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Developing and Evaluating Deep Learning Algorithms for Object Detection: Key Points for Achieving Superior Model Performance

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In recent years, artificial intelligence, especially object detection-based deep learning in computer vision, has made significant advancements, driven by the development of computing power and the widespread use of graphic processor units. Object detection-based deep learning techniques have been applied in various fields, including the medical imaging domain, where remarkable achievements have been reported in disease detection. However, the application of deep learning does not always guarantee satisfactory performance, and researchers have been employing trial-and-error to identify the factors contributing to performance degradation and enhance their models. Moreover, due to the black-box problem, the intermediate processes of a deep learning network cannot be comprehended by humans; as a result, identifying problems in a deep learning model that exhibits poor performance can be challenging. This article highlights potential issues that may cause performance degradation at each deep learning step in the medical imaging domain and discusses factors that must be considered to improve the performance of deep learning models. Researchers who wish to begin deep learning research can reduce the required amount of trial-and-error by understanding the issues discussed in this study.

Keywords: Deep learning; Object detection; Diseases with small sizes; Disease subclass; Image modality; Deep learning workflow; Data augmentation; Hyperparameter optimization

INTRODUCTION

Recently, deep learning technologies in computer vision have rapidly developed owing to the advances in and widespread use of graphic processor units optimized for parallel operation [1]. Object detection [2] is a deeplearning task that simultaneously identifies the location and label of a target object. Interesting results for object detection have been reported in various studies, such as

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This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (https://creativecommons.org/licenses/by-nc/4.0) which permits unrestricted non-commercial use, distribution, and reproduction in any medium, provided the original work is properly cited. face detection [3], recognition [4], pedestrian detection [5], and car detection [6]. Furthermore, object detection has been applied in the medical imaging domain, which has shown remarkable results in developing models to predict lesions, such as brain cancer [7], liver disease [8], and wrist, rib, and pediatric skull fractures [9–12] using various imaging modalities, such as radiography, computed tomography (CT), and magnetic resonance imaging (MRI).

While majority of deep learning studies in the medical image domain have demonstrated remarkable results, certain approaches have exhibited poor performance [13]. In a previous study [14], deep-learning-based false-positive reduction demonstrated lower performance than rulebased false-positive reduction. Performance degradation can be caused by various factors, such as insufficient data, unoptimized hyper-parameters, or the application of an incorrect evaluation strategy. However, it is often difficult to understand the cause of poor performance because humans cannot understand the intermediate process of the deep learning model owing to the black-box problem



[15]. Moreover, conducting comparative experiments on all variables that can affect deep learning performance has practical limitations.

The purpose of this article is to explain and demonstrate the key considerations for applying object detection to the medical imaging domain across each step of deep learning research. These considerations are typically performed in the following order: target disease selection, data collection, data labeling, deep learning network training, and performance evaluation (Fig. 1). We hope that this article can help junior researchers who aim to apply object detection in the medical image domain to understand the potential issues that may occur at each step and efficiently conduct their research by reducing trial-and-error.

Target Disease Selection

Suitability of Object Detection

Deep learning techniques in medical imaging analysis can be categorized into classification, object detection, and segmentation. Examples of previous studies that have applied these three deep learning methods in medical image analysis are summarized in Supplementary Table 1, and an example explaining the differences between the three methods are shown in Figure 2.

Object detection (Fig. 2A) detects trained objects in



Fig. 1. Flowchart of the process for deep learning research by applying the issues introduced in this study. The flowchart organizes the issues that should be considered when conducting deep learning research in a sequential manner, and categorized into target disease selection, data collection, data labeling, network training, and performance evaluation. 2D = two dimensional, 2.5D = two and a half dimensional, 3D = three dimensional, CT = computed tomography, MRI = magnetic resonance imaging, E2E = end-to-end



Fig. 2. An example showing the differences between object detection, classification, and segmentation. The object detection method **(A)** shows the result with a blue bounding box with the label "BM", the classification method **(B)** presents the result with the label "Brain Metastases", and the segmentation method **(C)** displays the result on the image using a red mask. BM = brain metastases.

an image using a bounding box or circle with its class. In contrast, the classification method (Fig. 2B) determines the class of the entire input image. Object detection offers the advantage of being able to identify multiple lesions in an image and does not suffer from the multilabel classification problem [16]. For example, previous studies have utilized object detection methods to detect lesions such as brain metastases [17], liver lesions [18], maxillary sinusitis [19], and cerebral microbleeds [14,20]. Other studies have applied it to identify the location of an object in an image, such as facial region [19], wrist region [9], and various organs, to extract a patch image. As shown in Figure 2C, the segmentation method yields results at the pixel level and is suitable for detecting lesions that require pixel-level evaluation, such as measuring the volume of the lesion [21,22] or supporting a radiotherapy plan. However, the cost of labeling is higher than that of the object detection method. Therefore, unless an evaluation in units of pixels is required, the object detection method is appropriate for detecting visible diseases or lesions in images, where the evaluation of the location information contained in images can help recognize each lesion separately.

