A fully automatic deep learning system for L3 slice selection and body composition assessment on abdominal computed tomography.

Jiyeon Ha
University of Ulsan College of Medicine

Taeyong Park
University of Ulsan College of Medicine

Hong-Kyu Kim
University of Ulsan College of Medicine

Youngbin Shin
Asan Institute for Life Sciences, Asan Medical Center

Yousun Ko
Asan Institute for Life Sciences, Asan Medical Center

Dong Wook Kim
University of Ulsan College of Medicine

Yu Sub Sung
Asan Medical Center

Jiwoo Lee
Asan Institute for Life Sciences, Asan Medical Center

Su Jung Ham
Asan Institute for Life Sciences, Asan Medical Center

Seungwoo Khang
Soongsil University

Heeryeol Jeong
Soongsil University

Kyoyeong Koo
Soongsil University

Jeongjin Lee
Soongsil University

Kyung Won Kim (medimash@gmail.com)
University of Ulsan College of Medicine

Research Article
Abstract

Background and aims: As sarcopenia research has been gaining emphasis, the need for quantification of abdominal muscle on computed tomography (CT) is increasing. Thus, a fully automated system to select L3 slice and segment muscle in an end-to-end manner is demanded.

We aimed to develop a deep learning model (DLM) to select the L3 slice with consideration of anatomic variations and to segment cross-sectional areas (CSAs) of abdominal muscle and fat.

Methods: Our DLM, named L3SEG-net, was composed of a YOLOv3-based algorithm for selecting the L3 slice and a fully convolutional network (FCN)-based algorithm for segmentation. The YOLOv3-based algorithm was developed via supervised learning using a training dataset (n=922), and the FCN-based algorithm was transferred from prior work. Our L3SEG-net was validated with internal (n=496) and external validation (n=586) datasets. L3 slice selection accuracy was evaluated by the distance difference between ground truths and DLM-derived results. Technical success for L3 slice selection was defined when the distance difference was <10 mm. Overall segmentation accuracy was evaluated by CSA error. The influence of anatomic variations on DLM performance was evaluated.

Results: In the internal and external validation datasets, the accuracy of automatic L3 slice selection was high, with mean distance differences of 3.7±8.4 mm and 4.1±8.3 mm, respectively, and with technical success rates of 93.1% and 92.3%, respectively. However, in the subgroup analysis of anatomic variations, the L3 slice selection accuracy decreased, with distance differences of 12.4±15.4 mm and 12.1±14.6 mm, respectively, and with technical success rates of 67.2% and 67.9%, respectively. The overall segmentation accuracy of abdominal muscle areas was excellent regardless of anatomic variation, with the CSA errors of 1.38–3.10 cm².

Conclusions: A fully automatic system was developed for the selection of an exact axial CT slice at the L3 vertebral level and the segmentation of abdominal muscle areas.

Introduction

The segmentation of muscle and fat areas on abdominal computed tomography (CT) has gained huge emphasis in the last decade, as sarcopenia research has been growing rapidly. According to the revised European Working Group on Sarcopenia in Older People (EWGSOP2) ¹, the muscle area on CT measured at the third lumbar vertebral level is used as a representative value because it can reflect the whole-body muscle mass ²⁻⁵.

Recently, the necessity to measure muscle and fat areas on CT has increased rapidly ⁶, increasing the demand for automatic muscle and fat measurement technologies such as deep learning model (DLM). Accordingly, there have been several previous studies that developed automatic segmentation for body composition analysis using DLM ⁷⁻¹³, and some of them are commercially available ¹⁴. These new automatic segmentation methods can reduce the time to measure abdominal muscle and fat areas to
some degree. Still, these techniques have required manual selection of L3 slice CT images, which might be the greatest hurdle to achieve fully automatic body composition measurements. In general, it takes several minutes (around three minutes) to find L3 slice level on abdominal CT even by experts\textsuperscript{15}.

So far, only a few studies have attempted to develop a fully automatic technique for L3 slice selection and muscle segmentation\textsuperscript{16,17}. However, these studies have not been clinically validated well; especially, it is unclear whether or not these studies have developed automatic L3 slice selection technique with consideration of thoracolumbar/lumbosacral variations. Thoracolumbar/lumbosacral variations may occur in around 20\% (4–30\%) of normal population\textsuperscript{18,19}. Therefore, developing a DLM-based automatic L3 slice selection technique requires training data with full consideration of anatomic variations.

