of results. It is clear that CE-Net is able to recognize a larger range of liver tumors due to the employment of SPP module with multi-scale receptive fields. In the third column, both U-net and CE-Net obtain more false liver areas, but U-Net++ shows better performance for large liver target recognition because it has stronger generalization capability and more dense feature representation. In the fourth column of Fig. 10, U-Net and U-Net++ are inaccurate in identifying liver boundaries, while CE-Net, which uses multiple atrous convolutional parallel modules, provides higher accuracy for liver boundary detection. Secondly, it is well known that 3D networks can provide better segmentation results of liver and liver tumors than 2D networks as they can capture the temporal information of volumetic data. In both the first and second columns, it can be seen that the tumor boundaries obtained by the 3D network are clearer than results provided by 2D networks, and the tumor boundaries obtained by V-Net are clearer and more accurate than results obtained by 3D U-Net due to the utilization of feature extraction block with residual connection. In the third column, 3D networks are clearly superior to 2D networks since the former do not suffer form the problem of over-detection. Finally, it is evident from the results obtained by DefED-Net in the first and second columns that they provide more accurate segmentations of both liver and liver tumors than the above mentioned 2D and 3D networks. In addition, in the third and fourth columns, the DefED-Net focuses on the relevant liver region while suppressing the influence of surrounding organs, it thus provides smoother segmentation boundaries than comparative approaches. In general, Fig. 10 shows that the DefED-Net achieves better feature encoding and context information extraction, which is helpful for improving the segmentation accuracy of liver and liver tumor.

TABLE VI Comparison of the efficiencies of different networks. The first two best values are in bold.

Network	Operations(GFLOPS)	Training Parameters	ModelSize (MB)
U-Net [28]	123.96	13,394,242	51.15
U-Net++ [36]	25.90	9,041,700	35.34
CE-Net [45]	35.78	29,003,668	110.77
3D U-Net [39]	1032.80	16,320,322	62.27
V-Net [40]	516.12	65,173,903	248.69
DefED-Net (without DSC)	547.81	35,580,976	139.19
DefED-Net	222.44	14,529,959	56.96

E. Model-size Comparison

We also counted the number of training parameters and computational costs of networks as shown in Table VI. Compared with 2D networks, 3D networks require much more memory and higher computational cost due to the employment of 3D convolutional kernels. The number of parameters of V-Net is greatly larger than one of 3D U-Net since V-Net uses a deeper network structure than 3DU-Net, uses more convolutions and uses residual connections. On the efficiency of models, the DefED-Net is similar to U-Net.

In fact, the DefED-Net adds Ladder-ASPP block compared to U-Net. The Ladder-ASPP is a densely connected block, and thus it shows high computational complexity and requires a large of parameters as shown in Table VI. In this paper, we utilize depth separable convolution to decouple the operation of spatial-dimension and channel-dimension, which efficiently reduces the number of parameters. Thus, the added Ladder-ASPP is a very small block compared to the size of U-Net. Finally, the DefED-Net achieves excellent liver and liver tumor segmentation with low computational cost.

IV. CONCLUSION

Liver and liver tumor segmentations attract attentions of many researchers due to their importance in medical image analysis. Deep convolutional neural networks, especially U-Nets, are very useful and popular for liver and liver tumor segmentations. For improved CNNs, deformable convolution is very important for the capture of context information, but received little consideration in liver segmentation. In this paper, we have introduced deformable convolution into U-Nets to achieve better feature encoding. Furthermore, although ASPP is effective for improving the context information, atrous convolution and pooling lead to the loss of detail information. We have suggested the Ladder-ASPP for feature encoding and fusion. The Ladder-ASPP is superior to ASPP due to the dense connection and atrous convolution with variable dilation rate. Finally, the proposed DefED-Net provides the best liver segmentation results without increasing the size of models. Our studies also show that utilization of spatial information is more important than feature fusion via modifying the network architecture for liver and liver tumor segmentations. Experiments demonstrate the advantages of the proposed DefED-Net on improving segmentation accuracies and reducing modelsize for liver and liver tumor segmentations.

