Cohesive Multi-Modality Feature Learning and Fusion for COVID-19 Patient Severity Prediction

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Abstract—The outbreak of coronavirus disease (COVID-19) has been a nightmare to citizens, hospitals, healthcare practitioners, and the economy in 2020. The overwhelming number of confirmed cases and suspected cases put forward an unprecedented challenge to the hospital's capacity of management and medical resource distribution. To reduce the possibility of cross-infection and attend a patient according to his severity level, expertly diagnosis and sophisticated medical examinations are often required but hard to fulfil during a pandemic. To facilitate the assessment of a patient's severity, this paper proposes a multi-modality feature learning and fusion model for end-to-end covid patient severity prediction using the blood test supported electronic medical record (EMR) and chest computerized tomography (CT) scan images. To evaluate a patient's severity by the co-occurrence of salient clinical features, the High-order Factorization Network (HoFN) is proposed to learn the impact of a set of clinical features without tedious feature engineering. On the other hand, an attention-based deep convolutional neural network (CNN) using pre-trained parameters are used to process the lung CT images. Finally, to achieve cohesion of cross-modality representation, we design a loss function to shift deep features of both-modality into the same feature space which improves the model's performance and robustness when one modality is absent. Experimental results demonstrate that the proposed multi-modality feature learning and fusion model achieves high performance in an authentic scenario.

Index Terms—Multimodality, COVID-19 severity prediction, factorization methods, attention mechanism, convolutional neural network.

I. INTRODUCTION

THE highly contagious coronavirus 2019 (COVID-19), first detected in Wuhan City, Hunan Province, China by the end of 2019, has caused thousands of causalities and tremendous economic damage worldwide [1], [2]. Hospitals

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Fig. 1. The number of COVID-19 cases reported weekly by the World Health Organization (WHO) Region, and global deaths, as of 23 February 2021. [1].

and clinics are soon overrun by confirmed patients which brings considerable stress on scarce medical resources. Many lives are lost not because of insufficient nucleic acid amplification testing reagents but the underestimation of the severity of their infection. Fig. 1 and Fig. 2 show the current global distribution of COVID-19 cases. At the beginning of the pandemic, professional clinicians require assessment reports and medical data obtained by sophisticated and expensive medical tests to determine the severity of a patient. Such tests become more and more inaccessible in face of the increasing number of patients. As a result, many patients were unaware or under-treated before they turn into a fatal stage. Therefore, the use of inexpensive and accessible medical data of a confirmed COVID-19 patient for the evaluation of their severity level can be a game-changer in such devastating situations. Hundreds of lives can be saved if the hospital can grasp the severity of their condition and make arrangement to the limited medical resources accordingly.

Although much work has been proposed to classify the infected patients or to detect lesion areas based on visual data [3]–[5], few works are providing a viable solution for inpatient's severity estimation that promotes the efficiency of medical resources allocation and reduces the possibility of cross-infection by potential fatal cases. To some extent, the automatic and accurate classification of COVID-19 infected patients can alleviate the excessive demand for the limited medical testing resources and facilitate the following clinical decision and treatment. However, these classification methods do not explicitly estimate the severity of the inpatients. The severity of the inpatients has

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Fig. 2. Distribution of global cases of coronavirus COVID-19 on 23 February 2021 [21].

been regarded as key guidance for rational allocation of medical resources, efficient clinical decision, and effective quarantine measurement under the pandemic situation [6]–[8]. Actually, most patients are experiencing obvious symptoms of COVID-19 disease before their admission to the hospital. Thus, assigning them to the proper ward, and providing them with appropriate treatments according to their severity level to avoid further damage should be a more necessary measure for the patients' protection. In other words, a reliable estimation of a inpatient's severity level can save lives and relieves the burden of hospitals and clinics caused by the overwhelming patients, especially under the situation of a pandemic. In this paper, we mainly focus on the multi-modality method for inpatient's COVID-19 severity prediction since medical test outcomes and the chest computerized tomography (CT) images are the two important sources of data that the clinician will refer to when determining the severity level of a COVID-19 patient.

The appropriate fusion of available information from multi-modality data is still an important way to increase the prediction accuracy of the COVID-19 patient severity, even if some data missing issues (e.g. modality missing) may be encountered [9], [10]. The missing issue of some patient's medical test outcomes (i.e. loss of medical test outcomes or chest CT image) is more likely to be encountered in overwhelmed hospitals due to the limitation of medical resources to perform medical testings. Thus, at such a crucial moment in epidemic prevention and controlling, the availability and accessibility of inexpensive multi-modality data, along with well-performing automatic data analytic and fusion method has become a vital topic in patient's severity estimation. Multi-modality data fusion and classification aim to integrate the data of different distributions, sources, and types into a global space in which both inter-modality and cross-modality can be represented in a unified manner [11]. However, multi-modality data fusion methods based on CT images and medical data has not been widely studied.

To the best of our knowledge, the most related works are using image and text description for visual question answering (VQA) [12], depression classification based on video and text records of questionnaire responses [13], as well as visual-based click-through rate (CTR) with visual commercial posters [14]. In particular, for CTR prediction and multi-modality depression classification, visual data and textual data are processed using two independent networks or machine learning models before applying concatenation operation. The combined features are then fed into a multi-layer perceptron (MLP) to generate the CTR prediction [15], [16] or used to compute some confidence score for the depression diagnosis [17], [18]. For VQA, a typical framework is designed to exploit a deep convolutional neural network for images along with a deep long-short term memory network (LSTM) for questions before fusing visual and textual features using element-wise production or feature vector concatenation [19], [20]. Unfortunately, most of the above-mentioned methods can not be applied directly to our COVID-19 severity prediction scenario for two reasons. On one hand, training a high-performing deep convolutional neural network (CNN) network or a language LSTM requires a large-scale dataset while patient's comprehensive data in the medical field is often unavailable especially when a new disease emerges. On the other hand, their methods only operate on complete multi-modality data, leading to a constrained classification model. Furthermore, when building the classification model, most existing methods simply concatenate features from different modalities into a single feature vector without considering their underlying associations.

