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CT Image Based Computer-Aided Lung Cancer Diagnosis

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1. Introduction

An early stage detection of lung cancer is extremely important for survival rate and quality of life (QOL) of patients (Naruke et al., 1988). Although a nationwide periodical group medical examination is conducted in Japan by diagnosing chest X-ray images, such group examination is not often good enough to detect the lung cancer accurately and thus there is a high possibility that the cancer at an early stage cannot be detected by using only the chest X-ray images. To improve the detection rate for the cancer at early stages, X-ray computed tomography (CT) has been used for a group medical examination as well (Iinuma et al., 1992; Yamamoto et al., 1993).

Using the X-ray CT, pulmonary nodules that are typical shadows of pathological changes of the lung cancer (Prokop and Galanski, 2003) can be detected more clearly compared to the chest X-ray examination even if they are at early stages. This is an advantage of the X-ray CT diagnosis. In fact, it has been reported that the survival rate of the later ten years can reach 90% after the detection at early stages using X-ray CT images (I-ECAP, 2006).

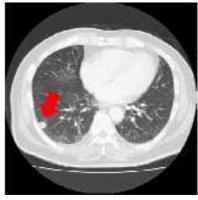
On the other hand, compared to the chest X-ray images diagnosis, the X-ray CT diagnosis may exhaust radiologists because the CT generates a large number of images (at least over 30 images per patient) and they must diagnose all of them. The radiologists' exhaustion and physical tiredness might cause a wrong diagnosis especially for a group medical examination where most of CT images are healthy and only very few images involve the pathological changes. Therefore, some computer-aided diagnosis (CAD) systems have been developed to help their diagnosis work (Okumura et al., 1998; Lee et al., 1997; Yamamoto et al., 1994; Miwa et al., 1999). Core techniques of CAD systems can be found in feature extraction and pattern recognition. Because of the fuzziness of the diagnosis target in the medical images, it often requires different methods from those for artificial targets.

Miwa et al. have developed a variable N-quoit filter to detect isolated pulmonary nodules (Miwa et al., 1999) and Homma et al. have further improved the detection accuracy by discriminating between the isolated nodules and blood vessels those are both in a circle-like shape in CT images (Homma et al., 2008). The discrimination was achieved by developing new feature extraction techniques and combining those features extracted by the techniques. These methods, however, aimed at detecting only isolated circle-like shapes with the some morphological features, and thus non-isolated nodules (pathological changes) may not be detected by such methods. Indeed, it has been demonstrated that the conventional methods can detect isolated nodules shown in Fig. 1 (a) (Homma et al., 2008), but cannot or hard to

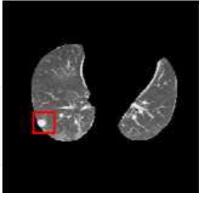
detect a non-isolated nodule shown in Fig. 1 (b). A schematic difference between isolated and non-isolated targets is depicted in Fig. 2.



(a) Red squares show locations of isolated-nodules

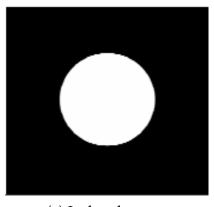


(b) Red arrow indicates a non-isolated nodule

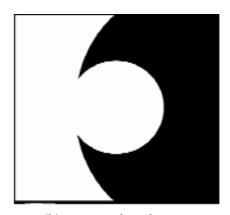


(c) Red square shows a converted isolated nodule from non-isolated one

Fig. 1. (a) Isolated and (b) non-isolated nodules, and the conversion (c) from non-isolated into isolated one.



(a) Isolated target



(b) Non-isolated target

Fig. 2. A schematic difference between isolated and non-isolated targets.

Although non-isolated nodules are not very often seen in lung cancer observations, they can be a lung cancer with a high possibility and should not be missed from the viewpoint of the early stages detection of cancers (I-ECAP, 2006).

In this chapter, to improve the detection rate of such non-isolated nodules, we propose a technique transforming the *non-isolated nodules* connected to the walls of the chest into *isolated ones* that can be detected more easily by the conventional CAD systems. The transformation of Fig. 2 (b) into (a) can be achieved by extracting the lung area from the original whole CT image as shown in Fig. 1 (c).

The rest of this chapter consists of as follows. In section 2, a fundamental theory of active contour models (Kass et al., 1998) that can be used for such extraction and its local optimum problem will be introduced. Then, by setting appropriate initial contours for solving the local optimum problem, a novel extraction technique based on the contour model will be developed in section 3. Experimental results using clinical data of X-ray CT images will be

discussed to demonstrate the usefulness of the proposed method in section 4. Concluding remarks will be given in section 5.