REFERENCES

- M. Seehawer, F. Heinzmann, L. D'Artista, J. Harbig, P. F. Roux, L. Hoenicke, *et al.*, "Necroptosis microenvironment directs lineage commitment in liver cancer," *Nature*, vol. 562, no. 7725, pp. 69-75, Oct. 2018.
- [2] A. B. Ryerson, C. R. Eheman, S. F. Altekruse, J. W. Ward, A. Jemal, R. L. Sherman, *et al.*, "Annual report to the nation on the status of cancer, 1975-2012, featuring the increasing incidence of liver cancer," *Cancer*, vol. 122, no. 9, pp. 1312-1337, Mar. 2016.
- [3] G. Li, X. Chen, F. Shi, W. Zhu, J. Tian, and D. Xiang, "Automatic liver segmentation based on shape constraints and deformable graph cut in CT images," *IEEE Trans. Image Process.*, vol. 24, no. 12, pp. 5315-5329, Dec. 2015.

- [4] D. Furukawa, A. Shimizu, and H. Kobatake, "Automatic liver segmentation method based on maximum a posterior probability estimation and level set method," in *Proc. Int. Conf. Med. Image Comput. Comput. Assist. Intervent. (MICCAI)*, pp. 117-124, 2017.
- [5] Y. LeCun, Y. Bengio, and G. Hinton, "Deep learning," *Nature*, vol. 521, pp. 436-444, May 2015.
- [6] D. Wong, J. Liu, Y. Fengshou, Q. Tian, W. Xiong, J. Zhou, et al., "A semi-knowledge method for liver tumor segmentation based on 2D region growing with knowledge-based constraints," in Proc. Int. Conf. Med. Image Comput. Comput. Assist. Intervent. (MICCAI), pp. 1-10, 2008.
- [7] C. Li, X. Wang, S. Eberl, M. Fulham, Y. Yin, J. Chen, *et al.*, "A likelihood and local constraint level set model for liver tumor segmentation from CT volumes," *IEEE Trans. Biomed. Eng.*, vol. 60, no. 10, pp. 2967-2977, Oct. 2013.
- [8] G. Chartrand, T. Cresson, R. Chav, A. Gotra, A. Tang, and J. A. De Guise, "Liver segmentation on CT and MR using Laplacian mesh optimization," *IEEE Trans. Biomed. Eng.*, vol. 64, no. 9, pp. 2110-2121, Sep. 2017.
- [9] A. Saito, S. Nawano, and A. Shimizu, "Joint optimization of segmentation and shape prior from level-set-based statistical shape model, and its application to the automated segmentation of abdominal organs," *Med. Image Anal.*, vol. 28, no. 33, pp. 46-65, Feb. 2016.
- [10] Y. Boykov, O. Veksler, and R. Zabih, "Fast approximate energy minimization via graph cuts," *IEEE Trans. Pattern Anal. Mach. Intell.*, vol. 23, no. 11, pp. 1222-1239, Nov. 2001.
- [11] S. Vicente, V. Kolmogorov, and C. Rother, "Graph cut based image segmentation with connectivity priors," in *Proc. IEEE Conf. Comput. Vis. Pattern Recogn.*, pp. 1-8, 2008.
- [12] S. Paris, F. X. Sillion, and L. Quan, "A surface reconstruction method using global graph cut optimization," *Int. J. Comput. Vis.*, vol. 66, no. 2, pp. 141-161, Jan. 2006.