In this paper, we propose an end-to-end learning method to learn and fuse multi-modality features for COVID-19 inpatient severity prediction that simultaneously exploit the information from different modalities (i.e., patient's chest CT images and the patient's electronic medical record (EMR)). As discussed earlier, major obstacles of high-performing severity perdiction are 1) the insufficiency of the large-scale dataset with complete multi-modality data, 2) the lack of viable ways to handle data missing, and 3) the unavailability of offthe-shelf methods to leverage complex EMR data without excessive feature engineering workload. To address these problems, a novel factorization-machine-based network is proposed to explicitly consider both low-level and high-level feature interactions without tedious feature engineering. Unlike a virus that only attacks specific organs of the patient, diseases like COVID-19 causes lesions and dysfunctions in multiple organs and eventually lead to fatality. With possible complications not being fully explored, automatically weighing the impact of high-level feature interactions contribute not only to better generalization but also potential clinical discovery of the disease. In addition, the inter-modality similarity loss is designed for modality-shared information learning and robustness against missing modality situations. Compared with the problem of binary classification that identify COVID-19 infected lungs from the healthy ones, the problem of estimating a patient's severity level is more challenging because more fine-grained and more discriminative features are required to perform multiclass severity prediction. Compared with other methods using multimodality for cross-domain analysis, our method is a pioneering work to address the problem of fusing EMR data and visual data. The main contributions of this paper can be summarized as follows:

- We propose a high-order factorization network (HoFN) for COVID-19 patient severity prediction based on EMR data.
- We propose an attention-based convolutional network for COVID-19 CT image recognition.
- A novel multi-modality method is proposed to learn and fuse EMR data and visual data for patient's severity prediction.
- Without the need for data augmentation, our method achieves high accuracy on real-life patient cohort data.

The rest of the paper is organized as follows. Section II reviews some related methods for conventional medical data analysis and off-the-shelf methods to handle multi-modality data. Section III demonstrates the proposed network in details. Section IV presents the experimental settings, results, comparison and discussion. Section V concludes this paper.

II. RELATED WORK

A. Medical Data Analysis

Medical data are produced during the operation process of a hospital which includes examination, testing, monitoring, and treatment in medical intensive care units (ICU). Despite the increasing accessibility and efficiency of admission testing, medical data still requires thorough inspections from medical experts before knowing the situation of some patients at the moment. To assist medical practitioners who have involved in these works, many methods for medical data analysis are proposed. Most of them use machine learning (ML) techniques such as logistic regression and decision tree to predict the daily situation of patients inside the hospital [22]. Due to their applicability, statistical reliability, and interpretability, these machine learning techniques are further exploited to predict the occurrence of critical illness for COVID-19 patients. In [23], by employing the Least Absolute Shrinkage and Selection Operation (LASSO) regression, significant medical features are selected to build a logistic regression model. Coefficients of the logistic regression model are further utilized to construct a COVID-19 risk score for online critical illness prediction. Although this model is computationally simple and interpretable, it ignores some higher-order intersection of the patient's medical status which can be further utilized to diagnose a patient's severity. Generally, the major drawback of medical data analysis using conventional ML techniques is the missing of the sophisticated feature engineering processes and the lack of expressive power.

Compared with conventional ML methods, deep neural networks can yield outstanding results without the need for significant domain-specific knowledge [24]. For the ease of data analysis, EMR is usually converted into categorical and continuous values after preprocessing. Therefore, some methods in the field of CTR prediction can be utilized to analyze both continuous and categorical data. To learn both the co-occurrence of categorical features and the transitivity of features' correlation, Cheng *et al.* [25] proposes the Wide & Deep learning framework, a hybrid network structure to combine linear model and deep learning model. In their model, although the intersection of different categorical features is used as input to the wide component, expertly feature

engineering is still required to construct a large number of intersecting features from the input data manually. To extend this framework, Chen et al. [26] propose to use a more efficient locally connected network to replace the deep component of the Wide & Deep learning framework. What's more, his work demonstrates the selection of useful features during the feed-forward process is an important step towards efficiency and higher performance. To automatically capture the linear and pairwise interactions between features. Guo et al. [27] propose deep factorization machines (DeepFM). By modeling both high-order and low-order feature interactions, DeepFM gains performance improvement without requiring tedious feature engineering. However, DeepFM only considers feature interaction in the same abstraction level and it may not be applicable in our patient severity prediction problem, which involves features from heterogeneous modalities (i.e., CT image and EMR data). To deal with the drawbacks of previous networks we propose a deep learning method that can utilize cross-layer interactions.

B. Deep Convolutional Neural Networks for Medical Images Analysis

Medical image analysis is a critical technique to give a quick detection of illness areas and abnormal conditions in human body tissues. As one of the popular medical imaging technology, CT plays an important role in the COVID-19 diagnosis. Inspired by the success of deep learning in image classification [28], [29], deep learning techniques, such as CNN, have been introduced into the field of medical image analysis and have received great interest from the community [30]–[32].

To perform effective medical diagnosis and treatment for COVID-19 patients, several works have exploited CNN for the task of COVID-19-related medical CT image analysis [33]-[35]. One key challenging issue comes from the insufficiency of training data. CT images for inpatient are normally not kept storage for privacy concerns, or the lack of data management mechanism. Without sufficient training data, the CNN models would not be effective and will tend to overfit. To alleviate this issue, most works employ the pre-trained weights in their CNN models. For example, Wang et al. [33] train a modified inception network (M-inception) for the chest CT image, by incorporating pre-trained network parameters from the large scale image dataset (i.e., ImageNet [36]). With the help of pre-trained weights, their method can still achieve outstanding performance with insufficient CT images. On the other hand, Song et al. [34] designed the Details Relation Extraction neural network (DRE-Net) based on a pre-trained ResNet-50 network to extract highly relevant details in the CT images. Last but not least, Wang et al. [35] propose COVID-Net that possesses selective long-range connectivity to improve representational capacity. In his work, patients are assigned into four categories including normal, bacterial infection, non-COVID, and COVID-19 infection so as to find the optimal treatment method for each patient.

Following current practices, pre-trained weights are also applied in our designed CNN model in order to extract more fine-grained features for severity prediction. Unlike the above methods, our method further incorporates an attention module on top of the convolutional network to facilitate recognition performance. This is because the additional attention mechanism is able to help the baseline CNN model to focus more on the lesion areas that may exist in the lung lobes.