The primary objective of this study was to develop a DLM to automatically select L3 slices on abdominal CT scans and then automatically segment areas of the abdominal muscle, visceral fat, and subcutaneous fat. The secondary objective was to validate the accuracy of DLM to select L3 slices with consideration of anatomic variations. The third objective was to validate the segmentation accuracy of DLM to measure muscle and fat areas at the L3 level.

### Materials And Methods

This study was approved by the institutional review boards of Asan Medical Center (AMC), Kyung Hee University Hospital (KHUH), Ajou University Hospital (AUH), and Ulsan University Hospital (UUH). The informed consent requirement was waived by the institutional review board of Asan Medical Center. The research have been performed in accordance with Declaration of Helsinki and all experimental protocols were carried according to the experimental guideline and regulations of Asan Institute for Life Sciences.

This article reports on and complies with the methods and terms described in the most recently published guidance on reading literature about machine learning for medical applications\textsuperscript{20}.

#### Data acquisition: study subjects

The datasets used for this study were as follows: (1) development dataset used for developing the DLM, which was further split into the training set and tuning set; (2) validation dataset for independent testing of model performance, including an internal validation set and an external validation set. An overview of dataset composition is described in Fig. 1.

The development dataset was composed of 922 patients (560 men and 362 women; mean age, 54.4 ± 14.0 years), with 1496 abdominal CT images. The development dataset was used in our previous study\textsuperscript{7}. The overlapped dataset composed of 922 patients was used for the development and internal validation of the DLM segmenting body composition area in the prior article. In this manuscript, the dataset was used as the development dataset for the DLM selecting L3 slice. The development dataset included patients with various diseases and healthy subjects who underwent CT scanning for potential kidney
To identify anatomic variations in accurate manner, we also obtained chest CT scans in 910 patients.

The internal validation set was composed of 500 healthy subjects who had both chest CT and abdominal CT scans acquired in our institution from March through December 2012. Four subjects who underwent interbody lumbar vertebra fusion surgery were excluded, and a total of 496 subjects with 496 CT scans were used for validation (301 men and 195 women; mean age, 53.7 ± 8.7 years). The external validation dataset included 600 patients who had both chest and abdominal CT scans, acquired between September 2011 and March 2019 from three other institutions (KHUH, AUH, and UUH). A total of 586 patients were included after excluding 14 subjects who underwent lumbar interbody fusion surgery (347 men and 239 women; mean age, 58.5 ± 12.3). The clinical characteristics of subjects included in the validation dataset are summarized in Table 1.

Healthy subjects conducted CT for evaluation for organ donation including liver and kidney. Abdominal CT scan is a part of routine clinical management for potential liver of kidney donors. Healthy subject who underwent CT scan for benign lesion were also included in internal and external validation group. Ultrasonography is a screening method widely used due to absence of radiation hazard. If a focal lesion detected on the ultrasonography, CT scan is usually conducted for further characterization of the lesion in clinical practice.

CT scanners from various manufacturers were used. Detailed specifications of the abdominal CT acquisition are summarized in Supplementary Table 1.
### Table 1
Subject characteristics of internal and external validation cohorts

| Characteristics                        | Development dataset | Internal validation dataset | External validation dataset |
|----------------------------------------|---------------------|-----------------------------|-----------------------------|
| Number of subjects                    | 922                 | 496                         | 586                         |
| Age (years)                            | 54.4 ± 14.0         | 53.7 ± 8.7                  | 58.5 ± 12.3                 |
| Female (% female:male)                 | 39.3% (362:560)     | 39.3% (195:301)             | 40.8% (239:347)             |
| **Anatomic variation**                 |                     |                             |                             |
| Normal anatomy group                   | 807 (87.5%)         | 438 (88.3%)                 | 505 (86.2%)                 |
| Anatomic variants group                | 115 (12.5%)         | 58 (11.7%)                  | 81 (13.8%)                  |
| Thoracolumbar variant                  | 48 (5.2%)           | 20 (4.0%)                   | 26 (4.4%)                   |
| Lumbosacral variant                    | 43 (4.7%)           | 29 (5.8%)                   | 43 (7.3%)                   |
| Numeric variant                        | 12 (1.3%)           | 4 (1.4%)                    | 7 (1.2%)                    |
| Combined variant                       | 12 (1.3%)           | 5 (1.7%)                    | 5 (0.9%)                    |
| Institution                            | AMC                 | AMC                         | UUH, KHUH, AUH              |
| Underlying disease (n)                 | 87                  | 496                         | 586                         |
| None                                   |                     |                             |                             |
| Gastric cancer                         | 436                 | 0                           | 0                           |
| Sepsis                                 | 245                 | 0                           | 0                           |
| Pancreatic cancer                      | 154                 | 0                           | 0                           |