2. Active contour model

The active contour model proposed by Kass ((Kass et al., 1998) uses a gradient decent-based optimal method. The optimality can be defined by an energy function. The time evolution of the model is controlled by the following partial differential equation.

$$\frac{\partial v}{\partial t} = -\eta \frac{\partial E}{\partial v} \tag{1}$$

where v(t,x,y) is a function of time t and coordinates x and y in the two dimensional space of the original image. η is a positive coefficient. The contour can be defined by a set of coordinates (x,y) satisfying a condition v=L where L is a constant. Obviously, the final contour evolved by (1) is depended on the energy function E.

A well known simple energy function is related to the edge of the original image and can be defined as follows.

$$E = -\int_{\Omega} |\nabla I(x, y)|^2 dxdy$$
 (2)

where I(x,y) is a pixel value at the coordinates (x,y) and $\nabla I(x,y)$ is the spatial gradient of the pixel value. Ω is a domain of the coordinates (x,y) on the contour, i.e, $\Omega = \{(x,y) | v(x,y) = L\}$. By using the energy function E in Eq. (2), the final contour may be on an edge of the original image in which the gradient of the pixel value is the local maximum (i.e., the local minimum for the energy function). Fig. 3 shows an example of the time evolution of the contour given by the active contour model where the energy function was defined by Eq. (2).

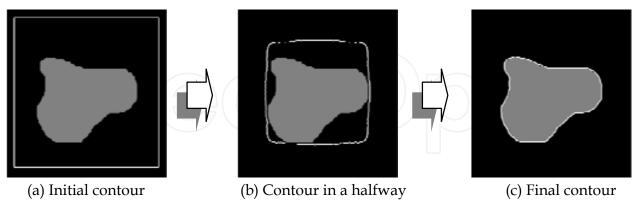
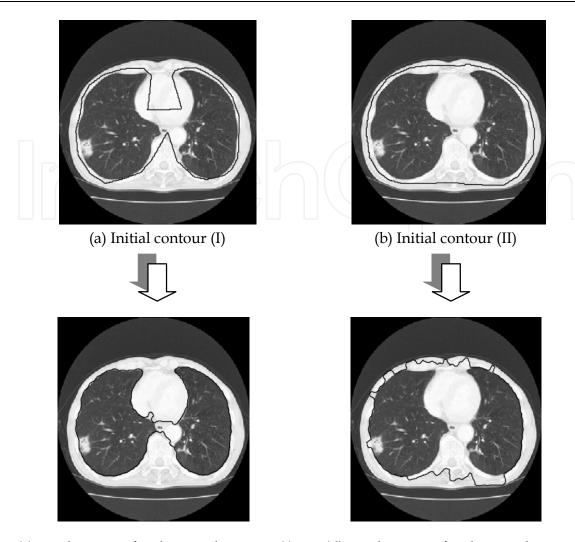


Fig. 3. A sample time evolution of active contour. White lines show contours.

Since the active contour model is controlled by a gradient-decent evolution as mentioned above, the final result is also depended on the initial settings of the contour. In other words, such model can converge to a local optimal solution instead of the global optimal one. Thus, as well as the right design of the energy function, an appropriate setting of the initial contour is required to obtain the desired contour. Fig. 4 shows an example illustrating



(c) Final contour for the initial contour (I) (d) Final contour for the initial contour (II)

Fig. 4. Effect of initial contours on the final results: Examples using the same lung X-ray CT image. Black lines near the walls on the CT images are contours.

results obtained from different initial contours for the same X-ray CT image. In fact, as is clear from this figure, the results are quite different from each other.

In addition, note that the result (I) in Fig. 4 (c) may be more desirable than the result (II) in Fig. 4 (d) because the result (I) seems more similar to the target contour inside the walls of the chest. This is because the initial contour (I) in Fig. 4 (a) is more similar to the target and thus appropriate than the initial contour (II) in Fig. 4 (b). Consequently, if an initial contour as similar as possible to the desirable contour could be given, it may be expected that the final result is the most desirable one since the number of local optimal contours encountered during the time evolution can be the minimum compared to those for the other initial settings.