C. Multi-Modality Medical Data Analysis

When a single modality can not provide sufficient information during analysis, multi-modality approaches become effective to leverage different sources of information for better contextual knowledge [11]. For example, magnetic resonance imaging (MRI) conveys information on the spatial structure of a brain while positron emission tomography (PET) shows function-related information of the brain [32]. Observing the same region from multiple perspectives enables a model to enlarge the feature space for exploring the ground-truth distribution. In the field of medical data diagnosis, professional medical practitioners make diagnoses to patients by utilizing several sources of information. For example, observation (visible symptoms), CT or CXR results (visual data), and medical test outcomes (EMR data).

With the advancement of deep learning methods, data collected from different modalities can be well-handled by neural networks. In particular, Dolz et al. [37] propose the HyperDense-Net for multimodality image segmentation. By properly combining information from both CT images and MRI, the HyperDense-Net achieves better performance than single-modality-based methods. However, the input to their models are limited in visual data, and they ignore the modality missing issue. Nie et al. [38] and Chen et al. [39] follow a similar approach and propose to increase the model's performance using different types of visual data from the same region. Liang et al. [40] put forward a multimodal Deep Belief Network (DBN) to analyze cancer data using an unsupervised learning scheme. The DBN can capture both intra-modality and cross-modality correlations and identify cancer subtypes from genomic data and clinical data. However, without higher-level feature fusion, their method may lack generalization capability if the training dataset is small. In addition, the above works have not addressed the missing modality problem in multi-modality scenarios. In our work, both CT images and EMR data used to predict the COVID-19 patients' severity level. Especially, if a sample encounters a modality missing issue (e.g., unavailable CT images), other available modalities can be a compliment.

III. PROPOSED METHOD

Medical test outcomes and medical imaging are the two major sources of information for early COVID-19 diagnoses as well as the key to judging the severity of the infection to date. To construct a more accurate and robust severity prediction model for COVID-19 patients, both visual data and EMR data should be taken into consideration. As such, an end-to-end multi-modality feature learning and fusion model is proposed to cope with features from diverse modalities.

This section demonstrates the three components in the proposed multi-modality feature learning and fusion method. The main pipeline of our method is illustrated in Fig. 3, after obtaining lung CT images and EMRs from a COVID-19 patient retrospective study, several preprocessing steps are applied to CT images as well as EMR. Then, the multi-modality data are converted into standardized lung image and medical data consisting of categorical and numerical features. Next, we arrive at the multi-modality feature learning and fusion process handled by a two-branch architecture. For the branch which handles visual data from CT images, an attention-based convolutional network is used to learn interpretive and discriminative features for severity level prediction. For the other branch handling panel data from EMR, a novel deep learning network is proposed to exploit the interaction of both low-order and high-order features. The prediction results of each branch are aggregated to obtain the final results via addition operator.

A. Medical Data Preprocessing

1) Data Overview: To obtain standardized panel data consisting of both continuous values and categorical values from EMRs, three types of factors including 194 examination indicators were extracted from each patient's medical record. These indicators can be sorted into three categories. They are demographic, clinical, and laboratory results. Indicators related to each of the three categories are listed as follows.

- **Demographic information** (9 **indicators):** This category contains medical history and census information of an impatient, including gender, age, presence or absence of a set of underlying health conditions (e.g., hypertension, diabetes, cerebral infarction, and heart disease).
- Clinical and course examinations (12 indicators): This category includes chest radiographs or CT scan, symptoms at admission (e.g., fever, dry cough, sputum production, and fatigue), and daily routine tests results (e.g., pulse, respiration rate, blood pressure, body temperature, oxygen saturation, and heart rate).
- Laboratory indicators (173 indicators): The last category contains indicators from blood routine examination. That includes complete blood count (CBC), coagulation profile, serum biochemical tests, liver function, renal function electrolyte, blood lipid, blood glucose, procalcitonin detection, fluorescence, glucose determination, coagulation, respiratory tract infection pathogen, influenza A/B virus antigen detection, and other related indicators.

2) Severity Definitions: We divide the patients into 4 severity levels according to the Diagnosis and Treatment Plan for Corona Virus Disease 2019 (The 7.0 Trial Edition) [41]. The severity levels and the corresponding clinical symptoms are listed as follows:

- Severity Level I Mild: Asymptomatic, body temperature lower than 37.3 degrees, few or no visible lesion areas in the lungs.
- Severity Level II Moderate: Fatigue, dryness, constant coughing, body temperature over 37.3 degrees, multiple early lesions appear in the lungs.
- Severity Level III Severe: Blood oxygen saturation (SaO_2) less than 93%, Asthma, coughing, phlegm, and the increment of lesion areas within both lung lobes.

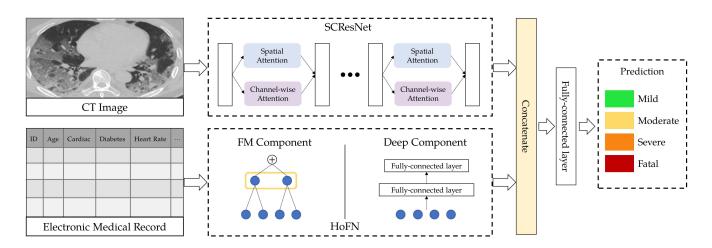


Fig. 3. The pipeline of the proposed multi-modality method. CT image and EMR data are used for multi-modality feature learning and fusion to predict COVID-19 patient's severity level.

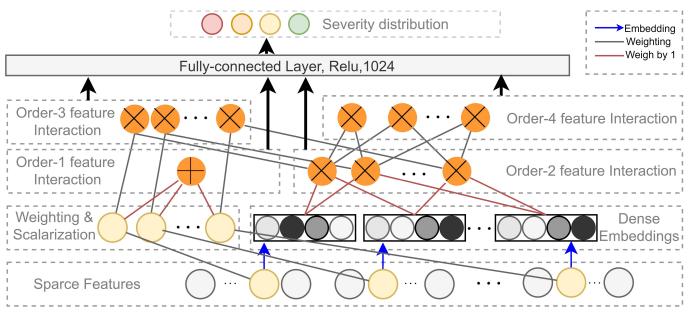


Fig. 4. Illustration of the high-order factorization component of the proposed method. A normal connection in black denotes a connection with weight to be learned. A connection in red indicate a connection with weight *s* by default. The blue arrow indicates a dense embedding is to learn. An addition sign in the yellow circle denotes the summation of all inputs and a cross sign in a yellow circle indicates the inner-product of the inputs.

• Severity Level IV Fatal: Multiple complications, requiring mechanical ventilation for live sustainment, and the lesion areas (mainly consolidation) continue to expand and occupy a large area of lung lobes.