Note.—AMC = Asan Medical Center, AUH = Ajou University Hospital, KHUH = Kyung Hee University Hospital, UUH = Ulsan University Hospital

### Generation of the ground truth

For each CT scan, the axial CT slice number of the third lumbar vertebra inferior endplate was annotated, and the lumbar vertebral anatomic variant was identified by a board-certified radiologist (J.H.) and double-checked by another radiologist (K.W.K.). In most cases, we counted the number of thoracolumbar spines and ribs in chest CT and abdominal CT scans to identify the anatomic variations accurately. Disagreement was resolved by reaching a consensus through discussion.

According to the vertebral anatomy, patients were divided into the normal anatomy group and the anatomic variant group. The anatomic variants were categorized into four subgroups as follows: (1)
thoracolumbar variant (twelfth rib aplasia/hypoplasia or rudimentary rib of L1), (2) lumbosacral variant (lumbarization or sacralization), (3) numeric variant (four or six lumbar vertebrae without transitional vertebra), and (4) combination of two different variants. Figure 2 summarized the type of anatomic variant. Morphologically normal ribs were defined as a pair of ribs that were 3.8 cm in length or more and originated from the facet between the pedicle and vertebral body. Lumbosacral transitional vertebrae were identified based on the criteria described by Castellvi et al. in 1984. Lumbar vertebrae without rudimentary or normal ribs and showing normal transverse processes were regarded as morphologically normal lumbar vertebrae.

An expert image analyst (S.J.H.) manually generated the GT segmentation map for skeletal muscle area (SMA), visceral fat area (Vfat), and subcutaneous fat area (Sfat). The segmentation map was double-checked by a supervising radiologist (K.W.K.).

Deep learning model development

Our DLM was composed of two algorithms, as follows: (1) a YOLOv3-based algorithm for selecting the L3 slice and (2) a fully convolutional network (FCN)-based algorithm for segmentation. These two algorithms were packaged in a DLM toolkit, named L3SEG-net.

Several preprocessing steps were used to generate input data to increase the effective dataset size and improve overfitting and accuracy. Data augmentation was performed to generate 15,226 maximum intensity projection (MIP) images from 1496 CT scans. Of these, 12,180 MIP images were used as a training set, and 3,046 images were used as a tuning set.

YOLOv3-based L3 slice selection algorithm

A YOLOv3-based algorithm was adopted because YOLOv3 can detect objects and extract features more efficiently than conventional convolutional neural networks, accomplished via object detection and classification. Our YOLOv3-based algorithm generated multiple bounding boxes to extract features from MIP images using a concept similar to feature pyramid networks. The L3 endplate was localized in a MIP image using extracted features of multiple bounding boxes and its relative coordinates. Network architecture and an example of bounding boxes are shown in Fig. 3.

FCN-based segmentation algorithm

Our FCN-based algorithm for automatic segmentation is described elsewhere. We added several post-processing steps to separate the intramuscular adipose tissue from the SMA based on Hounsfield units. The network architecture of our FCN-based algorithm is illustrated in Supplementary Fig. 1. Our FCN-based segmentation algorithm yielded cross-sectional areas (CSAs) of SMA, Vfat, and Sfat in cm² at the selected L3 slice CT images. Currently, the FCN-based segmentation algorithm is available as a web-based iAID toolkit.
Validation of deep learning model

Accuracy of automatic L3 slice selection

In both internal and external validation cohorts, the L3 slice selection accuracy of the YOLOv3-based algorithm was evaluated by the absolute distance difference between the GT and the DLM-derived CT slice. The differences in CT slice numbers between the GT and the DLM-derived results were calculated and multiplied by slice thickness to generate the actual distance difference in millimeters. Technical success was defined when the distance difference between the GT and the DLM-derived results was less than 10 mm (Supplementary Fig. 2). The distance difference and technical success were separately evaluated in the normal anatomy group and anatomic variant group.