Target Disease with a Small Size

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Small-object detection is a fundamental challenge in computer vision [23]. A small object is defined as an object with a size less than or equal to 32 x 32 pixels [24], which results in issues such as indistinguishable features, low resolution, and limited context information, making it difficult to detect the target object [25]. Therefore, using an object detection algorithm for detecting diseases with small sizes, such as small calcifications and early-stage cancers, may result in poor performance. Examples of previous studies that demonstrated lower performance on small lesions than on large lesions are listed in Table 1 [17,22,26-28]. Zhou et al. [17] reported that four deep-learning networks exhibited lower sensitivity (10%-40%) in detecting brain metastases smaller than 3 mm. In another study on the detection of breast calcifications, Akselrod-Ballin et al. [26] reported that removing calcifications with radii smaller than 10 pixels can significantly improve performance. To address the challenges of detecting small objects, applying the patch process to increase the proportion of lesions in an image may improve performance [29]. Moreover, using specialized models for small objects, such as M2Det [30], multi-scale deconvolutional single-shot detector (SSD) [31], and improved faster region-based convolutional neural network (R-CNN) for small object detection [32], which have been recently published, may enhance the detection of small lesions. However, to the best of our knowledge, these models have not yet been applied in the medical imaging domain.

Identifying Data Distribution of Subset Groups

Certain target diseases in the medical imaging domain can be sub-classified. For example, maxillary sinusitis can be sub-classified into full opacification, air/fluid level, cysts, and mucosal thickening [19]; cancer labels can be grouped based on lesion size [17]; and metastases can be grouped according to their origin. If the object detection model is trained with integrated labels, the performance for each subclass might vary owing to the differences in the lesion features and amount of training data. In addition, the

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d Lesions	Contents	Four deep learning networks showed lower sensitivity (10%-40%) for detecting brain metastases smaller than 3 mm while higher sensitivity (92%-98%) for larger than 6 mm.	The performance of the proposed models for the INBreast dataset yielded a significant improvement by the removal of small calcification (radius < 10 pixels).	Two deep learning models that performed the best did not recognize any parts of the tumors in five cases with a size smaller than 1.5 cm^3 .	The sensitivity of the CE + NECT model was 41.5% for < 3 mm lesions; 81.6% for $3-6$ mm lesions; and 96.5% for ≥ 6 mm lesions.	45% of nodules smaller than 2.0 cm were detected by the algorithm, while 63% of nodules larger than 2.0 cm were detected.
g Small-sized	Lesion Size	< 3 mm	< 10 pixels	< 1.5 cm ³	< 3 mm	≤ 2.0 cm
1 to Lesion Detection, Includin	Algorithm	SSD SSD + ResNet50 SSD + focal loss RetinaNet	Faster R-CNN	Attention and 2D U-net with GMM	SSD	Commercially available deep learning-based algorithm (Lunit INSIGHT CXR; Lunit)
p Learning Approacl	Deep Learning Task	Object detection	Object detection	Segmentation	Object detection	1
s that Applied a Dee	Target Disease	Brain metastases	Small calcifications	Meningioma	Brain metastases	Lung cancer
of Previous Studie	Modality	MRI	Breast mammography	MRI	CL	Chest radiography
able 1. Examples	Studies	hou et al. [17]:	kselrod-Ballin et al. [26]	ang et al. [22]	akao et al. [27]	lam et al. [28]

integrated performance of the test dataset may be altered if the composition of the subclass is different from that of the training dataset. If the composition of the subclass with lower accuracy is higher in the external validation set, the performance may be lower than that of the internal validation set, which can be mistakenly considered as the result of overfitting (Fig. 3). Therefore, the ratio of the subclasses should be verified during the composition of the test dataset, and the performance of each subclass must be investigated.

Data Collection

Evaluation of Diminished Performance Owing to Insufficient Data

Previous studies that applied the deep learning approach used various amounts of data (Supplementary Table 1). A deep learning network is trained with features from the data; therefore, training with more data can improve performance. However, data collection in the medical image domain is difficult compared with that in the general image domain, and studies that apply deep learning are often performed using a limited amount of data. Hence, the guestion "How much data is sufficient?" is commonly asked by researchers interested in artificial intelligence, and the amount of data required may vary depending on the target disease and imaging modality. For example, if the target disease has numerous variables, such as various sizes, locations in different regions, and varying lesion textures, specifying the amount of data required is difficult.

To address this issue, it is possible to evaluate whether the data are insufficient. If the target disease is determined and a certain amount of data is collected at the initial stage of the study, the relationship between the amount of data and performance of the deep learning model can be investigated by training and evaluating each model while increasing the amount of training data (Fig. 4). By estimating the amount of data required, a researcher can determine whether the amount of data for the study must be increased and how much data must be collected. For example, Cho et al. [33] investigated the relationship between accuracy and the amount of training data. Their estimation results predicted 98% accuracy for a training data size of 1000 per body class.