3) CT Data Preprocessing: Image preprocessing is an essential step in medical image analysis tasks. In our case, proper localization of the lung area reduces the interferences from redundant information such as background noises and other organs. Following the widely-used setting in lung CT image processing [42], [43], it is effective to crop CT images based on the lung regions. In this paper, the input CT images are first converted into grayscale for both noise reduction and computational efficiency. After greyscale conversion, binarization, image erosion, and dilation are performed to separate the lung area from the rest of the image. Finally, image cropping and resizing are applied to eliminate background interferences while adjusting the cropped image to the size of 224×224 .

B. High-Order Factorization Network

To analyze EMR data, we propose a novel factorizationmachine-based neural network called High-order Factorization Network (HoFN). Unlike the conventional Deep Factorization-Machine (DeepFM) [27] and the Wide and Deep Model (Wide & Deep) [25], HoFN exploits the high-order interactions of features explicitly. To avoid possible confusion in mathematic notation, the term order-n in this paper denotes the interaction of n features from independent fields. Learning the consequence of interacting features are key to medical data analysis as well as modern diagnostics. For example, experiencing constant blood pressure rising is not something worrisome when a person is engaging in some high-pressure work. But, it is dangerous if a person also exhibits symptoms of myocardial infarction and hyperlipidemia. Frequently-used statistical models in medical data analysis (e.g., linear regression, logistic regression or survival analysis), however, fail

to generalize in the future when a large-scale dataset is not available [44]. In other words, statistically significant variables may be more easily overfitted towards the specific patient cohort in a certain region or certain health conditions if only low-order variable interaction is considered. As a result, there will be considerable uncertainty about the scientific merit of these findings. To alleviate this problem, the proposed HoFN network aims to leverage the high-order interaction of medical features as well as the nonlinearity in deep learning framework to make dataset-specific adaption.

Similar to the preliminary settings in most factorizationmachine-based models, we also utilize the EMR dataset for training. The EMR dataset contains a set of instances (χ, y) . $y \in \{0, 1, 2, 3\}$ corresponding to the 4 severity level of a patient defined in subsection III-A.2 while χ is an *m*-field EMR record representing the medical conditions of a patient.

After proper preprocessing as described in subsection III-A, a patient's condition $\chi = [x_1, x_2, \dots, x_m]$ includes *s* categorical fields and *d* numerical fields (m = s + d). Each categorical field is encoded using a one-hot encoder and is converted into a sparse representation while the numerical field is represented by its value. Since a categorical field has many potential values, its one-hot vector will exhibit a high level of sparsity. That's why they are often called sparse features. On the other hand, each continuous field is represented as the value itself, or as a one-hot vector after discretization.

With a patient's status χ , the task of predicting COVID-19 severity using EMR data can be formulated as $\hat{y} = HoFN(\chi)$. The proposed HoFN is consists of an FM component and a Deep component. Both components of HoFN are illustrated in more details as follows:

1) FM Component: The FM component of HoFN is a factorization machine to explicitly learn high-order interactions among a patient's clinical conditions for COVID-19 severity prediction. It models 4 levels of interactions explicitly. Linear (order-1) interactions, pairwise (order-2) interactions, triple-wise (order-3) interactions and biquadratic (order-4) interactions.

For low-order interactions (order-1 and order-2 interactions), the FM component models linear and pairwise feature interactions using inner projection of respective feature vectors. Given the *d*-dimensional input sparse feature list as $x = [x_1, x_2, \dots, x_d]$, the output of order-1 interaction $l^{(1)}$ is modelled by the linear combination of the sparse input features:

$$l^{(1)}(x) = \langle w, x \rangle = \sum_{i}^{d} w_i x_i, \qquad (1)$$

where $\langle \cdot, \cdot \rangle$ denotes the inner product operation, $w \in \mathbb{R}^d$ is the learnable weighting variables in the order-1 interaction, and each scalar in the weighting variables w_i is used to weigh the importance of an input sparse feature for computing the order-1 feature output. Meanwhile, a latent vector $V_i \in \mathbb{R}^{d \times e}$ is defined for each sparse feature x_i to encode the sparse feature as a dense embedding, where e is the pre-defined embedding size. Consequently, The embedding measures the sparse feature's impact on other sparse features when generating the order-2 interaction. In form, the output of the order-2 interaction $l^{(2)}$ is computed as follows:

$$l^{(2)}(x) = \sum_{j_1 > j_2}^d \langle V_{j_1}, V_{j_2} \rangle x_{j_1} x_{j_2}.$$
 (2)

To this end, it is noteworthy that when implementing the order-2 interaction, we use a more efficient algorithm (as Eq. 3) to reduce its time-complexity so that we don't need to compute the inner product of each pair of features as $p_{i,j} = \langle V_i, V_j \rangle x_i x_j$.

$$l^{(2)}(x) = \frac{1}{2} (\parallel V^{\mathrm{T}} x \parallel^2 - \sum_{i}^{d} V_i x_i)$$
(3)

For generating the higher-order interactions (order-3 and order-4 interactions), the embedding V and weights w from order-2 interaction are reused. The order-3 interaction considers the interaction among order-1 and order-2 features while the order-4 interaction considers the interaction between a pair of order-2 feature which resembles a hierarchical decision-making process. In form, the output of order-3 and order-4 interaction is calculated as follows:

$$l^{(3)}(x) = \sum_{j_1 > j_2 > j_3} u_{j_3} w_{j_3} x_{j_3} \cdot u_{(j_1, j_2)} \langle V_{j_1}, V_{j_2} \rangle x_{j_1} x_{j_2}$$

$$= \sum_{j_1 > j_2 > j_3} w_{j_3} \langle V_{j_1}, V_{j_2} \rangle x_{j_1} x_{j_2} x_{j_3}, \qquad (4)$$
$$l^{(4)}(x) = \sum_{j_1 > \dots > j_4} \mu_{(j_1, j_2)} p_{j_1, j_2} \cdot \mu_{(j_3, j_4)} p_{j_3, j_4}$$

$$= \sum_{j_1 > \dots > j_4} \mu_{(j_1, j_2)} p_{j_1, j_2} \cdot \mu_{(j_3, j_4)} p_{j_3, j_4}$$

$$= \sum_{j_1 > \dots > j_4} \mu_{(j_1, j_2)} \mu_{(j_3, j_4)} \langle V_{j_1}, \cdots, V_{j_4} \rangle x_{j_1} \cdots x_{j_4},$$
(5)

where $u \in R^{d+C_d^2}$ and $\mu \in R^{C_d^4}$ is the weighing for order-3 and order-4 interactions. The main benefit of the high-order interaction is to allow the model to learn the importance of feature combinations of a particular severity level that were not observed in the training dataset. Another benefit is the automation of feature engineering without tedious manual effort for feature searching.