3. Advanced active contour model for lung cancer diagnosis

As expected in the last paragraph of section 2, the local optimum problem can be avoided by starting from the appropriate initial contours. Note that a lung shape changes smoothly in axial direction as shown in Fig. 5 and recently the interval between X-ray CT slices next to each other is at most 10 [mm] in the direction. Then lung shapes in CT slices (axial tomography)

next to each other are almost the same or at least similar as shown in original CT images of Fig. 6. Thus a novel technique proposed here initializes the contour by using such anatomical characteristics of the lung shape. That is, the resulting contour obtained from the active contour model on the CT slice next to a target slice can be an appropriate candidate for the initial contour of the target CT slice. This is a key idea of the proposed initialization. Let us define, in this chapter, a lung area as inside the thorax that includes the center area of heart and aorta, and consider the walls of the chest that does not include the center area.

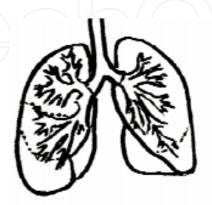


Fig. 5. A schema of a human lung.

A flowchart of the proposed algorithm for extracting the lung area is shown in Fig. 7. In this algorithm, only the first CT slice is needed to be initialized in a specific way and called the *initial slice* of a series of the slices. Because of the specific initialization, steps (i) and (ii) in the flowchart for the initial slice are different from those of the other slices. In the followings, it is assumed, for simplicity, that the algorithm processes the series of CT slices from the head to the legs in the axial direction, but the algorithm is the same for the reverse direction.

- (i). Selection of the target slice: If the current target is the initial slice of the series, select a slice without non-isolated nodules connected to the walls of the chest. Otherwise, select the slice below the previous target slice.
- (ii). **Initialization:** There are many local optima during the time evolution of the model due to the edges created by the costae (bones) in the walls of the chest as shown in Fig. 4 (d). The resulting contour of the previous target slice can be a good candidate for the initial contour of the current slice as described above. The initialization except for the initial slice can thus be done easily by setting the candidate.

There is, however, no previous final contour for the initial slice. In this case, to remove such undesirable edges, an equalization of the pixel values that are larger than a threshold is conducted within the walls of the initial slice. The equalization can be given as follows.

$$\vec{I}(x,y) = \begin{cases}
I_{max}, & (I(x,y) > I_{Th}) \\
I(x,y), & (otherwise)
\end{cases}$$
(3)

where I'(x,y) denotes a new pixel value after the equalization, I_{Th} is the threshold, and I_{max} is the maximum pixel value that usually represents the white color.

As shown in Fig. 8, lung area of the initial slice can be extracted by using a mask processing. Then, a good result can be obtained from any contour outside the mask

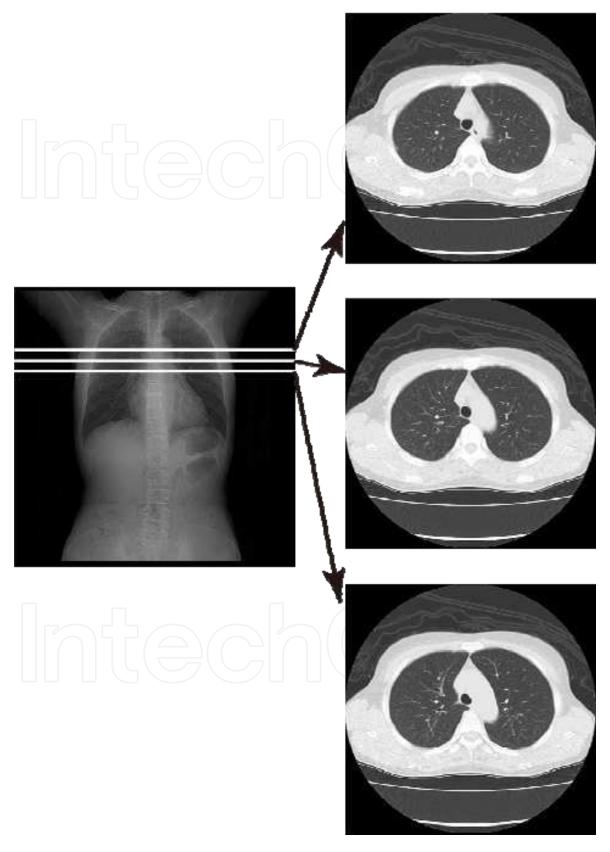


Fig. 6. Similar lung shapes between CT slices next to each other.