Single-Slice Images Such as Radiographs and Patch Process

A radiography image is relatively large compared with

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MRI = magnetic resonance imaging, SSD = single-shot detector, R-CNN = region-based convolutional neural network, GMM = Gaussian mixture model, CT = computed tomography,

+ NECT = contrast-enhanced + non-enhanced computed tomography

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Fig. 3. Simple example of the data imbalance issue for a subclass. A: The scenario where subclass data imbalance can be misinterpreted as overfitting. (B-E) The predicted results of the object detection model for cyst cases, which are a subclass of sinusitis. The sample images for the internal (B, C) and external (D, E) datasets indicate ground truth and predicted results as blue and yellow bounding boxes, respectively. ACC = accuracy



Fig. 4. Relationship between the amount of data and performance of the object detection deep learning network for brain metastases. The black dots represent the number of data used for model training and their corresponding sensitivity. The blue dashed line represents the trend line of the black dots, and the black dashed line represents the number of data required to achieve a 95% sensitivity. The results, shown as a logarithmic function, indicate that the sensitivity of the deep learning model increases as the amount of input data increases. In this example, to reach a sensitivity of 95%, the researcher can predict that they need the data of approximately 54 participants.

the average size of an image in ImageNet (approximately 1600-2000 pixels in the horizontal and vertical axes vs. approximately 400 x 350 pixels, respectively) [34]. A large image size significantly increases the computational power required and may also result in a dimensionality problem [35]. Additionally, radiographs usually contain a substantial portion of background that is unrelated to the diagnosis of the disease. Applying the patch process, which crops only the essential part of the image, can produce a cropped image without the loss of lesion information owing to shrinking, and unnecessary parts, such as the background, can be removed. However, the patch process has traditionally been performed by humans. In the medical imaging domain, certified radiologists or other medical doctors performed the manual patch process, which is time-consuming. As a result, the associated costs are considerably high.

Recently, an automated patch process was applied in the medical imaging domain by employing the object detection approach to address the disadvantages of large images, such as radiographs. Table 2 lists previous studies that used a manual patch process or applied deep learning approaches [9,19,36-42]. Previous studies [9,19,39-42] have applied an

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Table 2. Examples of Deep Learning Approach-based Studies that Applied the Patch Process