Intuitively, the computation of order-3 interaction takes polynomial time in the number of input features since it involves C_d^3 combinations and the factorization technique used in Eq. 3 is no longer applicable. To solve this problem, this paper also provides an efficient algorithm to compute the output of order-3 interaction using ANOVA kernel [45], [46] and dynamic programming (DP). We first define the ANOVA kernel of degree $2 \le m$ by

$$A^{m}(V, x) := \sum_{j_{1} > \dots > j_{m}} \prod_{t}^{m} v_{j_{t}} x_{j_{t}}$$
$$A^{0}(V, x) := 1$$
$$A^{1}(V, x) := \langle v, x \rangle$$
(6)

where V is the embedding of input features. With multi-linearity as the key property of the ANOVA kernel (Eq. 7), we can compute the result recurrisvely. To compute

Algorithm 1 Order-3 Interaction

Input: order-1 weights $w \in R^d$, order-2 embeddings V ∈ $R^{d \times e}$, input $x \in R^d$, order-3 weights $u \in R^{2d}$; $L1_i \leftarrow w_i x_i, \forall i \in [1, 2, \dots, d]$ $L2_i \leftarrow V_i x_i, \forall i \in [1, 2, \dots, d]$ $L \leftarrow L1 \cup L2$ $a_{j,t}, b_{j,t}, c_{j,t} \leftarrow 0, \forall t \in [1, 2, 3], t < j$ $a_{j,0} \leftarrow 1, \forall j \in [1, 2, \dots, 2d]$ $b_{j,0}, c_{j,0} \leftarrow 1, \forall j \in [1, 2, \dots, d]$ for t := 1, ..., 3 do for j := t, ..., 2d - 1 do if j < d then $a_{j,t} \leftarrow a_{j-1,t} + u_j w_j L_j a_{j-1,t-1}$ $b_{j,t} \leftarrow b_{j-1,t} + u_j w_j b_{j-1,t-1}$ else $a_{j,t} \leftarrow a_{j-1,t} + u_j V_j L_j a_{j-1,t-1}$ $c_{j,t} \leftarrow c_{j-d-1,t} + u_j w_{j-d} c_{j-d-1,t-1}$ end if end for end for result $\leftarrow a_{2d,m} - b_{d,m} - c_{d,m}$ Output: result

the above high-order interactions efficiently, we present algorithm 1 using ANOVA kernel and dynamic programming (DP) algorithm. As a result, the time complexity of computing the order-3 interaction has been reduced from $\mathcal{O}(d^3)$ to $\mathcal{O}(d)$.

$$A^{m}(V, x) = A^{m}(V_{\neg j}, x_{\neg j}) + V_{j}x_{j}A^{m-1}(V_{\neg j}, x_{\neg j})$$
(7)

2) Deep Component: As shown in Fig. 5, different from the FM component which focuses on characterizing the feature's interaction at different levels, the deep component aims to capture the non-linear abstraction from the EMR data. To handle continuous features, a feed-forward multi-layer perceptron (MLP) is used to learn task-specific features. Specifically, each fully-connected layer in the MLP computes:

$$a^{(l+1)} = f(W^l a^l + b^l), (8)$$

where f is an activation function, a^{l} is the input for the l^{th} layer, W^{l} and b^{l} are the weight and bias a the l^{th} layer respectively. For l = 1, the input a^{1} is the concatenated numerical features.

C. Spatial-Channel Attention Residual Network

Empirically, professional medical practitioners focus on specific lesions around the lung region in a CT Image. For example, the large area of consolidation and fibrosis in a patient's lung lobes can be evidence for heavy infection. To concentrate on those critical lesion areas and yield discriminative visual features for severity prediction, we proposed to use the spatial-channel attention residual network (SCResNet) for the visual branch of our multi-modality feature learning and fusion model. The SCResNet emphasizes the crucial details and the lesion areas along the channel and spatial dimension which are beneficial for severity prediction. Fig. 6

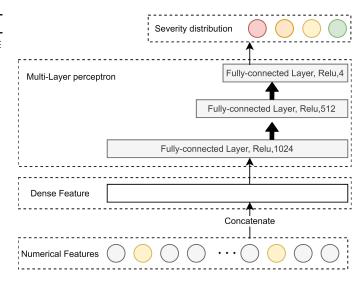


Fig. 5. Illustration of the deep component. Numerical features are concatenated into a dense feature vector before fed to an MLP. The MLP is consists of 3 fully-connected layers.

illustrates the spatial-channel attention module (SCAM). Given an output feature map of a convolutional layer $F \in \mathbb{R}^{C \times H \times W}$. we calculate the channel-wise attention $A_c \in \mathbb{R}^{C \times 1 \times 1}$ and spatial attention $A_s \in \mathbb{R}^{1 \times H \times W}$ separately. A_c and A_s are then utilized to update F_a which is the final output of the SCAM. Note that the SCAM can be applied after every convolutional layer. In our case, it is the residual layers. Mathematically, The output of the Spatial-channel attention F_a can be formularized as:

$$F_a = F \otimes A_c \otimes A_s, \tag{9}$$

where \otimes denotes element-wise multiplication.

1) Channel-Wise Attention: Channel-wise attention A_c extracts the inter-channel relationship of features. In convolution layers, each output channel of a feature map is generated by a convolution kernel that incorporates a learnable filter to scan through the input image. Meanwhile, each convolution kernel learns to extract a specific kind of internal feature from the input feature during the back propagation. Attention across channels enables the deep convolutional network to learn the right channel to focus on during the feed-forward process. The process of channel-wise attention is defined as follows.

