

Making Data Big for a Deep-learning Analysis: Aggregation of Public COVID-19 Datasets of Lung Computed Tomography Scans

Francesca Lizzi^{1,2}, Francesca Brero^{3,4}, Raffaella Fiamma Cabini^{3,5}, Maria Evelina Fantacci^{2,6}, Stefano Piffer^{7,8}, Ian Postuma³, Lisa Rinaldi^{3,4} and Alessandra Retico²

¹*Scuola Normale Superiore, Pisa, Italy*

²*National Institute for Nuclear Physics (INFN), Pisa Division, Pisa, Italy*

³*INFN, Pavia Division, Pavia, Italy*

⁴*Department of Physics, University of Pavia, Pavia, Italy*

⁵*Department of Mathematics, University of Pavia, Pavia, Italy*

⁶*Department of Physics, University of Pisa, Pisa, Italy*

⁷*Department of Biomedical Experimental Clinical Science "M. Serio", University of Florence, Florence, Italy*

⁸*INFN, Florence Division, Florence, Italy*

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Abstract: Lung Computed Tomography (CT) is an imaging technique useful to assess the severity of COVID-19 infection in symptomatic patients and to monitor its evolution over time. Lung CT can be analysed with the support of deep learning methods for both aforementioned tasks. We have developed a U-net based algorithm to segment the COVID-19 lesions. Unfortunately, public datasets populated with a huge amount of labelled CT scans of patients affected by COVID-19 are not available. In this work, we first review all the currently available public datasets of COVID-19 CT scans, presenting an extensive description of their characteristics. Then, we describe the design of the U-net we developed for the automated identification of COVID-19 lung lesions. Finally, we discuss the results obtained by using the different publicly available datasets. In particular, we trained the U-net on the dataset made available within the COVID-19 Lung CT Lesion Segmentation Challenge 2020, and we tested it on data from the MosMed and the COVID-19-CT-Seg datasets to explore the transferability of the model and to assess whether the image annotation process affects the detection performances. We evaluated the performance of the system in lesion segmentation in terms of the Dice index, which measures the overlap between the ground truth and the predicted masks. The proposed U-net segmentation model reaches a Dice index equal to 0.67, 0.42 and 0.58 on the independent validation sets of the COVID-19 Lung CT Lesion Segmentation Challenge 2020, on the MosMed and on the COVID-19-CT-Seg datasets, respectively. This work focusing on lesion segmentation constitutes a preliminary work for a more accurate analysis of COVID-19 lesions, based for example on the extraction and analysis of radiomic features.

1 INTRODUCTION

Lung Computed Tomography (CT) is a very sensitive medical imaging technique to detect lung lesions due to COVID-19. It can be used for the diagnosis, prognosis and for monitoring the disease evolution over time. Despite the use of CT for diagnosis is not recommended by the World Health Organization (World Health Organization, 2020), lung CT analysis can be very informative regarding the severity of the disease and its time evolution (Fang et al., 2021). The use of CT in clinical practice for COVID-19 diagnosis in symptomatic patients has been explored. Since

the unexpected outbreak of the pandemic, physicians tried to use CT imaging of the chest to diagnose COVID-19 disease. The first publication describing in details radiological findings of CT was published in January, the 24 of 2020 (Huang et al., 2020) and it describes the radiological findings of the majority of COVID-19 hospitalized patients of this study, such as bilateral multiple lobular and subsegmental areas of consolidation and bilateral ground-glass opacity. Afterwards, several studies have been published to describe the radiological findings of COVID-19 chest CT (Carotti et al., 2020). A summary of all possible findings and their incidence is reported in Table 1.

Table 1: Summary of COVID-19 chest CT findings and their incidence on the population. The normal chest CT findings are also associated to symptomatology (Huang et al., 2020).

Findings	Incidence
Normal chest CT findings	10.6% (95% CI: 7.6%, 13.7%)
Ground-Glass opacity, Lower lobe involvement, Bilateral abnormalities, Vascular enlargement, Posterior predilection,	High incidence (More than 70%)
Consolidation, linear opacity, septal thickening and/or reticulation, crazy-paving pattern, air bronchogram, pleural thickening, halo sign, bronchiectasis, nodules, bronchial wall thickening, reversed halo sign	Intermediate incidence (between 70% and 10%)
Pleural effusion, lymphadenopathy, tree-in-bud sign, central lesion distribution, pericardial effusion, cavitating lung lesions	Low incidence (less than 10%)