StudiesModalityTarget Disease or Object.Patch RegionPatch MethodContents1:1-151Wate's viewMarklay sinustifyFacial regionHard-caftAnadcafted patch process induring oth maculing vinusse.1:1-151Wate's viewMarklay sinustifyHard-caftHard-caftAnadcafted patch process induring oth maculing vinusse.1:1-151Wate's viewMarklay sinustifyHard-caftHard-caftHard-caftHard-caft1:1-151Wate's viewMarklay sinustifyHard-caftHard-caftHard-caftHard-caft1:1-151Hard ArbHard caftHard-caftHard-caftHard-caftHard-caft1:1-151Hard ArbHard caftHard-caftHard-caftHard-caftHard-caft1:1-151Hard ArbHard caftHard-caftHard-caftHard-caftHard-caft1:1-151Hard ArbHard CaftHard-caftHard-caftHard-caftHard-caft1:1-151Hard CaftHard-caftHard-caftHard-caftHard-caftHard-caft1:1-151Hard Vate'sHard-caftHard-caftHard-caftHard-caftHard-caft1:1-151Hard Vate'sHard-caftHard-caftHard-caftHard-caftHard-caft1:1-151Hard Vate'sHard-caftHard-caftHard-caftHard-caftHard-caftHard-caft1:1-151Hard Vate'sHard-caftHard-caftHard-caftHard-caftHard-caftHard-caftHard-caft <trr>1:1-151</trr>						
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td. [37]Water's viewMaxillary sinustisMaxillary sinustisMand-caftMand-caftS40 Water's view images were cooped to 7 x 7 cm based ontd. [38]CTRenal calculMand-caftMand-caftMand-caftMand-caftMand-cafttd. [31]Hand AP, PAHandRenal calculMand-caftMand-caftMand-caftMand-cafttd. [32]Hand AP, PAHandHand maskMand-caftMand-caftMand-caftMand-cafttd. [33]Hand AP, PAHandHand maskMand-caftMand-caftMand-caftMand-cafttd. [34]Mand-PDWrist radiographsMand-caftMand-caftMand-caftMand-cafttd. [34]Mand-PDMand-PDMand-PDMand-caftMand-caftMand-cafttd. [34]Mand-PDMand-PDMand-PDMand-caftMand-caftMand-cafttd. [30]Mand-PDMand-PDMand-PDMand-PDMand-PDMand-cafttd. [30]Mand-PDMand-PDMand-PDMand-PDMand-PDMand-PDtd. [30]Mand-PDMand-PDMand-PDMand-PDMand-PDMand-PDtd. [40]Mand-PDMand-PDMand-PDMand-PDMand-PDMand-PDtd. [41]Mand-PDMand-PDMand-PDMand-PDMand-PDMand-PDtd. [42]Mand-PDMand-PDMand-PDMand-PDMand-PDMand-PDtd. [43]Mand-PDMand-PDMand-PDMand-PDMand-PDMan	et al. [36]	Water's view	Maxillary sinusitis	Facial region	Hand-craft	A handcrafted patch process including both maxillary sinuses was performed for 5020 Water's view images.
tal. [38]CTRend. cucleRend. cucleR	et al. [37]	Water's view	Maxillary sinusitis	Maxillary sinus	Hand-craft	9540 Water's view images were cropped to 7 x 7 cm ² based on the central coordinates of the bilateral maxillary sinuses.
tal. [39]Hand AP, FAHandHand maskDispectedImages were copped based on the hand mask patch generated by multiple-processing steps, including normalization, object detection on hand radiographs, and reconstruction.n et al. [30]Wrist radiographsWrist radiographsWrist radiographs, and reconstruction.n et al. [41]Wrist radiographsWrist regionMachine learningPeep tearningtar.MamogramBreast cancerMast regionMachine learningPeep tearningtar.MamogramBreast cancerMast regionMachine learningPeep tearningtar.MamogramBreast cancerMast regionMachine regions from the trained YOL0 model for masstar.MamogramBreast cancerMast regionMachine regions from the trained YOL0 model for masstar.MamogramBreast cancerMast regionMachine regions from the trained YOL0 model for masstar.MamogramBreast cancerMast regionMachine regions from the trained YOL0 model for masstar.Galowell and Water'sSinusitisPeep tearningInput images had cropped based on bounding power whichet al. [41]ViewSinusitisPeep tearningInput images had cropped based on bounding power whichet al. [42]ViewSinusitisPeep tearningInput images were used for tho califring each sinus area.et al. [43]Water's viewMacrist sinus sinus sinus area.Input images were used for tho califring each sinus area.et al. [43]Water's viewMacrist sinus sinus sinu	et al. [38]	L	Renal cancer	Renal cell carcinoma region	Hand-craft	Images were cropped based on the region of interest for the renal cell carcinoma lesion drawn by the radiologist.
net al. [9]Wist fractureWist fra	it al. [39]	Hand AP, PA	Hand	Hand mask	Object detection-based preprocessing engine	Images were cropped based on the hand mask patch generated by multiple-processing steps, including normalization, object detection on hand radiographs, and reconstruction.
LitManmogramBreast cancerMass regionDep learningThe predicted regions from the trained YOL0 model for massLitLitManmogram(object detection)(object detection)(bep learning)detection were cropped for the mass segmentation stage.et al. [41]Caldwell and Water'sSinusitisFontal, maxillary,Deep learningmant images had cropped based on bounding boxes which were predicted from a detector for localizing each sinus area, and ethmoidet al. [41]ViewSinusitisInput images were used for the second network that cancellaring each sinus area, and ethmoidet al. [42]Water's viewMaxillary sinusitisDeep learninget [42]ManmogramManmographyPhantom regionDeep learninget al. [42]ManmogramManmographyPhantom regionDeep learningetal [42]ManmogramManmographyPhantom regionDeep learningetal [42]ManmogramManmogramPhantom regionPhantom region to extract a patch for the next step.	1 et al. [9]	Wrist radiographs (PA and LAT)	Wrist fracture	Wrist region	Machine learning	The patch process consisted of global search (RFRV) and local search (RFCLM).
et al. [41]Caldwell and Water's oviewSinusitis and ethmoid and ethmoid sinusitisDeep learning object detection object detection classifies each sinus patch with diagnostic labels.or [19]Water's viewMaxillary sinusitis sinusitisDeep learning object detectionInput images had cropped based on bounding boxes which and cropped images were used for the second network that classifies each sinus patch with diagnostic labels.al. [19]Water's viewMaxillary sinusitisFacial regionDeep learning object detectionIne YOLD v2 detection network was applied to detect the facial region that includes both maxillary sinuses.al. [42]MamogramMamographyPhantom regionDeep learning (object detection)The YOLD v2 detection network was applied to detect the region that includes both maxillary sinuses.	tari ıl. [40]	Mammogram	Breast cancer	Mass region	Deep learning (object detection)	The predicted regions from the trained YOLO model for mass detection were cropped for the mass segmentation stage.
tal. [19] Water's view Maxillary sinusitis Facial region Deep learning The YOLO v2 detection network was applied to detect the facial region tal. [42] Mammogramy Mammogramy Phantom region Deep learning The YOLO v2 detection network was applied to detect the facial region tal. [42] Mammogramy Mammogramy Phantom region Deep learning The YOLO v2 detection network was applied to detect the value of the next step.	et al. [41]	Caldwell and Water's view	Sinusitis	Frontal, maxillary, and ethmoid sinusitis	Deep learning (object detection)	Input images had cropped based on bounding boxes which were predicted from a detector for localizing each sinus area, and cropped images were used for the second network that classifies each sinus patch with diagnostic labels.
: al. [42] Mammogram Mammography Phantom region Deep learning The YOLO v2 detection network was applied to detect the phantom image (object detection) phantom region to extract a patch for the next step. evaluation	: al. [19]	Water's view	Maxillary sinusitis	Facial region	Deep learning (object detection)	The YOLO v2 detection network was applied to detect the facial region that includes both maxillary sinuses.
	: al. [42]	Mammogram	Mammography phantom image evaluation	Phantom region	Deep learning (object detection)	The YOLO v2 detection network was applied to detect the phantom region to extract a patch for the next step.



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automated patch process based on deep learning or machine learning approaches, such as facial region detection [19], phantom region detection [42], radius location identification [9], and mass detection [40], as pre-processing to limit the area of the disease location, reducing the loss of lesion information caused by shrinking and removing the background that is unrelated to the disease.