- [13] J. Stawiaski, E. Decenciere, and F. Bidault, "Interactive liver tumor segmentation using graph-cuts and watershed," in *Proc. Int. Conf. Med. Image Comput. Comput. Assist. Intervent. (MICCAI)*, 2008.
- [14] E. Wisse, "An electron microscopic study of the fenestrated endothelial lining of rat liver sinusoids," J. Ultrastruct. Res., vol. 31, no. 1-2, pp. 125-150, May 1970.
- [15] R. Kéchichian, S. Valette, and M. Desvignes, "Automatic multiorgan segmentation via multiscale registration and graph cut," *IEEE Trans. Med. Imag.*, vol. 37, no. 12, pp. 2739-2749, Dec. 2018.
- [16] A. Khadidos, V. Sanchez, and C. T. Li, "Weighted level set evolution based on local edge features for medical image segmentation," *IEEE Trans. Image Process.*, vol. 26, no. 4, pp. 1979-1991, Apr. 2017.
- [17] G. Chen, L. Gu, L. Qian, and J. Xu, "An improved level set for liver segmentation and perfusion analysis in MRIs," *IEEE Trans. Inf. Technol. Biomed.*, vol. 13, no. 1, pp. 94-103, Jan. 2009.
- [18] C. Li, R. Huang, Z. Ding, J. C. Gatenby, D. N. Metaxas, and J. C. Gore "A level set method for image segmentation in the presence of intensity inhomogeneities with application to MRI," *IEEE Trans. Image Process.*, vol. 20, no. 7, pp. 2007-2016, Jul. 2011.
- [19] B. Wang, X. Yuan, X. Gao, X. Li, and D. Tao, "A hybrid level set with semantic shape constraint for object segmentation," *IEEE Trans. Cybern.*, vol. 49, no. 5, pp. 1558-1569, May 2019.
- [20] D. Shen, Y. Zhan, and C. Davatzikos, "Segmentation of prostate boundaries from ultrasound images using statistical shape model," *IEEE Trans. Med. Imag.*, vol. 22, no. 4, pp. 539-551, Apr. 2003.
- [21] D. F. Polan, S. L. Brady, and R. A. Kaufman, "Tissue segmentation of computed tomography images using a Random Forest algorithm: a feasibility study," *Phys. Med. Biol.*, vol. 61, no. 17, pp. 6553, Aug. 2016.
- [22] E. Vorontsov, N. Abi-Jaoudeh, and S. Kadoury, "Metastatic liver tumor segmentation using texture-based omni-directional deformable surface models," in *Proc. Int. Conf. Med. Image Comput. Comput. Assist. Intervent. (MICCAI)*, pp. 74-83, 2014.
- [23] S. Tomoshige, E. Oost, A. Shimizu, H. Watanabe, H. Kobatake, and S. Nawano, "Relaxed conditional statistical shape models and their application to non-contrast liver segmentation," in *Proc. Int. Conf. Med. Image Comput. Comput. Assist. Intervent. (MICCAI)*, pp. 126-136, 2012.
- [24] X. Zhang, J. Tian, K. Deng, Y. Wu, and X. Li, "Automatic liver segmentation using a statistical shape model with optimal surface detection," *IEEE Trans. Biomed. Eng.*, vol. 57, no. 10, pp. 2622-2626, Oct. 2010.
- [25] G. E. Hinton, and R. R. Salakhutdinov, "Reducing the dimensionality of data with neural networks," *Science*, vol. 313, no. 5786, pp. 504-507, Jul. 2006.