We underline that the dataset used by (Huang et al., 2020) contains a very limited number of CT scans (41 patients) and it is private. Most of the chest CT findings cannot be related exclusively to COVID-19 because they are nonspecific signs of disease and they are strongly related to the stage of the disease. This means that there are other forms of pneumonia that may have the same signs such as SARS-CoV-1 and MERS-CoV. For this reason, the World Health Organization (WHO) defined as “confirmed case” the patient that have been tested positive for COVID-19 RT-PCR, irrespective of clinical signs and symptoms (World Health Organization, 2020). Furthermore, it is necessary to differentiate the COVID-19 infections not only from other viral pneumonia but also from bacterial pneumonia, such as mycoplasma pneumonia (Ishiguro et al., 2019). The use of chest CT to diagnose COVID-19 is, hence, under discussion since it implies the use of ionizing radiation (Scientiae et al., 2020) while its ability to monitor the progression of the disease seems to be a promising way to use lung CTs (Adams et al., 2020). Artificial Intelligence (AI) is a powerful instrument that allows to analyse a huge quantity of data, such as CT scans and, hence, it can be used to monitor and study COVID-19 CT signs (Gülbay et al., 2021). Unfortunately,

AI implementations require a great amount of data, which may be not easily available. This is especially true when deep-learning methods are used. Since the beginning of the pandemic, some lung CT scans of COVID-19 patients have been released by different institutions following different guidelines for both image acquisition and annotation (ground truth). In this work, all the public lung COVID-19 CT datasets, to the best of our knowledge, suitable for training AI-based systems are reviewed. In this work, we present an extensive description of the currently publicly available datasets, which present different characteristics, and discuss the segmentation results obtained by using them. In particular, we trained a U-net on the dataset released within the COVID-19 Lung CT Lesion Segmentation Challenge 2020 (An et al., 2020) and tested it on data from MosMed (Morozov et al., 2020) and COVID-19-CT-Seg datasets (Ma et al., 2020). Finally, the limits and the advantages of aggregating this kind of data are discussed.

2 AI AND MEDICAL IMAGE DATASET ISSUES

AI has been used to analyse and process CT to diagnose COVID-19, to segment lesions inside the lungs and, also, in longitudinal studies to track the evolution of the disease (Ma et al., 2020). AI based methods, especially deep-learning ones, need a huge amount of labelled data that are not easy to collect and share. As already described in the introduction, some studies use private datasets which do not allow a fair comparison with other AI based systems. Furthermore, the characteristics of CT images depend on the scanner, on the acquisition and the reconstruction protocols and on other information which may not be available. This can be due also to the anonymization process needed to preserve subjects’ privacy or to the use of image format different from DICOM (Standard DICOM, 2021), such as the NIfTI format. DICOM is the most used image format for medical images and it contains several meta-data in its header. The DICOM header stores many information, some of which is Protected Health Information (PHI) or private keys that are inserted and encoded by the manufacturer and may contain PHI as well. On the other hand, some meta-data, such as anode characteristics or X-ray parameters, do not contain PHI and they can be useful in analysing images. For all these reasons, anonymizing a DICOM file is not a trivial problem and dataset may include images in a different format such as NIfTI (Moore et al., 2015). Deep learning based methods often require the association with a la-

bel depending on the task we want to solve. Many approaches are based on supervised learning and, hence, image annotation plays a crucial role. Usually, medical image labels are given by one or more radiologists with experience in the specific field, and image annotation is a very time-consuming task. This is the reason why there is a general lack of public labelled datasets of medical images. In order to save time, it may happen that the labelling is made with the support of an automatic tool and then labels are adjusted manually by one or more physicians.

3 LUNG CT DATASETS

In this section, the currently available datasets of lung CT and their annotation process are reported. The dataset are: COVID-19 Lung CT Lesion Segmentation Challenge 2020 Dataset, MosMed Dataset, COVID-19-CT-Seg Dataset and TCIA-COVID-19-AR.