Multi-Slice Images Such as CT or MRI

CT and MRI scans involve acquiring a multi-slice image instead of a single image and using 2D, 2.5D, or 3D methods depending on the type of image input required for the deep learning model. Compared with the 3D method, a 2D-based deep learning network can be used with a larger image as an input, for example, 256 x 256 or 512 x 512 images [14,17]. However, besides extremely small lesions, most target lesions are located over multiple image slices. The 2D deep learning model predicts the lesion in each image individually; therefore, to evaluate the lesion on a mass unit, post-processing that evaluates the lesion in an adjacent slice as a mass must be performed [18]. The 3D method has the advantage that it can train a deep learning model using more information from adjacent slices, and its performance can be improved. The disadvantage of the 3D method is that it requires a significant amount of computational power; therefore, small-sized input data are generally used [43]. The 2.5D method uses a 2D-based deep learning model in three orthogonal directions—coronal, sagittal, and axial—and can improve the performance of a deep learning algorithm through a majority decision step with three deep learning networks. However, the labeling process must be performed for all three directions, and post-processing for labeling must be performed to apply the label to all three directions. Moreover, a majority decision step must be performed during additional post-processing [44].

Single- and Multi-Channel Input Data

Special imaging techniques, such as dual-energy CT [45] and MRIs with multiple different sequences [46], have made it possible to acquire the same images with different intensities. Examples of previous studies that have investigated multi-channel input data for the deep-learning approach are listed in Table 3 [47-51]. Using images with different intensities as multi-channel input data, the deep learning model can be trained with more data and patterns, and its performance can be improved [49]. For example, if images before and after using a contrast agent are used

as a multichannel input, the performance of the deep learning model can be improved by recognizing the intensity differences before and after using the contrast agent [52].

However, using multi-channel data for performance improvement does not guarantee a statistically significant difference [48]. In a scenario where a model is being developed for detecting lesions, if images that are not related to the diagnosis of the lesion are included in the multi-channel input data, the required computational power increases, which increases the dimension of the data. This, in turn, may lead to a decrease in the performance of the deep learning network [47]. Therefore, researchers who aim to utilize multi-channel data in developing deep learning models should exclude unnecessary data during the data collection process and evaluate the performance using data that can affect the performance.

Data Labeling

Labeling Verification for Ambiguous Objects

Object detection requires training with the correct labels, and its performance may decrease when the training data include noisy or incorrect labels. According to Rolnick et al. [53], the performance of a classification model using the ImageNet dataset with 5% incorrect labels decreases by approximately 20%.

In the medical imaging domain, the labeling process is typically performed by a radiologist, and most lesions are clearly labeled. However, several lesions may be ambiguous to diagnose or label, and using labels for ambiguous lesions in training and evaluation can lead to performance degradation. This degradation can be overcome by verifying ambiguous labels. Verification can be performed by comparing with other imaging modalities or biopsy results, and labels can be determined by aggregating the opinions of several raters, which reduces ambiguity. In a previous study, Kim et al. [37] used paranasal sinus CT scans as the reference standard for sinusitis to compare the overall diagnostic performance of a deep-learning algorithm with that of radiologists. In another study [54], subtype labels were confirmed by the pathological examination of surgically removed tumors to diagnose kidney cancer.

Labeling Small Objects

In object detection, the labeling process is typically performed by drawing a bounding box or circle around a target object. For small objects, such as cerebral microbleeds

Table 3. Example:	s of Previous Sti	udies that Compared Mul	ti-channel Input Data for the Deep Learning Approach	H:	
Study	Imaging	Purpose	Comparison Models	Performance	Comments
Al-masni et al. [47]	Brain MRI	Cerebral microbleeds detection	One-channel: SWI Two-channel: SWI + Phase Three-channel: SWI + Phase + Magnitude Two-channel: SWI and Complement phase Two-channel: SWI and Complement phase with averaging of adjacent slices*	91.67 97.22 88.89 94.44 100.00* (Sensitivity)	
Fei et al. [48]	Brain MRI	Synthesize the FLAIR modality	T1 T1 + T2* T1 + T2 + T1c models*	23.70 ± 2.16, 0.86 ± 0.02 24.80 ± 1.85, 0.88 ± 0.02* 24.93 ± 1.96*, 0.87 ± 0.02 (PSNR, SSIM)	The triple-input model achieved the highest PSNR values, and the T1 + T2 group achieved the highest SSIM values.
Feng et al. [49]	Breast MRI (DWI, DCE)	Classifying the breast cancer	OA patch PT patch AP-DCE patch DCE ensemble*	70.0 78.2 80.0 84.6* (Accuracy)	The classification performance of multi-channel input was always better than using only a single-channel input.
Chen et al. [50]	Brain MRI	Brain segmentation for GM, WM, CSF	T1 T1-IR T2-FLAIR All All + auto-context*	86.96, 89.70, 79.58 80.61, 85.89, 76.44 81.13, 83.21, 75.34 88.08, 90.93, 82.51 88.50, 91.06, 82.70* (Dice coefficient for GM, WM, CSF, respectively)	
Park et al. [51]	Brain MRI	Segmentation for brain metastases	3D BB + 3D GRE* 3D BB 3D GRE	93.1* 92.6 76.8 (Sensitivity)	
*The detection ne inversion recovery = peripheral tissue blood, GRE = grad	tworks with the , PSNR = peak s e, AP-DCE = all p ient echo	e best performance for ea signal-to-noise ratio, SSII phases of DCE-MRI, GM =	ch test dataset. MRI = magnetic resonance imaging, S M = structural similarity index, DWI = diffusion-weight gray matter, WM = white matter, CSF = cerebrospinal f	SWI = susceptibility-weighted imaging ted image, DCE = dynamic contrast-en fluid, T1-IR = T1 inversion recovery, 3	g, FLAIR = fluid-attenuated hanced, OA = over-appearance, PT 8D = three dimensional, BB = black