Segmentation accuracy of the DLM

In both internal and external validation datasets, the CSA error was calculated to evaluate the accuracy of the DLM-derived segmentation, which is a result of a combination of the YOLOv3-based L3 slice selection and the FCN-based segmentation of abdominal muscle and fat. The CSA error is a standardized percentage difference in measured areas of muscle and fat between the GT values and the DLM-derived values. Thus, a low CSA error implies a high segmentation accuracy. The CSA error was calculated using the following equation:

$$\text{CSA error (\%)} = \frac{|\text{ground truth} \cdot \text{CSA} - \text{DLM} \cdot \text{CSA}|}{\text{ground truth} \cdot \text{CSA}} \times 100$$

In subjects with concordant L3 levels, i.e., identical CT slice numbers from both the GT and the DLM-derived results, the Dice similarity coefficient (DSC) was also used to evaluate the segmentation accuracy of our DLM.

Subgroup analysis according to anatomic variation

The influence of anatomic variation on the performance of the DLM when selecting the L3 slice and segmenting muscle and fat areas were explored by subgroup analysis. The whole validation cohort, i.e., combined internal and external validation cohorts, was divided according to spinal anatomic variations. The accuracy of L3 slice selection and the segmentation accuracy of the DLM was compared between these subgroups.

Statistical analysis

The average values of distance differences between the GT and the DLM-derived L3 slices were compared between the normal anatomy group and the anatomic variant group using a Student t-test. The technical success rate, i.e., the percentage of subjects with technical success among all subjects, was compared between the normal anatomy group and the anatomic variant group using the chi-square test.
The average CSA values of SMA, Sfat, and Vfat were compared between the GT and the DLM-derived results using paired t-tests. The CSA errors were compared between subjects with technical success and subjects with technical failure in the internal and external validation datasets.

Agreement in the measured CSAs of muscle and fat between GT values and DLM-derived values was evaluated with Bland–Altman plots with 95% limits of agreement. R version 3.6.3 (R Foundation for Statistical Computing, Vienna, Austria), MedCalc 12.7.0 (MedCalc Software, Mariakerke, Belgium) were used for statistical analysis. A p-value < 0.05 was regarded as statistically significant.

Results

Accuracy of automatic L3 slice selection

The accuracy of the YOLOv3-based algorithm for automatic L3 slice selection in the internal and external validation datasets is summarized in Fig. 4. The mean distance differences between the GT and the DLM-derived L3 slices were 3.7 ± 8.4 mm and 4.1 ± 8.3 mm for the internal and external validation cohorts, respectively. Subjects with normal spinal anatomy yielded smaller distance differences than those with anatomic variants in the internal (2.5 ± 6.1 vs. 12.4 ± 15.4 mm, p < 0.001) and external (2.8 ± 5.9 vs. 12.1 ± 14.6 mm, p < 0.001) validation sets. The maximum distance difference was 40 mm, equivalent to the height of a vertebral body.

Technical success was achieved for 93.1% (463/496) and 92.3% (541/586) of subjects in the internal and external validation datasets, respectively. The normal anatomy group yielded higher technical success rates than the anatomic variant group in the internal (96.6% vs. 67.2%, p < 0.001) and external (96.2% vs. 67.9%, p < 0.001) validation datasets.

Segmentation accuracy of DLM-derived abdominal muscle and fat areas

The average CSAs of SMA, Sfat, and Vfat derived from GT and DLM are presented in Table 2. In all subjects of internal and external validation datasets, there were no significant differences in CSAs between the GT and DLM-derived measurements in SMA, Sfat, and Vfat (p > 0.05 for all comparisons). Even for subjects with technical failure of L3 slice selection, the CSAs did not differ significantly between the GT and the DLM-derived measurements (p > 0.05 for all comparisons).