To begin with, pooling operations is used to squeeze the spatial dimension. Both average-pooling and max-pooling are used for distinctive features conservation. Afterward, the pooling output is fed to a MLP for channel-wise attention learning. Finally, the sigmoid function is applied to scale the importance of each channel to a scale of [0, 1]. In form, the channel-wise attention is computed as follow:

$$A_{c} = \sigma \left(W_{1}(W_{0}(AvgPool(F) + b_{0}) + b_{1}) + W_{1}(W_{0}(MaxPool(F) + b_{0}) + b_{1}) \right)$$
(10)

where σ is the sigmoid function, F is the output feature map of the previous convolution layer, AvgPool and MaxPool

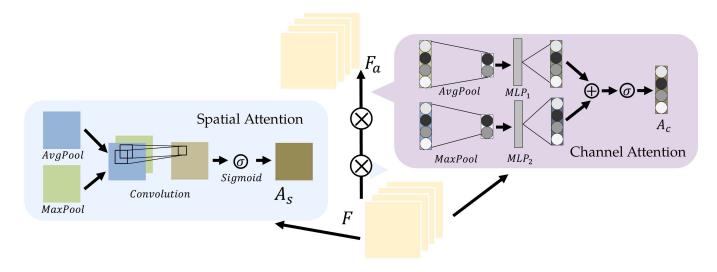


Fig. 6. Illustration of the attention module. The output feature map of a previous convolution layer is used to compute the spatial and channel-wise attention.

denote average-pooling and max-pooling operation respectively, W_0 , b_0 and W_1 , b_1 are the weights and biases of two fully-connected layers in the MLP. The details of which can be referred to in [47].

2) Spatial Attention: To attend the areas of interest in an input feature map, aggregation of spatial clues and computation of inter-spatial relationships are performed in the spatial attention module. Similar to the channel-wise attention, average-pooling and max-pooling operation are adopted for channel aggregation before convolution and normalization. In form, the output of the spatial attention A_s is calculated by the following equation:

$$A_s = \sigma(f^{3\times3}([AvgPool(F); MaxPool(F)]))$$
(11)

where $f^{3\times3}$ refers to a 3×3 filter size convolution, and [;] denotes vector concatenation. After A_c and A_s are computed, channel-wise and spatial attention weights are multiplied with the original input feature map for adaptive feature refinement.

3) SCResNet Architecture: The SCResNet, a pre-trained ResNet-50 integrated with spatial and channel-wise attention, is developed to classify chest CT images into different severity levels. Table I shows the configuration for each layer. Following each residual block, both the spatial attention and channel-wise attention are computed to capture significant details in each block. Finally, the fully-connected layer is modified for the classification of the COVID-19 severity level.

D. Inter-Modality Similarity Loss

The cohesion of a patient's lung condition assessment and clinical data diagnosis is of utmost importance for a reliable severity estimation algorithm. In the scenario of field diagnosis, given a clinical report of a patient, medical practitioners can circumscribe his lung condition shown in the chest CT image and vice versa. This correspondence indicates that visual or clinical clues share a similar abstract concept of the

TABLE I The Configuration Setting for Spatial-Channel-Attention ResNet

Layer name	Output size	Configuration	
input	256 × 256	-	
conv1	64×64	7×7 , 64, stride 2	
COIIVI	04 × 04	3×3 , max pool, stride 2	
		$1 \times 1,64$	
		$3 \times 3,64$	
conv2	64×64	$1 \times 1,256$	$\times 3$
		channel(avg&max, MLP)	
		spatial(avg&max, 3×3 conv)	
		$1 \times 1, 128$	
		$3 \times 3, 128$	
conv3	32×32	$1 \times 1,512$	$\times 4$
		channel(avg&max, MLP)	
		spatial(avg&max, 3×3 conv)	
		$1 \times 1,256$	
		3 imes 3,256	
conv4	16×16	$1 \times 1,1024$	$\times 6$
		channel(avg&max, MLP)	
		spatial(avg&max, 3×3 conv)	
		$1 \times 1,512$	
		$3 \times 3,512$	
conv5	8×8	$1 \times 1,2038$	$\times 3$
		channel(avg&max, MLP)	
		spatial(avg&max, 3×3 conv)	
hidden layer	1024	fc(1024,Relu)	
hidden layer	512	fc(512,Relu)	
hidden layer	4	fc(4,Relu)	
output	4	softmax	
¹ fc denotes f	ully-connected	layer.	

² conv denotes convlution layer.

 3 avg and max denote the average and max pooling operation respectively. ⁴ channel(·) and spatial(·) denote the channel-wise and spatial attention respectively.

patient's condition. However, many widely-used loss function in deep learning models, such as the cross-entropy loss, fails to extract modality-shared information effectively. In this paper, instead of solely using cross-entropy loss to train each branch of the network and extract severity-level-specific information for classification, we propose a new loss function to exploit the intra-class similarity and the cross-modality cohesion for severity prediction. To learn modality-shared information, we design the following loss function which enforces that the feature learning from EMR data and CT image data to be performed in the shared feature space by reducing the cosine similarity between both features. Meanwhile, we penalize the distance between features from two modalities to improve intra-class similarity in each severity level by using the hetero-centre loss in our inter-modality similarity loss (IMSLoss). To this end, the IMSLoss L_{ims} is readily defined in the following equation:

$$L_{ims} = \sum_{i}^{|I|} 1 - \cos(h_i^{emr}, h_i^{ct}) + \sum_{c}^{C} (z_c^{emr} - z_c^{ct}), \quad (12)$$

where $cos(h_i^{emr}, h_i^{ct})$ is the cosine similarity of the EMR feature h_i^{emr} and CT image feature h_i^{ct} extracted from the i^{th} training sample, I denotes the training dataset, |I| is the size of the dataset, and C denotes the number of classes, which is 4 in our scenario. z_c^{emr} and z_c^{ct} denote the center of EMR and CT features belonging to the c^{th} class. Let I^c denotes the subset of the training samples belonging to the c^{th} class. The center of both feature's distribution are computed as $z_c^{emr} = \frac{1}{|I^c|} \sum_{k}^{|I^c|} h_k^{emr}$ and $z_c^{ct} = \frac{1}{|I^c|} \sum_{k}^{|I^c|} h_k^{et}$. As a result, a cross-modality model can learn to capture information shared by different modalities during the training process. More details of the hetero-centre loss can be refferred to [48]. Dispite using the IMSLoss for refined multimodality feature learning, the cross-entropy loss L_{ce} (Eq. 13) is also used for training the classification model.

$$L_{ce} = -\log \prod_{i=1}^{|I|} p(y_i | \chi_i, M_i),$$
(13)

where |I| denotes the size of the training dataset, χ_i and M_i are the EMR data and CT image from the *i*th training sample respectively, y_i indicates the *i*th severity label. Finally, the total loss *L* used to train the proposed multi-modality model is defined as:

$$L = \lambda L_{ims} + L_{ce}, \tag{14}$$

where λ is a factor to control the impact of the IMSLoss in the training process. In our experiments, λ is set to 0.1.