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Fig. 5. Example of intersection over union (IoU) degradation with different boundary box sizes for small-object detection. The ground truth and predicted results are represented by blue and green boundary boxes, respectively. The IoU for the small bounding boxes **(A)** and bounding boxes that include a sufficient area around the lesion **(C)** was estimated to be 0.605, and 0.614, respectively. For the small bounding boxes **(B)** and bounding boxes that include a sufficient area around the lesion **(D)**, the predicted boundary box was moved upward by two pixels and the IoU was estimated to be 0.485, and 0.511, respectively.

or cancers less than 3 mm in size, the size of the label box is significantly small. The intersection over union (IoU) [55] in the small label box can be easily decreased, even with a difference of 1–2 pixels between the ground truth and prediction results (Fig. 5). In Figure 5B, the IoU decreased to 0.485 owing to the upward movement of the prediction box by two pixels. If an IoU of 0.5 is used as the threshold for a true positive, it will not be evaluated as a true positive even if the prediction box includes the lesion. Therefore, when the labeling process is performed for small objects, the stability of the IoU can be improved by labeling a large area that includes a sufficient area around the lesion. In the study of [47], to detect cerebral microbleeds based on a deep learning approach, a bounding box with a size of 20 x 20 was applied for the labeling process, which included a sufficient area around the cerebral microbleeds.

Network Training

Object Detection Networks

Object detection architectures have been continuously developed, and several review papers have described their progress from early models to state-of-the-art technologies [56-58]. Therefore, this article does not provide a detailed description of each model. Instead, this article briefly describes the classification of object detection architectures into 2-stage and 1-stage detectors, and a brief description of several object detection models is presented. Table 4 summarizes the performance of some well-known object detection architectures [2,59-69].

The 2-stage detector performs localization and classification separately, whereas the 1-stage detector performs them simultaneously. Generally, a 2-stage detector

is recognized for achieving higher accuracy but lower speed. The R-CNN family is a representative 2-stage detector. The R-CNN [59] was the first model to apply a CNN to object detection and consists of region proposal (selective search), feature vector acquisition using a CNN, class classification using a support vector machine, and bounding box regression. However, R-CNN has the disadvantage of long training time owing to the multiple stages of learning. To improve this, a fast R-CNN [2] with a region of interest pooling and faster R-CNN [62] with a region proposal network were developed. Although not in the R-CNN family, the region-based fully convolutional network (R-FCN) model [65], which performs position-sensitive pooling using position-sensitive score maps, showed similar performance to the faster R-CNN but was 2.5 to 20 times faster. Moreover, the feature pyramid network (FPN) [64], which employs a method for recognizing target objects of various sizes, and mask R-CNN [63], which adds a mask branch to enable instance segmentation in the bounding box, have been introduced.

For real-time screening, a 1-stage detector, which has the advantage of high speed, is appropriate. It is known that the 1-stage detector shows lower performance than the 2-stage detector. However, owing to recent developments in the 1-stage detector, its accuracy has become similar to that of the 2-stage detector. The you only look once (YOLO) family is a representative 1-stage detector. YOLO [66], which is the first introduced model, redefines localization and classification, which are separately performed in a 2-stage detector, as a single-regression problem. Consequently, a single neural network predicts the bounding box and class probability using a single process. However, the YOLO model exhibits a lower mean average precision (mAP) value with



Proposed Model	Region Proposal	Trained Dataset	Training Time, h	Backbone	Test Dataset	mAP, %	Run-time, s
2-stage detector							
R-CNN [59]	Selective search	ILSVRC2012 + ILSVRC2013	13	-	ILSVRC2013	31.4	60 (CPU)
		ILSVRC2012 + VOC 2012		-	VOC 2010	53.7	
				-	VOC 2007	58.5	
				OxfordNet	VOC 2007	66.0	
Fast R-CNN [2]	Selective search	VOC 2007 + 2012	9.5	VGG16	VOC 2012	70.0	0.3
				VGG16	VOC 2010	68.8	
				VGG16	VOC 2007	68.4	
Faster R-CNN [62]	RPN	VOC 2007 + 2012 + COCO	-	VGG16	VOC 2012	75.9	0.2
				VGG16	VOC 2007	78.8	
				VGG16	C0C0	42.7	
R-FCN [65]*	RPN	VOC 2007 + 2012 + COCO	-	ResNet101	VOC 2012	82.0	0.42
				ResNet101	VOC 2007*	83.6*	
				ResNet101	C0C0	53.2	1
FPN [64]	RPN	0000	8 (8 GPUs)	ResNet101	C0C0	57.1	0.148
Mask R-CNN [63]	RPN	0000	44 (8 GPUs)	ResNeXt101FPN	C0C0	60.0	0.2
1-stage detector							
YOLO [66]	-	VOC 2007 + 2012	-	-	VOC 2012	57.9	0.02
				-	VOC 2007	63.4	
YOLO v2 [67]	-	ImageNet	-	Darknet19	VOC 2012	73.4	0.025
				Darknet19	VOC 2007	78.6	
				Darknet19	C0C0	44.0	
YOLO v3 [68]	-	-		Darknet53	C0C0	57.9	0.05
YOLO v4 [69]	-	0000	-	CSPDarknet53	C0C0	65.7	0.03
SSD [61]*	-	VOC 2007 + 2012 + COCO	-	VGG16	VOC 2012*	82.2*	
				VGG16	VOC 2007	83.2	0.045
				VGG16	C0C0	48.5	
RetinaNet [60]	-	0000	10-35	ResNeXt101FPN	C0C0*	61.1*	0.198