The average CSA errors of SMA, Sfat, and Vfat in all subjects of the internal and external validation datasets ranged from 1.38–4.54% (Table 2), indicative of excellent segmentation accuracy of the DLM. When we divided them into subjects with technical success and subjects with technical failure in terms of L3 slice selection, the average CSA errors of subjects with technical failure were higher than those with technical success in both the internal and external validation groups (p < 0.05 for all comparisons).
However, such CSA errors in subjects with technical failure were relatively small in the SMA, compared with the Sfat and Vfat in both the internal and external validation datasets.

In both internal and external validation cohorts, the Bland–Altman plots also demonstrated that agreement of CSAs between the GT and DLM was higher for subjects with technical success than for subjects with technical failure (Fig. 5 and Supplementary Fig. 3).

The mean difference of SMA between GT and DLM-derived results ranged from 0.2 to 3.0 % regardless of technical success on Bland-Altman plot. The mean difference of Sfat ranged from −5.6 to 2.2 %, and Vfat ranged from −3.5 to 1.9 %. The mean differences between GT and DLM-derived results were probably within an acceptable range of measurement variability.

The DSC values in subjects with concordant L3 level between the GT and DLM-derived results were very high. The DSC values of SMA, Sfat, and Vfat were 0.98, 0.98, and 0.98, respectively, in the internal validation dataset and were 0.96, 0.97, and 0.97, respectively, in the external validation dataset.
Table 2
Cross-sectional area segmentation using the ground truth–derived and DLM–derived levels

| Parameter                  | Internal validation dataset | External validation dataset |   |   |   |
|----------------------------|-----------------------------|-----------------------------|---|---|---|
|                            | SMA                         | Sfat                        | Vfat | SMA | Sfat | Vfat |
| All subjects (n = 1082)   |                             |                             |      |     |     |
| CSA from GT (cm²)         | 140.88 ± 34.53              | 140.90 ± 56.71              | 114.53 ± 65.05 | 132.76 ± 31.25 | 133.15 ± 62.16 | 110.59 ± 64.29 |
| CSA from DLM (cm²)        | 140.53 ± 34.20              | 141.98 ± 56.60              | 115.93 ± 65.40 | 130.07 ± 31.07 | 135.54 ± 62.64 | 110.72 ± 65.19 |
| p value*                  | 0.874                       | 0.764                       | 0.736 | 0.139 | 0.492 | 0.973 |
| CSA error (%)             | 1.38 ± 1.46                 | 3.51 ± 5.41                 | 4.00 ± 6.35 | 3.10 ± 2.85 | 4.54 ± 6.34 | 4.26 ± 6.47 |
| Subjects with technical success (n = 1004) |
| CSA from GT (cm²)         | 141.20 ± 34.46              | 138.85 ± 55.86              | 112.42 ± 64.73 | 132.75 ± 31.15 | 133.99 ± 62.82 | 110.88 ± 64.18 |
| CSA from DLM (cm²)        | 140.87 ± 34.06              | 140.47 ± 55.72              | 114.11 ± 64.95 | 130.14 ± 31.00 | 136.73 ± 63.15 | 111.13 ± 65.06 |
| p value*                  | 0.883                       | 0.659                       | 0.692 | 0.167 | 0.474 | 0.950 |
| CSA error (%)             | 1.22 ± 1.08                 | 2.31 ± 2.21                 | 2.97 ± 3.21 | 2.86 ± 2.57 | 3.39 ± 2.78 | 3.36 ± 4.68 |
| Subjects with technical failure (n = 78) |
| CSA from GT (cm²)         | 136.33 ± 35.18              | 169.78 ± 60.54              | 144.23 ± 62.27 | 132.97 ± 32.3 | 123.03 ± 52.30 | 107.10 ± 65.61 |
| CSA from DLM (cm²)        | 135.77 ± 35.80              | 163.24 ± 64.41              | 141.53 ± 66.37 | 129.21 ± 31.93 | 122.66 ± 54.42 | 105.78 ± 66.50 |
| p value*                  | 0.949                       | 0.672                       | 0.865 | 0.579 | 0.974 | 0.924 |
| CSA error (%)             | 3.68 ± 3.19                 | 20.42 ± 8.14                | 18.37 ± 15.68 | 6.01 ± 4.18 | 18.28 ± 15.16 | 15.06 ± 12.56 |
| p value§                  | < 0.01                      | < 0.01                      | < 0.01 | < 0.01 | < 0.01 | < 0.01 |
| Parameter | Internal validation dataset | External validation dataset |
|-----------|-----------------------------|-----------------------------|
|          | SMA | Sfat | Vfat | SMA | Sfat | Vfat |

Note.—Data are presented as mean ± standard deviation

* The p-value is calculated from Student t-test comparing the GT CSA and the CSA determined using the DLM.