3.1 COVID-19 Lung CT Lesion Segmentation Challenge 2020 Dataset

The COVID-19 Lung CT Lesion Segmentation Challenge 2020 (Challenge dataset) dataset is a public dataset made by 199 unenhanced chest CT with positive RT-PCR for SARS-CoV-2 patients (An et al., 2020), published as training set in the occasion of the COVID GrandChallenge (<https://covid-segmentation.grand-challenge.org/>). Each CT is annotated voxel-wise and indicates all the COVID-19 lesions in a unique mask. Data has been provided by The Multi-national NIH Consortium for CT AI in COVID-19 via the NCI TCIA public website in NIFTI format. Annotations have been made using a COVID-19 segmentation model provided by NVIDIA that takes a full CT chest volume and produces pixel wise segmentation masks of COVID-19 lesions. These segmentation masks have been adjusted manually by a board of certified radiologists in order to give 3D consistency to the lesion masks. The annotations of the training set have been published in the context of the challenge while the system performance has been evaluated by challenge organizers on an independent validation set of 50 CT scans, for which the lesion annotations were not publicly released. A third set, an independent test set consisting of 46 CT scans, was used to the define the final ranking among the participants, and, also in this case, the lesion segmentation annotations were not publicly released.

3.2 MosMed Dataset

MosMed (Morozov et al., 2020) is a dataset of COVID-19 Chest CT scans collected by the Research and Practical Clinical Center for Diagnostics and Telemedicine Technologies of the Moscow Health Care Department. It includes 1110 CT studies taken from 1110 patients and it is provided with a labelling that consists of 5 classes, based on the percentage of involved lung parenchyma. A small subset of class CT-1 cases (50 patients) has been annotated by expert radiologists with the support of Med-Seg software (2020 Artificial Intelligence AS). The image annotations consist of binary masks in which white voxels represent both Ground-Glass opacities and consolidation. Both CT scans and annotations were provided in NIFTI format. During the DICOM-to-NIFTI conversion only one every 10th image was preserved (MosMed, 2020).

3.3 COVID-19-CT-Seg Dataset

The COVID-19-CT-Seg dataset is a collection of CT scans made available by the Coronacases Initiative and Radiopaedia (Ma et al., 2020) and contains 20 CT scans of patients resulted positive for RT-PCR COVID-19 infection. It is a public dataset which contains annotations related to both lung and infection localization. The ground truth has been made in three steps: first, junior radiologists (1-5 years of experience) delineated the annotations of lungs and infections, then two radiologists (5-10 years of experience) refined the labels and finally the annotations were verified and optimized by a senior radiologist (more than 10 years of experience in chest radiology). The annotations have been produced with ITK-SNAP software. Ten cases of this dataset were provided in 8-bit depth which are not commonly used in clinical practice.

3.4 TCIA-COVID-19-AR

The TCIA-COVID-19-AR (Desai et al., 2020) is a dataset of COVID-19 cases taken from a rural population, which is often underrepresented in public datasets. It contains 24 CT scans of patients with both lung lesions due to COVID-19 and control cases. Each patient is described by a set of clinical data correlates that includes key radiology findings. Moreover, for each patient the information about Intensive Care Unit (ICU) admission is included while annotations on images are not included in this dataset.

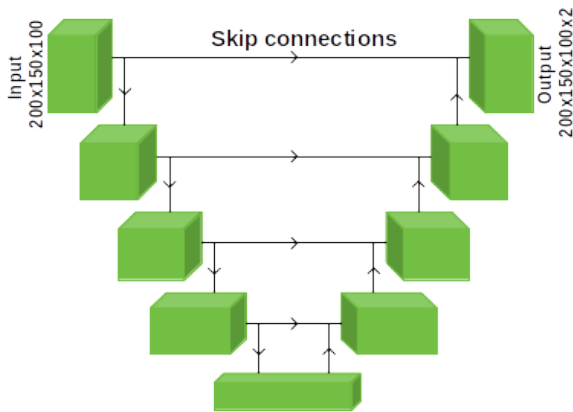


Figure 1: U-net summary: the U-shaped neural network is made of 5 levels of depth. In the left path (compression), the input is processed through convolutions, activation layers (ReLU) and instance normalization layers, while in the right one (decompression), in addition to those already mentioned, 3D Transpose Convolution (de-convolution) layers are also introduced. Each block (green) is made of 3 convolutional layers.

4 COVID-19 LESION SEGMENTATION

We developed an automatic system which can segment COVID-19 lesions based on a U-net (Ronneberger et al., 2015) in the framework of the COVID-19 Lung CT Lesion Segmentation Challenge 2020 (GrandChallenge, 2020). First, a bounding box which contains the lungs has been built for each CT scan to reduce as much as possible the background from the images. An in-house lung segmentation algorithm based on active contours was developed for this purpose and implemented in *matlab* (The MathWorks, Inc.). This algorithm, which accurately segments the lung parenchyma in absence of lesions, has very limited performance on CT scans of subjects with COVID-19 lesions. The CT images have been cropped to the bounding boxes, resized to a matrix of $200 \times 150 \times 100$ voxels and a CT windowing in $[-1000, 300]$ range of Hounsfield Units has been applied on them to enhance the COVID-19 lesions. A schematic representation of the used U-net is reported in Figure 1.