Table 4. Well-known Object Detection Networks and Their Performances in the Literature

*The detection networks with the best performance for each test dataset. mAP = mean average precision, R-CNN = region-based convolutional neural network, CPU = central processing unit, RPN = region proposal network, R-FCN = region-based fully convolutional network, FPN = feature pyramid network, GPU = graphic processor unit, YOLO = you only look once, SSD = single-shot detector

missing small objects. To overcome the disadvantages of the YOLO model, an SSD consisting of a multiscale feature layer and default box was proposed [61]. Subsequently, several improvements were introduced: YOLO v2 [67] improved performance by applying batch normalization and using an anchor box; RetinaNet [60] used focal loss to solve the class imbalance problem caused by the difference in the number of positive/negative samples used during model training; YOLO v3 [68] improved performance by using DarkNet53 as the backbone architecture and three feature maps; and YOLO v4 [69] combined various methods that affect performance.

If there is no time constraint, it is appropriate to select 2–3 recent and well-known models supported in the relevant development environment, compare them, and select the model with satisfactory performance. In addition, the proposed object detection network can be modified by applying models such as VGG-19 [70], ResNet-50 [71], or Inception v3 [72] as the backbone.

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Deep Learning Workflow Based on the Diagnostic Process Performed by Radiologists

Radiologists are empirically trained in anatomy, pathology, imaging techniques, and disease patterns, and they make decisions based on their own experiences and criteria [73]. However, a deep-learning network is not trained with any criteria with anatomical and pathological bases, and the weights of each node are adjusted by training on a specific dataset. Consequently, a deeplearning network cannot provide any criteria for its decision, and even if a deep-learning network presents a correct decision, it may not be accepted if the deep learning model does not provide any decision criteria. Therefore, rather than simply developing a deep learning network to detect disease, one can take a step towards developing a more reliable algorithm by understanding each process of disease diagnosis performed by a radiologist and applying it in similar manner to the deep learning workflow. In this way, the deep learning model can present the results of each step to the radiologists, helping them to better understand and trust the model's results.

Previous studies have proposed deep-learning workflows that mimic the diagnostic processes of radiologists for maxillary sinusitis assessment and mammography phantom image evaluation. For the assessment of maxillary sinusitis [19], the diagnostic processes of a radiologist include finding the facial region, adjusting the window level, increasing the contrast difference, diagnosing the lesion, and generating a clinical report. These processes were imitated and applied to the deep learning workflow in the form of preprocessing, facial patch detection, facial region extraction, image intensity normalization, maxillary sinusitis detection, and detection result generation, which highlights the image with a bounding box and a report regarding the original image space. Figure 6 shows a diagram that compares the process of maxillary sinusitis detection by radiologists and that of the deep learning model. For a mammography phantom image [42], the evaluation processes of a radiologist include finding the phantom region, adjusting the window level and width, evaluating each phantom object, summing phantom scores for each group according to the guidelines of the American College of Radiology digital mammography quality control, and generating reports. These processes were applied to the deep learning workflow in the form of phantom region detection, image intensity normalization, phantom object detection that yields location information as a bounding box with its group and score, summation of each phantom score for each group, and generating reports.

Imaging Data Argumentation

Data collection in the medical field is often limited, and data augmentation is performed to compensate for the insufficient amount of data in the training dataset. Data augmentation may be applied selectively or randomly and includes image processing, such as flipping, rotating, translating, and scaling the image size (magnification or reduction). The labels in the augmented image are identical to those in the original image but with slightly different features, thereby allowing the deep learning model to learn a wider variety of patterns, which can improve its performance. Yadav et al. [74] investigated the effect of data augmentation to distinguish pneumonia images from normal images in a chest X-ray dataset. They set two different augmentation models using different augmentation parameters, and the model that included the augmentation parameters of rotation range, shear range, zoom range, horizontal flip, and vertical



Fig. 6. Diagram of the diagnostic process for maxillary sinusitis performed by a radiologist and deep learning workflow that mimics the radiologist's diagnostic process. The upper row represents the diagram of the diagnostic process conducted by a radiologist, while the bottom row represents the diagram of the deep learning workflow designed to mimic each step of the radiologist's diagnostic process.

flip showed better results. Liu et al. [75] trained a deep learning network to detect cerebral microbleeds, and their data augmentation comprised 3D rotation, translation, and random left-to-right flipping to avoid overfitting. In a study evaluating the diagnostic performance of deep learning networks on panoramic radiographs, Yang et al. [76] augmented their training set by applying horizontal and vertical flipping, translation, and scaling.