§ The p-value is calculated from Student t-test comparing CSA errors between subjects with technical success and subjects with technical failure.

CSA = cross-sectional area, DLM = deep learning model, GT = ground truth, Sfat = subcutaneous fat area, SMA = skeletal muscle area, Vfat = visceral fat area

Subgroup analysis according to anatomic variation

Anatomic variation significantly influenced L3 slice selection by the DLM. The technical success rate was highest in the normal anatomy group (96.5%), followed by the thoracolumbar variation (82.6%), lumbosacral variation (63.9%), numeric variation (54.5%), and combined variation (40%) subgroups. The mean distance differences were 2.6, 7.4, 13.4, 16.5, and 21.4 mm for the normal anatomy, thoracolumbar variation, lumbosacral variation, numeric variation, and combined variation groups, respectively.

Regarding the CSA errors, anatomic variation significantly influenced Sfat and Vfat measurement, with CSA error higher than 5%, while less significantly influenced SMA measurement, with CSA error less than 5% (Table 3). Specifically, the average CSA errors between GT and DLM-derived results were 2.22% in normal anatomy subgroup and ranged from 2.37–4.06% in subgroups with anatomic variations.
Table 3
Subgroup analysis according to spine anatomy

| Subgroup                        | Distance difference (mm) | Technical success (%) | CSA error (%) | Bland-Altman (mean ± limits of agreement) |
|---------------------------------|--------------------------|-----------------------|---------------|------------------------------------------|
|                                 |                          |                       | SMA           | Sfat                                      | Vfat | SMA | Sfat | Vfat |                       |
| Normal anatomy (n = 943)        | 2.6 ± 6.0                | 96.5                  | 2.22 ± 2.46   | 3.46 ± 4.78                              | 3.57 ± 5.58 | 1.68 ± 7.22 | -2.29 ± 13.13 | -0.84 ± 10.03 |
| Thoracolumbar variation (n = 46) | 7.4 ± 11.9               | 82.6                  | 2.73 ± 2.24   | 5.83 ± 8.79                              | 5.87 ± 7.04 | 2.23 ± 7.90 | 2.41 ± 34.33 | -2.69 ± 24.10 |
| Lumbosacral variation (n = 72)  | 13.4 ± 15.2              | 63.9                  | 3.04 ± 2.49   | 8.72 ± 10.63                             | 7.94 ± 9.23 | 1.40 ± 10.56 | 2.17 ± 35.93 | 0.86 ± 24.84  |
| Numeric variation (n = 11)      | 16.5 ± 16.1              | 54.5                  | 2.37 ± 2.11   | 10.87 ± 7.62                             | 10.36 ± 10.19 | -0.22 ± 7.10 | -3.93 ± 46.02 | -1.82 ± 26.79 |
| Combined variation (n = 10)     | 21.4 ± 17.0              | 40                    | 4.06 ± 2.92   | 11.86 ± 12.66                            | 14.95 ± 17.03 | -2.53 ± 14.78 | -7.15 ± 58.06 | 10.82 ± 67.40 |

Note.—CSA = cross-sectional area, Sfat = subcutaneous fat area, SMA = skeletal muscle area, Vfat = visceral fat area.

Discussion

We were able to develop the L3SEG-net, a fully automatic DLM for selecting axial CT slice at L3 vertebral level and segmenting abdominal muscle area in an end-to-end manner. The L3 slice selection accuracy was accurate with mean distance differences less than 5 mm between GT and DLM-derived results. The overall segmentation accuracy of abdominal muscle areas was also excellent, with the average CSA errors of 1.38–3.10 cm² between GT and DLM-derived results.