- [26] E. Shelhamer, J. Long, and T. Darrell, "Fully Convolutional Networks for Semantic Segmentation," *IEEE Trans. Pattern Anal. Mach. Intell.*, vol. 39, no. 4, pp. 640-651, Apr. 2017.
- [27] D. Nie, L. Wang, Y. Gao, and D. Shen, "Fully convolutional networks for multi-modality isointense infant brain image segmentation," in *Proc. IEEE Conf. Int. Symp. Biomed. Imag.*, pp. 1342-1345, 2016.
- [28] O. Ronneberger, P. Fischer, and T. Brox, "U-net: Convolutional networks for biomedical image segmentation," in *Proc. Int. Conf. Med. Image Comput. Comput. Assist. Intervent. (MICCAI)*, pp. 234–241, 2015.
- [29] K. Simonyan, and A. Zisserman. "Very deep convolutional networks for large-scale image recognition," arXiv preprint arXiv:1409.1556, 2014.
- [30] K. He, X. Zhang, S. Ren, and J. Sun, "Deep residual learning for image recognition," in *Proc. IEEE Conf. Comput. Vis. Pattern Recogn.*, pp. 770–778, 2016.
- [31] G. Huang, Z. Liu, L. Van Der Maaten, and K. Q. Weinberger, "Densely connected convolutional networks," in *Proc. IEEE Conf. Comput. Vis. Pattern Recogn.*, pp. 2261–2269, 2017.
- [32] N. Abraham, and N. M. Khan, "A novel focal tversky loss function with improved attention u-net for lesion segmentation," in *Proc. IEEE Conf. Int. Symp. Biomed. Imag.*, pp. 683-687, 2019.
- [33] O. Oktay, J. Schlemper, L. L. Folgoc, M. Lee, M. Heinrich, K. Misawa, et al., "Attention u-net: Learning where to look for the pancreas," arXiv preprint arXiv:1804.03999, 2018.
- [34] Q. Jin, Z. Meng, C. Sun, L. Wei, and R. Su, "RA-UNet: A hybrid deep attention-aware network to extract liver and tumor in CT scans," arXiv preprint arXiv:1811.01328, 2018.
- [35] M. Z. Alom, M. Hasan, C. Yakopcic, T. M. Taha, and V. K. Asari, "Recurrent residual convolutional neural network based on u-net (R2Unet) for medical image segmentation," *arXiv preprint arXiv*:1802.06955, 2018.
- [36] Z. Zhou, M. M. R. Siddiquee, N. Tajbakhsh, and J. Liang, "UNet++: Redesigning Skip Connections to Exploit Multiscale Features in Image Segmentation," *IEEE Trans. Med. Imag.*, 2019, doi: 10.1109/TMI.2019.2959609.
- [37] H. Seo, C. Huang, M. Bassenne, R.Xiao, and L. Xing, "Modified U-Net (mU-Net) with Incorporation of Object-Dependent High Level Features for Improved Liver and Liver-Tumor Segmentation in CT Images," *IEEE Trans. Med. Imag.*, 2019, doi: 10.1109/TMI.2019.2948320.
- [38] S. Ji, W. Xu, M. Yang, and K. Yu, "3D Convolutional neural networks for human action recognition," *IEEE Trans. Pattern Anal. Mach. Intell.*, vol. 35, no. 1, pp. 221-231, Jan. 2013.
- [39] Ö. Çiçek, A. Abdulkadir, S. S. Lienkamp, T. Brox, and O. Ronneberger, "3D U-net: Learning dense volumetric segmentation from sparse annotation," in *Proc. Int. Conf. Med. Image Comput. Comput. Assist. Intervent.* (*MICCAI*), pp. 424–432, 2016.
- [40] F. Milletari, N. Navab, and S. A. Ahmadi, "V-Net: fully convolutional neural networks for volumetric medical image segmentation," in *Proc. Int. Conf. 3D Vis.*, pp. 565-671, 2016.
- [41] S. Chen, K. Ma, and Y. Zheng, "Med3D: transfer learning for 3D medical image analysis," arXiv preprint arXiv:1904.00625, 2019.
- [42] Q. Dou, L. Yu, H. Chen, Y. Jin, X. Yang, J. Qin, et al., "3D deeply supervised network for automated segmentation of volumetric medical images," *Med. Image Anal.*, vol. 41, pp. 40–54, Oct. 2017.
- [43] T. Lei, W. Zhou, Y. Zhang, R. Wang, H. Meng, and A. K. Nandi,, "Lightweight V-Net for liver segmentation," *IEEE Int. Conf. Acoust. Speech Signal Process. (ICASSP)*, pp. 1379-1383, 2020.
- [44] X. Li, H. Chen, X. Qi, Q. Dou, C. W. Fu, and P. A. Heng, "H-DenseUNet: Hybrid densely connected UNet for liver and tumor segmentation from CT volumes," *IEEE Trans. Med. Imag.*, vol. 37, no. 12, pp. 2663-2674, Dec. 2018.
- [45] Z. Gu, J. Cheng, H. Fu, K. Zhou, H. Hao, Y. Zhao, *et al.*, "CE-Net: context encoder network for 2D medical image segmentation," *IEEE Trans. Med. Imag.*, vol. 38, no. 10, pp. 2281-2292, Oct. 2019.
- [46] H. Zhao, J. Shi, X. Qi, X. Wang, and J. Jia, "Pyramid scene parsing network," in *Proc. IEEE Conf. Comput. Vis. Pattern Recogn.*, pp. 2881-2890, 2017.
- [47] J. Dai, H. Qi, Y. Li, G. Zhang, H. Hu, and Y. Wei, "Deformable convolutional networks," in *Proc. IEEE Conf. Comput. Vis. Pattern Recogn.*, pp. 764–773, 2017.
- [48] T. Lei, X. Jia, Y. Zhang, S. Liu, H. Meng and A. K. Nandi, "Superpixel-Based Fast Fuzzy C-Means Clustering for Color Image Segmentation," *IEEE Trans. Fuzzy Syst.*, vol. 27, no. 9, pp. 1753-1766, 2019.
- [49] P. Bilic, P. F. Christ, E. Vorontsov, G. Chlebus, H. Chen, Q. Dou, et al., "The liver tumor segmentation benchmark (LiTS)," arXiv preprint arXiv:1901.04056, 2019.