Hyperparameter Optimization

A hyperparameter [77] that affects the performance of a deep learning algorithm is not the main variable optimized through the training process but is instead a variable that humans set as a priori knowledge before training the network. Hyperparameters include the activation function. batch size, dropout rate, number of dense nodes, input image size, epochs, initial learning rate, and a factor for L2 regularization [19,78]. Manual search [79], grid search [80], random search [81], and Bayesian optimization are known hyperparameter optimization methods. The Bayesian optimization method [82,83] uses prior knowledge to generate a statistical model based on experimental results, and it effectively determines the next search direction for the optimal hyperparameters by evaluating the objective function [84]. It has the advantage of efficiently finding the optimal hyperparameters in a shorter time than random or grid searches [85]. In our previous study [19], to enhance the performance of the maxillary sinusitis detector, Bayesian hyperparameter optimization was applied using the following parameters: input image size, number of anchor boxes, maximum epochs, initial learning rate, and a factor for L2 regularization. Bayesian hyperparameter optimization usually attempts to find values of hyperparameters that minimize an objective function and to find hyperparameters that increase the accuracy during the Bayesian optimization process. Ait Amou et al. [78] applied Bayesian optimization to obtain optimal hyperparameters for the complete training of their model to distinguish brain tumors. The activation function, batch size, dropout rate, number of dense nodes, and gradient descent optimization function were selected for Bayesian hyperparameter optimization, and their accuracy was evaluated as an objective function.

Performance Evaluation

Quantitative Performance Metric

For the object detection task, IoU, precision, recall,

average precision (AP), and mAP are mainly used to evaluate model performance. IoU is a metric that evaluates how much the predicted boxes overlap with the ground-truth bounding boxes and can be represented by Eq. (1) The IoU is used as a criterion to determine true and false positives and is the most popular evaluation metric used for object detection [55]. In general image domains, such as the PASCAL VOC [86] and MS COCO benchmark challenges [87], the performance of object detection models is commonly evaluated using a fixed IoU threshold of 0.5 [88] or multiple thresholds [62,69]. However, in the medical imaging domain, a fixed IoU threshold of 0.5 [89] or lower, such as 0.2 [17], may be used, depending on the specific study. However, it is also important to ensure that the lesions are included in the prediction box of the deep learning model. Precision (Eq. (2)) indicates the proportion of true positives among the total number of objects predicted by deep learning, and recall (Eq. (3)) indicates the proportion of true positives among all ground truths. The area under the curve in the precision-recall graph is calculated and expressed as an AP to quantitatively evaluate the model's performance [90]. The mAP is the mean of the AP values of each target class [91], and it is the same as the AP when only one target class exists. The false positive rate is calculated by dividing the total number of false positives by the number of slices or participants.

Grou	Ground truth \cap Predicted boxes				
100 = Grou	Ground truth \cup Predicted boxes				
Precision =	True positives	Fa (2)			
	Whole predictions by model	Ld. (L)			
Recall (Sens	sitivity) = True positives	Fa (3)			
needati (Sens	-9.(3)				

Ground truth

In a clinical setting, the developed computer-aided diagnostic algorithm is often evaluated using sensitivity, specificity, and area under the receiver operating characteristic (AUROC), which are evaluation metrics for distinguishing between normal and abnormal individuals. However, from Eq. (2) and (3), the precision and recall for the performance evaluation of the object detection model are the metrics evaluated in the target object unit.

To evaluate the performance for distinguishing normal and abnormal using the object detection model, secondary processing steps, such as considering predictions as normal or abnormal according to the presence or absence of the prediction result, should be performed. The researcher should then evaluate the accuracy, specificity, AUROC, etc.

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Evaluation of Deep Learning Models

In a mammography phantom image evaluation study [42], the inter-rater correlation coefficient for the total group score of the deep learning model and radiologist was 0.54–0.62, which is in the poor-to-acceptable range. However, as a result of evaluating each of the 16 phantom objects, the agreement between the deep learning model and ground truth was low only at the ambiguous point, and a similar pattern was observed in the results from humans. In previous studies on brain metastasis detection [17], although the overall detection sensitivity was 81%, the sensitivity for the small metastasis group (< 3 mm) was only 15%. By not only evaluating the integrated result but also investigating the deep learning performance for subclasses, researchers can identify deep learning that performs well.

CONCLUSION

In this study, we have addressed the potential challenges and important considerations that arise at each step of deep learning research when employing object detection methods. Although recent studies that have applied deep learning have shown remarkable performance, they have not always guaranteed the best results. Researchers can more efficiently perform deep learning research by identifying issues that may pose problems in each step of the research, thereby reducing trial-and-error.

Supplement

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Conflicts of Interest

The authors have no potential conflicts of interest to disclose.

Author Contributions

Conceptualization: Kyung Mi Lee. Formal analysis: Jang-Hoon Oh. Funding acquisition: Kyung Mi Lee, Hyug-Gi Kim. Methodology: Hyug-Gi Kim, Jang-Hoon Oh. Project administration: Kyung Mi Lee. Supervision: Hyug-Gi Kim, Kyung Mi Lee. Validation: Hyug-Gi Kim. Visualization: Jang-Hoon Oh. Writing original draft: Jang-Hoon Oh. Writing—review & editing: Kyung Mi Lee.

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