We trained the network on the Challenge training dataset of 199 CT scans, using a weighted cross-entropy as loss function, and we tested it on the Challenge validation set (independent from the training set). In order to have a sufficient number of samples, we applied data augmentation with rotations, zooming and elastic transformation to the training set. We tested the network also on the 50 annotated cases of MosMed and on the 10 annotated cases of the

COVID-19-CT-Seg-Dataset. The MosMed dataset contains images and labels taken in a very different way with respect to those of the Challenge dataset. The COVID-19-CT-Seg dataset has been built in a more similar way to the Challenge one for both data characteristics, such as slice thickness, and labelling process. We evaluated the segmentation performance of the trained network model in terms of Dice index (Equation 1) defined as:

$$\text{Dice}_{metric} = \frac{2 \cdot |M_{true} \cap M_{predict}|}{|M_{true}| + |M_{pred}|} \quad (1)$$

where M_{true} is the ground truth mask and M_{pred} is the predicted one.

We participated in the challenge, obtaining a Dice index equal to 0.67 on the challenge validation set (GrandChallenge, 2020). Then, we computed the segmentation performance of the trained model on the MosMed dataset obtaining a Dice of 0.42, and on the COVID-19-CT-Seg-Dataset, obtaining a Dice of 0.58.

We show in Figure 2 a visual comparison between the reference COVID-19 lesion masks and the ones predicted by the trained U-net for a representative CT scan of the MosMed and of the COVID-19-CT-Seg dataset.

5 DISCUSSION AND CONCLUSIONS

We obtained good results in terms of the Dice index as regards the segmentation of the lung lesions related to COVID-19 infection on the Challenge dataset compared to literature (Ma et al., 2020). The results obtained on the other two datasets are not good as the first one. We underline that on the dataset more similar to the Challenge one, the COVID-19-CT-Seg dataset, we obtained better results compared to MosMed. As expected, we conclude that aggregating data from different sources can be difficult if labelling has been performed using different guidelines. In fact, medical images have many parameters to be considered, such as the resolution of pixels and the size of the Field Of View (FOV). These parameters can be studied in order to attempt a standardization of images from different datasets, by contrast, different annotation styles can not be easily standardized. Since CT image characteristics can be variable, deep learning is a useful method to analyse them and their aggregation. Moreover, U-nets allows a quantification of the volumes of both COVID-19 lesions and lungs. On the other hand, the use of deep learning based methods requires a huge amount of homogeneous or harmonized data both to carry out an optimal training