There are several unique characteristics in the L3SEG-net. First, the L3SEG-net is composed of two algorithms running sequentially as one process: a YOLOv3-based L3 slice selection algorithm and a FCN-based segmentation algorithm. When we upload one or multiple series of full abdominal CT images in the L3SEG-net, it automatically selects L3 slice CT images, segments muscle and fat areas, and provides color maps with measurement values. The L3SEG-net can process approximately 1,000 abdominal CT scans per day in a setting of Intel® Core™ i7–7700K GPU (8M Cache, 4.20 GHz, Santa Clara, CA, USA). Thus, the L3SEG-net can be helpful to perform large-scale researches.

Second, we trained the L3SEG-net for L3 slice selection with accurate information of anatomic variations. To identify the anatomic variations accurately, we obtained chest CT and abdominal CT scans in almost all training and validation cases and counted number of all thoracolumbar spines and ribs. Thus, the
L3SEG-net is a unique model which can spotting L3 slice level with consideration of anatomic variations. Nevertheless, the normal anatomy group yielded much higher technical success rates than the anatomic variant group in the internal (96.6% vs. 67.2%) and external (96.2% vs. 67.9%) validation datasets. Among the abnormal variant subtypes, the thoracolumbar junction variant subgroup, including T12 rib hypoplasia/aplasia and L1 rudimentary rib, yielded similar performances to the normal anatomy group, whereas the lumbosacral junction variant subgroup and other numeric variant subgroup yielded lower technical success rates. The lower technical success of the lumbosacral junction variant subgroup may be attributable to our training process component to make the algorithm assume the iliac create as the L4 level. In near the future, we will keep training the L3SEG-net for automatic spine labelling using further data.

Third, we demonstrated that the L3SEG-net's overall segmentation accuracy of muscle areas is accurate regardless of anatomic variation in both internal and external validation cohorts. We used CSA error as a representative value of segmentation accuracy, instead of DSC. DSC evaluation was limited on the group showed the same CT slice of GT and L3SEG-net selection. Then DSC value can present only accuracy of segmentation algorithm. Thus we suggested CSA error as an indicator reflecting accuracies of both L3 selection algorithm and segmentation algorithm, regarding clinical impact. The average CSA errors between the GT and DLM-derived results were 2.22% in normal anatomy subgroup and ranged from 2.37–4.06% in subgroups with anatomic variations. These results may be attributable that the distance difference between GT and DLM was less than the height of a vertebral body, as the maximum distance difference was 40 mm. According to a recent study, the muscle area measurements were similar between the L2 inferior endplate level and L4 inferior endplate level.

Overall segmentation accuracy of SMA was consistent regardless of CT parameters or machine. The results were reported in prior study. Various CT machine and parameters from four other hospital were used in this study, but only portal phase abdominal CT scans were used for the analysis. The segmentation accuracy was consistent measuring SMA, Vfat and Sfat.

There have been two prior studies which reported performance of automatic L3 level slice selection models. However, these studies did not consider the anatomic variations in the training and validation process. Belharbi et al. compared the performances of various convolutional neural networks (CNNs) for L3 slice selection with a dataset of 642 CTs of a single institution. The mean distance difference was 1.8 to 10.5 CT slices, equivalent to 3.6 to 50.5 mm. This study was limited to the task of L3 slice selection and did not have segmentation algorithm. Bridge et al. reported deep learning models for the L3 slice selection and automatic segmentation, developed based on a training cohort (n = 595) and a testing cohort (n = 534). The mean localization error was 9.4 mm. Compared to these two prior studies, our L3SEG-net showed higher accuracy in L3 slice selection.

Our study had some limitations. First, we used a relatively small size of data for training and validation of L3SEG-net deep learning model. Thus, we plan to develop a sustainable training system and keep training our L3SEG-net model using prospectively collecting CT images. Second, healthy subjects were only
included for the internal and external validation cohorts. The performance of the developed DLM may require validation with large samples of patients with various diseases.

**Conclusion**

In conclusion, our new deep learning model, L3SEG-net, was developed for the selection of an axial CT slice at the L3 vertebral level and the segmentation of abdominal muscle areas in an end-to-end manner. The L3SEG-net performed well regardless of anatomic variations with high L3 slice selection accuracy and segmentation accuracy. The L3SEG-net will be open for non-profit research as a web-based toolkit with hope that it can help large scale sarcopenia research.