IV. EXPERIMENTS

To validate the proposed method, we collected the training and testing data as described in Section III-A and evaluate the proposed algorithm on the dataset. The experiments are implemented on the environment of Python 3.6 and operating system of Windows 10. Preprocessing like greyscale transformation and resizing are executed through OpenCV 3.4.2 for the ease of further analysis. The proposed model is implemented using the PyTorch deep learning platform and is trained on a platform with AMD Ryzen 7 2700X Eight-Core 3.7 GHz in CPU and NVIDIA GeForce GTX1080 Ti with CUDA 9.0 in GPU. All parameters of the proposed model are optimized by the adaptive subgradient method (Adagrad) [49] for faster convergence speed and better performance. The learning rate is set to 0.01. On average, the model can fully converged at around 50 epochs.

A. Wuhan COVID-19 Retrospective Dataset

The dataset used to evaluate the effectiveness of the proposed method is originated from a retrospective study consisting of 733 patients diagnosed with COVID-19 disease. Patients in the retrospective study dataset were admitted to Huangpi Hospital of Traditional Chinese Medicine (Wuhan, China) from January 2020 to March 2020. This dataset is desensitized and approved by the First Affiliated Hospital of Guangxi Medical University Hospital Ethics Committee and the requirement for informed consent was waived. The retrospective dataset is consists of the complete blood count (CBC) test outcomes, diagnosis, clinical characteristics, the patient's recovery situation, and their chest CT images. These multimodality data sources can be utilized to train and test our proposed severity prediction model. For each patient's record contained in the dataset, the patient's severity level is annotated according to their corresponding relationship with the symptom as illustrated in subsection III-A and their recovery situation. A list of sample images is shown in Fig. 7. The COVID-19 disease causes 6 major types of lung lesions including ground-glass opacification (GGO), halo sign (HS), reversed halo sign (RHS), septal thickening (ST), consolidation, and fibrosis. More detailed information about the appearance and morphological characteristics of each type of lesion is included in Table VI and illustrated in Fig. 7. As shown in Fig. 7, few or no visible lesions exhibits in the lungs of mild patients. As the severity level increases, the lesions area of the lung lobes gradually increases until it covers the majority of the lung. As a result of diversity among individuals' physical condition, the lung condition may not be the only indicator to determine the COVID-19 severity. For instance, although both lungs have obvious consolidation lesions, the 7th patient in the first row of Fig. 7 has no other underlying diseases. At the same time, since his blood oxygen saturation, heart rate and other medical indicators are also within normal range he only exhibits mild symptoms and has recovered very well. These observation also validate the necessity of utilizing multi-modality data in our proposed method.

Similar to existing COVID-19 research datasets like the TriNetX COVID dataset [50], [51], both datasets contain the same groups of medical data such as a patient's CBC outcomes, characteristics (i.e., age, gender), and their prior (or coexisting) conditions. However, minor differences remain in the components in each group. For example, TriNetX COVID dataset contains more underlying medical conditions such as dementia, rheumatic disease and cancer that are not displayed in our dataset while our dataset contains more detailed outcomes from the CBC test. These differences are due to varying data entry protocols and bespoke implementations of different EMR systems. In general, however, the overall quality of EMR data in our dataset is consistent with existing publicly available COVID-19 datasets. In addition, our dataset also contains CT images that further help us to perform a more comprehensive multimodality analysis.

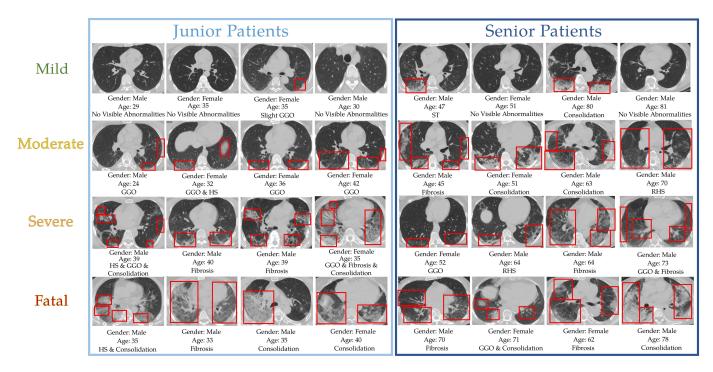


Fig. 7. Samples of the chest CT images in the dataset after preprocessing. Chest CT image from patients under mild, moderate, severe and fatal levels are shown from top to bottom. Gender, age and type(s) of lesion of the inpatients are also provided under each CT image.

TABLE II

DIVISION BASIS OF THE D	DATASET
-------------------------	---------

	Patient being asymptomatic or only	
Mild	exhibiting mild respiratory symptoms.	
Milia	Lung CT shows little or no visible pneumonia.	
	Patient survive and recover	
	Patient experience obvious fever,	
Moderate	respiratory disease and other related symptoms	
Wilderate	Lung CT shows visible signs of pneumonia.	
	Patient eventually survive	
	and most lung lesions being recovered.	
	Increased lung CT lesion area.	
Severe	Patient experiences heavy respiratory distress,	
	or requires auxiliary equipment for oxygen.	
	Patient eventually survive.	
	Patient experience one of the following	
	situations during the hospitalization:	
Fatal	Shock	
raidl	Respiratory failure	
	Need for ventilation machine for life substance	
	Death	

To evaluate the proposed method, our dataset is divided into two subsets for training and validation. Distribution among severity levels in the training subset and the validation subset are shown in Table III. Table II shows the basis for the patient's division. In our dataset, The majority of patients were experiencing mild and moderate COVID-19 symptoms while 73 of them were having severe or fatal symptoms. Given this relatively small dataset scale, using a pre-trained network parameter for our convolutional network to reduce overfitting is quite necessary.

B. Metrics

In order to compare the effectiveness of the proposed method quantitatively, the accuracy P of the convergence

TABLE III DISTRIBUTION OF SEVERITY LEVEL IN THE TRAINING SUBSET AND THE TEST SUBSET

Severity	Number of sample	Training Samples	Test Samples
Mild	120	84	36
Moderate	540	378	162
Severe	40	28	12
Fatal	33	23	10
Total	733	513	220

model can be obtained by:

$$P = \frac{Num_{correct}}{Num},\tag{15}$$

where Num and $Num_{correct}$ are the total number of ground truth labels and the number of correct prediction in the validation subset.