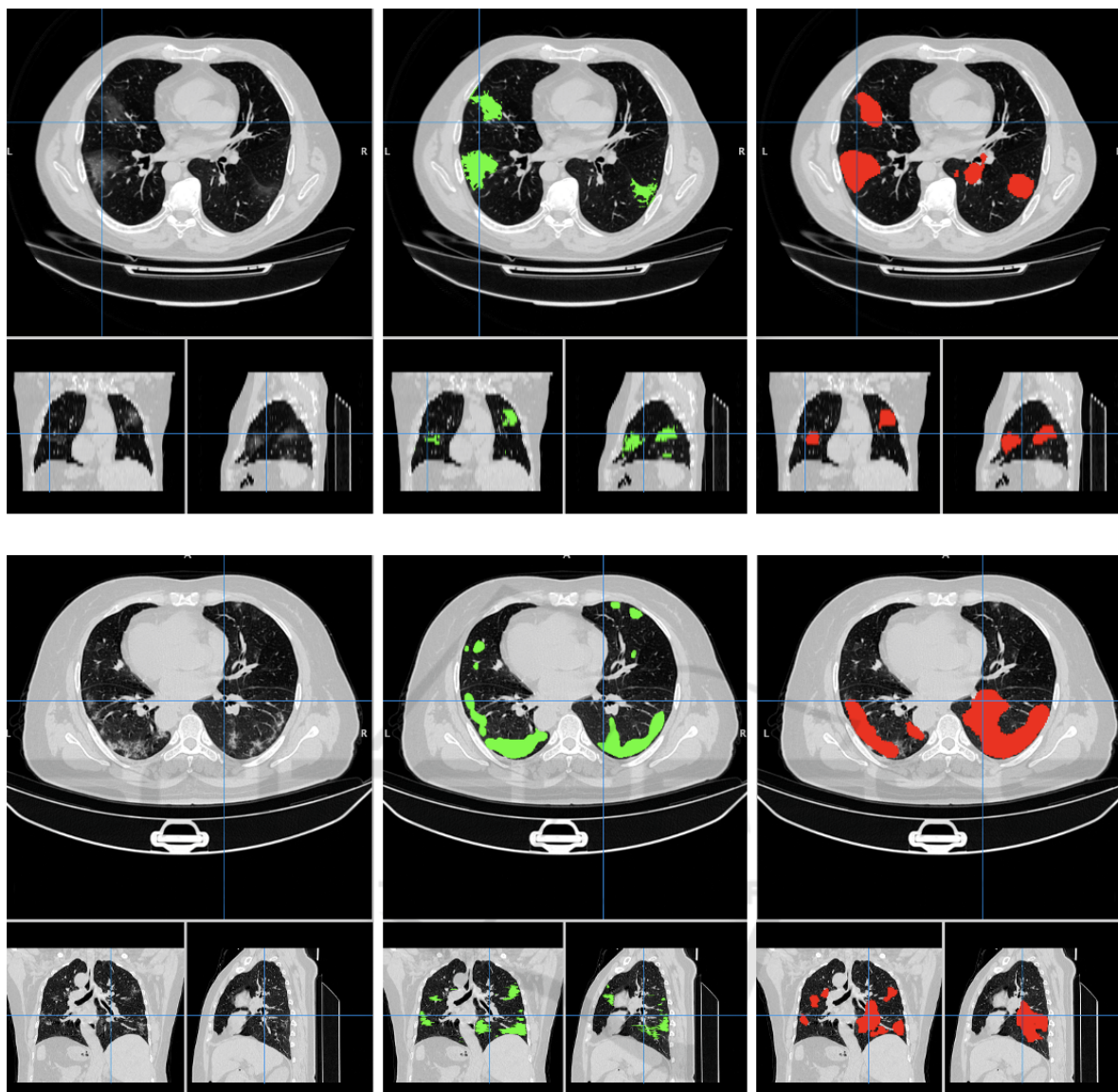


Figure 2: Visual comparison between the reference COVID-19 lesion masks (green) and the ones predicted (red) by the trained U-net for a representative CT scan of the MosMed (first row, study-0255.nii) and of the COVID-19-CT-Seg (second row, coronacases-001.nii) datasets. The original CT scans are shown on the left as a reference.

process and to implement a fair representation of the population to be studied.

This preliminary study has been useful to understand which parameters should be considered as the most critical ones in training a neural network model. Lesion labeling and data selection criteria are crucial for this kind of segmentation problems because of the lack of largely populated public datasets, impacting in a relevant way on the performances. In conclusion, we reviewed all the public available datasets (at the best of our knowledge in April 2021), i.e. COVID-19 Lung CT Lesion Segmentation Challenge 2020, MosMed, COVID-19-CT-Seg Dataset

and TCIA-COVID-19-AR. We used the Challenge data to train and evaluate a U-net for COVID-19 lung lesion segmentation, and we carried out an independent test of the MosMed and the COVID-19-CT-Seg datasets, obtaining good performances, as compared to other results available in literature (Ma et al., 2020). We are going to improve our system by adding a module for lung segmentation which could help in quantifying the percentage of lung tissue affected by COVID-19 lesions. We also plan to let radiologists evaluate the application of this algorithm on a part of public CT datasets without labelling. Furthermore, segmentation of COVID-19 lesions is a starting point

for an accurate radiomic analysis for the prediction, based on radiological signs, of the clinical outcome of patients affected by COVID-19 pneumonia.