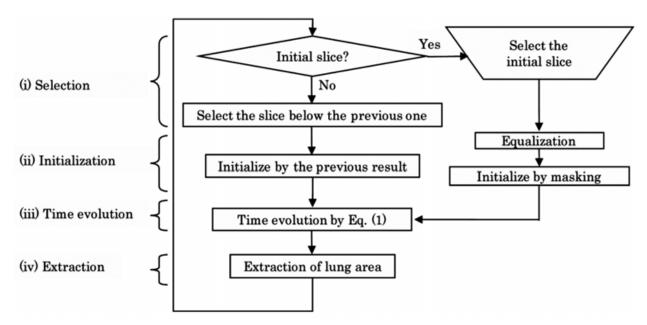


Fig. 7. Flowchart of the proposed method.

area. Note that lung area, however, could not often be extracted correctly if there is a non-isolated nodule connected to the walls of the chest as shown in Fig. 9. In this case, the non-isolated nodule that we want to detect is regarded as outside the lung area and thus cannot be detected by the mask processing. This is only the reason why we need to select the initial slice manually.

- (iii). **Time evolution:** By using Eq. (1), the resulting contour for the current target slice selected in step (i) can be obtained from the contour initialized in step (ii).
- (iv). Extraction: The lung area for the current target is extracted as the inside the resulting contour obtained in step (iii).

Steps (i) - (iv) are repeatedly conducted until all lung areas in all CT slices are extracted.

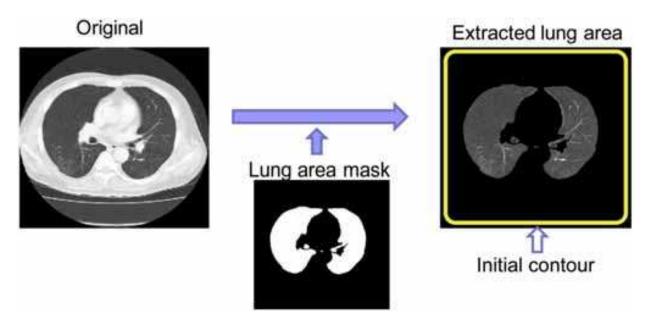


Fig. 8. A mask processing to extract the lung area.

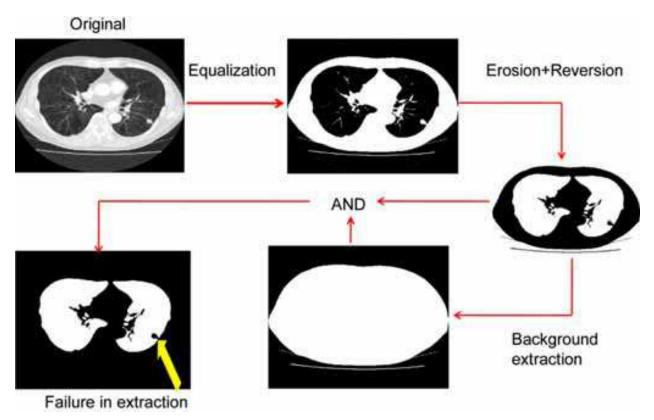


Fig. 9. A failure case of the mask processing for a slice where there is a non-isolated nodule connected to the walls of the chest.

4. Application to lung cancer diagnosis

We have tested the proposed method using an extraction task in which the clinical CT images (https://imaging.nci.nih.gov/ncia/faces/baseDef.tiles) including non-isolated nodules connected to the walls of the chest are used. Examples of the extraction results are shown in Figs. 10 and 11. It is clear that the proposed method can extract the lung area including the non-isolated nodules.

Extracted areas by the initial and the resulting contours for the original slice in Fig. 10 (c) are shown in Fig. 12. Note that the initial contour that is the resulting contour obtained in the previous slice in Fig. 10 (f) is similar enough to the target and thus, the final result in Fig. 10 (g) is good enough.

On the other hand, there are a few examples in which non-isolated nodules were not extracted as the lung area, but regarded as within the walls. In such case, still non-isolated nodules cannot be detected by the conventional CAD systems aiming at the isolated nodules detection. This problem may, however, be solved by designing a further appropriate energy function. For example, the contour curvature of the walls changes smoothly in general, but the curvature involving the connected nodules changes more sharply. Differences in the curvature may be incorporated into a new energy function to discriminate such non-isolated nodules from the walls of the chest.