- [50] K. Han, Y. Wang, Q. Tian, J. Guo, C. Xu, and C. Xu, "GhostNet: More features from cheap operations," in *Proc. IEEE Conf. Comput. Vis. Pattern Recogn.*, pp. 1580-1589, 2020.
- [51] Q. Shao, L. Gong, K. Ma, H. Liu, and Y. Zheng, "Attentive CT lesion detection using deep pyramid inference with multi-scale booster," in *Proc. Int. Conf. Med. Image Comput. Comput. Assist. Intervent.* (*MICCAI*), pp. 301-309, 2019.
- [52] A. G. Howard, M. Zhu, B. Chen, D. Kalenichenko, W. Wang, T. Weyand, et al., "Mobilenets: Efficient convolutional neural networks for mobile vision applications," arXiv preprint arXiv:1704.04861, 2017.
- [53] M. H. Vu, G. Grimbergen, T. Nyholm, and T. Löfstedt, "Evaluation of multi-slice inputs to convolutional neural networks for medical image segmentation," *Med. Phys.*, 2020.
- [54] S. Ren, P. Laub, Y. Lu, M. Naganawa and R. E. Carson, "Atlas-Based multiorgan segmentation for dynamic abdominal PET," *IEEE Trans. Radiat. Plasma Med. Sci.*, vol. 4, no. 1, pp. 50-62, Jan. 2020.
- [55] Z. Guo, X. Li, H. Huang, N. Guo and Q. Li, "Deep Learning-Based image segmentation on multimodal medical imaging," *IEEE Trans. Radiat. Plasma Med. Sci.*, vol. 3, no. 2, pp. 162-169, March 2019.
- [56] C. De Sio, J. J. Velthuis, L. Beck, J. L. Pritchard and R. P. Hugtenburg, "r-UNet: Leaf Position Reconstruction in Upstream Radiotherapy Verification," *IEEE Trans. Radiat. Plasma Med. Sci.*, doi: 10.1109/TRPMS.2020.2994648.
- [57] P. H. Conze, F. Rousseau, V. Noblet, F. Heitz, R. Memeo and P. Pessaux, "Semi-automatic liver tumor segmentation in dynamic contrast-enhanced CT scans using random forests and supervoxels," *Int. Workshop Mach. Learn. Med. Imag.*, pp. 212-219, 2015.
- [58] P. H. Conze, V. Noblet, F. Rousseau, F. Heitz, V. de Blasi and R. Memeo, "Scale-adaptive supervoxel-based random forests for liver tumor segmentation in dynamic contrast-enhanced CT scans," *Int. J. comput. assist. radio. surg.*, vol. 12, no. 2, pp. 223-233, 2017.
- [59] Y. Li, S. Hara and K. Shimura, "A machine learning approach for locating boundaries of liver tumors in CT images," *Int. Conf. Pattern Recogn.*, pp. 400-403, 2006.
- [60] P. H. Conze, A. E. Kavur, E. C. L. Gall, N. S. Gezer, Y. L. Meur, M. A. Selver, *et al.*, "Abdominal multi-organ segmentation with cascaded convolutional and adversarial deep networks," *arXiv preprint arXiv:2001.09521*, 2020.
- [61] X. Xiao, S. Lian, Z. Luo, S. Li, "Weighted res-unet for high-quality retina vessel segmentation," *Int. Conf. Inf. Technol. Med. Educ. (ITME*), pp. 327–331, 2018.
- [62] M. Sun, G. Zhang, H. Dang, X. Qi, X. Zhou and Q. Chang, "Accurate Gastric Cancer Segmentation in Digital Pathology Images Using Deformable Convolution and Multi-Scale Embedding Networks," *IEEE Access*, vol. 7, pp. 75530-75541, 2019.