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REFERENCES

- Adams, H. J., Kwee, T. C., Yakar, D., Hope, M. D., and Kwee, R. M. (2020). Chest CT Imaging Signature of Coronavirus Disease 2019 Infection: In Pursuit of the Scientific Evidence. *Chest*, 158(5):1885–1895.
- An, P., Xu, S., Harmon, S. A., Turkbey, E. B., Sanford, T. H., Amalou, A., Kassin, M., Varble, N., Blain, M., Anderson, V., Patella, F., Carrafiello, G., Turkbey, B. T., and Wood, B. J. (2020). CT Images in COVID-19.
- Carotti, M., Salaffi, F., Sarzi-Puttini, P., Agostini, A., Borgheresi, A., Minorati, D., Galli, M., Marotto, D., and Giovagnoni, A. (2020). Chest CT features of coronavirus disease 2019 (COVID-19) pneumonia: key points for radiologists. *Radiologia Medica*, 125(7):636–646.
- Desai, S., Baghal, A., Wongsurawat, T., Al-Shukri, S., Gates, K., Farmer, P., Rutherford, M., Blake, G., Nolan, T., Powell, T., Sexton, K., Bennett, W., and Prior, F. (2020). Data from Chest Imaging with Clinical and Genomic Correlates Representing a Rural COVID-19 Positive Population [Data set].
- Fang, X., Kruger, U., Homayounieh, F., Chao, H., Zhang, J., Digumarthy, S. R., Arru, C. D., Kalra, M. K., and Yan, P. (2021). Association of AI quantified COVID-19 chest CT and patient outcome. *International Journal of Computer Assisted Radiology and Surgery*.
- GrandChallenge (2020). COVID-19 Lung CT Lesion Segmentation Challenge - 2020, <https://covid-segmentation.grand-challenge.org/COVID-19-20/>.
- Gülbay, M., Özbay, B. O., Mendi, B. A. R., Baştuğ, A., and Bodur, H. (2021). A CT radiomics analysis of COVID-19-related ground-glass opacities and consolidation: Is it valuable in a differential diagnosis with other atypical pneumonias? *PloS one*, 16(3):e0246582.
- Huang, C., Wang, Y., Li, X., Ren, L., Zhao, J., Hu, Y., Zhang, L., Fan, G., Xu, J., Gu, X., Cheng, Z., Yu, T., Xia, J., Wei, Y., Wu, W., Xie, X., Yin, W., Li, H., Liu, M., Xiao, Y., Gao, H., Guo, L., Xie, J., Wang, G., Jiang, R., Gao, Z., Jin, Q., Wang, J., and Cao, B. (2020). Clinical features of patients infected with 2019 novel coronavirus in Wuhan, China. *The Lancet*, 395(10223):497–506.
- Ishiguro, T., Kobayashi, Y., Uozumi, R., Takata, N., Takaku, Y., Kagiya, N., Kanauchi, T., Shimizu, Y., and Takayanagi, N. (2019). Viral pneumonia requiring differentiation from acute and progressive diffuse interstitial lung diseases. *Internal Medicine*, 58(24):3509–3519.
- Ma, J., Wang, Y., An, X., Ge, C., Yu, Z., Chen, J., Zhu, Q., Dong, G., He, J., He, Z., Nie, Z., and Yang, X. (2020). Towards Efficient COVID-19 CT Annotation: A Benchmark for Lung and Infection Segmentation. pages 1–7.
- Moore, S. M., Maffitt, D. R., Smith, K. E., Kirby, J. S., Clark, K. W., Freymann, J. B., Vendt, B. A., Tarbox, L. R., and Prior, F. W. (2015). De-identification of medical images with retention of scientific research value. *Radiographics*, 35(3):727–735.
- Morozov, S. P., Andreychenko, A. E., Pavlov, N. A., Vladzimirskyy, A. V., Ledikhova, N. V., Gombolevskiy, V. A., Blokhin, I. A., Gelezhe, P. B., Gonchar, A. V., and Chernina, V. (2020). MosMedData: Chest CT Scans with COVID-19 Related Findings Dataset. *medRxiv*, page 2020.05.20.20100362.
- MosMed (2020). MosMed dataset website, <https://mosmed.ai/en/>.
- Ronneberger, O., Fischer, P., and Brox, T. (2015). U-net: Convolutional networks for biomedical image segmentation. *Lecture Notes in Computer Science (including subseries Lecture Notes in Artificial Intelligence and Lecture Notes in Bioinformatics)*, 9351:234–241.
- Scientiae, D. C.-S., Adams, H. J. A., Kwee, T. C., Hope, M. D., Kwee, R. M., Hja, A., Tc, K., and Yakar, D. (2020). Systematic Review and Meta- in the Diagnosis of Coronavirus. (December):1342–1350.
- Standard DICOM (2021). DICOM standard.
- World Health Organization (2020). WHO Interim guidance 20 March 2020 - Global Surveillance for COVID-19 disease caused by human infection with novel coronavirus (COVID-19). *Who*, (January):1–4.