Furthermore, the active contour model has an ability of making a smooth contour line even if the initial contour has a sharp corner with a high curvature. We can then select the initial slice in an automatic way, i.e., random selection, the first (top), middle, or last (bottom) slice

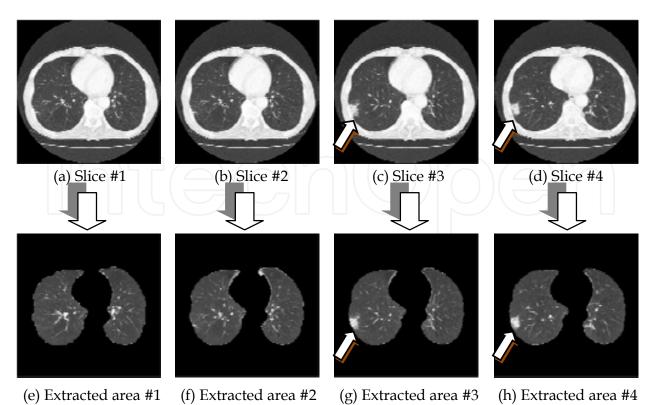


Fig. 10. Extracted results for case 1 by the proposed active contour method. (a) - (d): Original CT images. (e) - (h): Extracted lung areas.

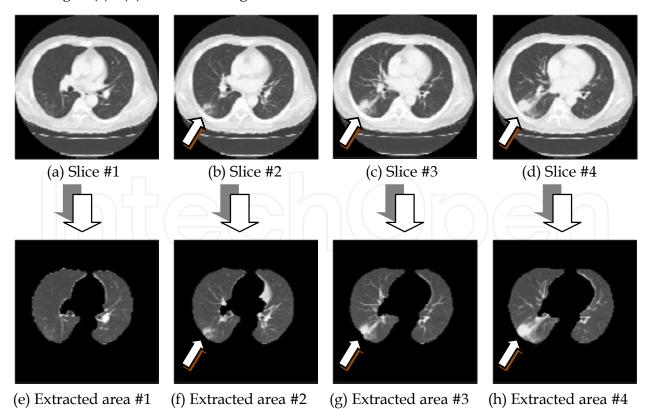
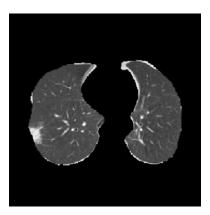


Fig. 11. Extracted results for case 2 by the proposed active contour method. (a) - (d): Original CT images. (e) - (h): Extracted lung areas.

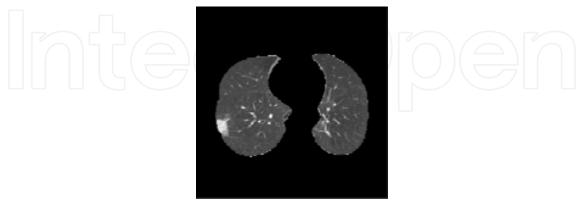
of the series, and so on. The masking problem with the initial slice including non-isolated nodules connected to the walls of the chest can be solved by applying the proposed algorithm with an appropriate parameters setting repeatedly to the same series. This direction of future works can be important for clinical use.



(a) Original CT image (same as in Fig. 10 (c))



(b) Extracted area by the initial contour that is the final contour of the above slice



(c) Extracted area by the final contour (same as in Fig. 10 (g))

Fig. 12. The appropriate initial contour and the final contour for the CT slice #3 in Fig. 10 (c).

5. Concluding remarks

In this chapter, we have taken into account non-isolated nodules connected to the walls of the chest that cannot be detected by the conventional CAD systems for lung cancer. To detect such nodules, we have proposed a technique to transform the non-isolated nodules into the isolated ones by using an active contour model to extract the lung area from the original CT image. The promising results suggest that the detection accuracy of the CAD systems can be further improved by incorporating the proposed technique.

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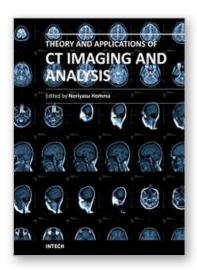
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Theory and Applications of CT Imaging and Analysis

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The x-ray computed tomography (CT) is well known as a useful imaging method and thus CT images have continuingly been used for many applications, especially in medical fields. This book discloses recent advances and new ideas in theories and applications for CT imaging and its analysis. The 16 chapters selected in this book cover not only the major topics of CT imaging and analysis in medical fields, but also some advanced applications for forensic and industrial purposes. These chapters propose state-of-the-art approaches and cutting-edge research results.

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