**Abbreviations**

CNN = Convolutional neural network, CT = computed tomography, DSC = Dice similarity coefficient, CSA = cross-sectional area, DLM = deep learning model, FCN = fully convolutional network, GT = ground truth, MIP = maximum intensity projection, Sfat = subcutaneous fat area, SMA = skeletal muscle area, Vfat = visceral fat area.

**Declarations**

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**Conflicts of interest statement and funding:**

1. Kim KW, Park T, Khang S, Jeong H, Koo K, Lee J, Sung YS, and Shin Y are inventors on patent issued by the Korean Intellectual Property Office (KR patent application No. 10-2018-0035284). All other authors declare no conflicts of interest.

2. Kim KW, Park T, Khang S, Jeong H, Koo K, Lee J, Shin Y are inventors on patent issued by the Korean Intellectual Property Office (KR patent application No. 10-2019-0009323). All other authors declare no conflicts of interest.

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**Author Contributions:**

Study concept and design: All authors.

Acquisition of data: JH, YK, Jiwoo Lee, SJH.
Software: TP, YS, YSS, SK, HJ, KK, Jeongjin Lee.

Statistical analysis and interpretation of data: JH, TP, YK, KKW.

Drafting of the manuscript: JH, TP

Critical revision of the manuscript for important intellectual content: KKW, Jeongjin Lee, KDW

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**Figures**

**Figure 1**
An overview of dataset composition.

| Normal Anatomy | Thoracolumbar Variant | Lumbosacral Variant | Numeric Variant | Combined Variation |
|----------------|-----------------------|---------------------|-----------------|-------------------|
| • Five morphologically normal lumbar vertebrae | • T12 rib hypoplasia or aplasia | • Lumbarization of S1 | • Four or six morphologically normal lumbar vertebrae without transitional vertebra T12 hypoplasia/aplasia | • Combined anatomic variation |
| • Rudimentary rib attached to L1 vertebra | | • Sacralization of L5 | | |

Table

- e.g., five morphologically normal lumbar vertebrae
- e.g., T12 rib hypoplasia
- e.g., sacralization of L5
- e.g., six morphologically normal vertebrae
- e.g., T12 rib hypoplasia and L5 sacralization

**Figure 2**

Anatomic lumbar spine variants. Examples of normal, thoracolumbar, lumbosacral, numeric, and combined variations are presented.
Figure 3

Example of multiple bounding boxes for training of the YOLOv3-based model and architecture of our YOLOv3-based network. Multiple bounding boxes were generated in the maximum intensity projection images based on the following prerequisites as illustrated in (A): (1) the L4 vertebra was located at the iliac crest level, (2) the L3 vertebra was located superiorly to the L4 vertebra, (3) the morphologies of the lumbar vertebrae were the same. The YOLOv3-based model used an objectness score for each bounding
box obtained from logistic regression to predict the width and height of the box as well as its location relative to grid cell. The sum of the squared error loss was used to train the model for minimizing differences between the ground-truth object and the bounding box. Any error between the bounding box over the ground-truth object was incurred for both classification and detection loss. Our model extracted features of the bounding boxes using the network architecture illustrated in (B). Our network architecture used successive 3×3 and 1×1 convolution layers and a set of residual blocks with shortcut connections. A total of 53 convolutional layers were formed like Darknet-53. YOLOv3 predicted boxes at three different scales to support detection on varying scales.

![Box plots of distance difference between ground truth and deep learning model (DLM) derived results in (A) internal validation dataset and (B) external validation cohorts.](image)

**Figure 4**

Box plots of distance difference between ground truth and deep learning model (DLM) derived results in (A) internal validation dataset and (B) external validation cohorts. The mean differences between the ground truth and the DLM-derived results were 3.7 mm ± 8.4 and 4.1 mm ± 8.3 for the internal, and external validation cohorts, respectively.
Figure 5

Bland Altman plots to evaluate agreement of SMA between the GT and DLM (A) In subjects with technical success in the internal validation cohort (B) In subjects with technical failure in the internal validation cohort (C) In subjects with technical success in the external validation cohort (D) In subjects with technical failure in the external validation cohort

Supplementary Files

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- SupplementaryTableandfigures0616.docx