C. Evaluation

In our experiment, we compare our proposed method with the following four widely-used machine learning methods. Categorical and numerical fields from the EMR data are simply concatenated together as input for these methods.

1) **Logistic Regression (LR)**: A Logistic regression method is employed for COVID-19 patient severity classification using the EMR data.

2) **Support Vector Machine (SVM)**: The features from the EMR data are fed into a SVM classifier using linear kernel, and C = 1.

3) **Decision Tree**: The decision tree is used to classify the clinical data into each severity category.

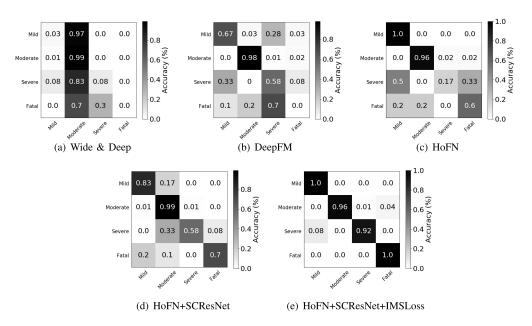


Fig. 8. Confusion matrices of Wide & Deep model, DeepFM, HoFN, HoFN+SCResNet, HoFN+SCResNet+IMSLoss on the Wuhan COVID-19 retrospective dataset.

TABLE IV Comparison With Different Methods

Method	Input	Accuracy(%)
Decision Tree	EMR	63.18
MLP	EMR	72.27
Logistic	EMR	72.73
SVM	EMR	73.64
Random Forest	EMR	73.64
Wide & Deep	EMR	74.09
DeepFM	EMR	85.91
HoFN	EMR	90.45
HoFN+SCResNet (Without MS Loss)	EMR+CT	93.18
HoFN+SCResNet (with MS Loss)	EMR+CT	96.36

4) **Random Forests (RF)**: The random forest classifier is applied to the medical features set for patient severity classification. The number of trees in a random forest is 100.

5) **Multi-Layer Perceptron (MLP)**: A MLP is applied to learn the raw input data. In particular, The MLP consists of 4 layers including an input layer and an output layer. The hidden layers consist of a fully-connected layer with 1024 perceptrons and a fully-connected layer with 512 perceptrons. Both hidden layers are using Relu as activation function.

6) Wide & Deep Network [25]: The wide and deep network is adopted. The categorical features are fed into the deep component after embedding while the numerical features are concatenated and fed to the deep component.

7) Factorization-Machine based neural network (DeepFM) [27]: The DeepFM is an end-to-end model which integrates factorization machines and MLP. It also captures linear and pairwise interactions between features and is adopted as a compared method in the ablation experiment since it doesn't model higher-order interactions.

As illustrated in Table IV, compared to conventional machine learning methods, our method achieves the best result in terms of accuracy. Compared to MLP, the baseline neural

network architecture, our method exceeds its performance by a large margin, indicating that the learning of feature interaction plays an essential role in feature selection and task-specific feature generation. Compared with Wide & deep neural network which is a widely-used method to analyze both numerical and categorical features, HoFN and DeepFM prevail since the factorization machine architecture is able to jointly leverage the co-occurrence of features. Without explicitly using the cross-feature intersection [25], the wide & deep network lacks the automatic feature engineering capacity. On the other hand, HoFN has an edge over DeepFM since more information is aggregated through the higher-order interactions.

For the sake of internal comparison, a HoFN model that only utilizes EMR data falls short compared to the joint utilization of both multimodality by 2.73%. Indicating that the proposed multi-modality feature learning and fusion model yields a significant improvement in performance. Furthermore, the use of multimodality loss further enhances the reliability of the proposed method, bringing the overall prediction accuracy to 96.36%.

Fig. 8 shows the confusion matrices corresponding to Wide & Deep network, DeepFM, HoFN, HoFN+SCResNet, and HoFN+SCResNet(IMSLoss) on the collected dataset. Fig. 8(a) shows that using the Wide & Deep Network, most patients in the test subset is classified as moderate. This is mainly due to the fact Wide & Deep can not make use of feature interactions without manual feature engineering. Compared with the Wide & Deep model (Fig. 8(a)), HoFN (Fig. 8(c)) and DeepFM (Fig. 8(b)) are more capable to distinguish some mild and severe patients from moderate ones since the explicit utilization of both low and high-order feature interactions. After incorporating visual features, the lung condition serves as an important factor to discriminate fatal and severe cases from the moderate ones. As depicted in Fig. 8(d), the accuracy of correctly recognizing the severe and fatal cases

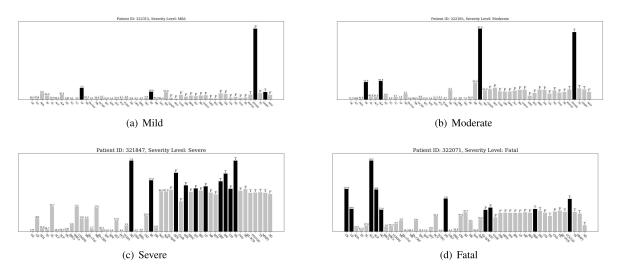


Fig. 9. Importance scores of input features (displayed in abbreviation form) are normalized and displayed by bar charts for different severity level at the bottom of each subgraph. Darker bar denotes the salient features, grey bar denotes the non-salient feature. Numbers or letters displayed on each bar are the original values of the corresponding input features. The letter T means the corresponding symptom exists while the letter F means the opposite. The full name of each feature displayed in the figures can be found in Table VII.

TABLE V Most Frequently Features Interactions

Level	Order	Features
		AD, PI, COUGH
	Order-3	PI, HL, COUGH
Severe		RT–PCR, PI, COUGH
Severe		ADV, COUGH, HL, CoxB
	Order-4	CoxB, RT–PCR, HL, COUGH
		CoxB, TS,AD, COUGH
	Order-3	ADV, RT–PCR, COUGH
		ADV, AD, COUGH
Fatal		ADV, PI, COUGH
1 aidi	Order-4	ADV, PI, HL, COUGH
		CoxB, AD, HL, COUGH
		ADV, COUGH, HL, CoxB

¹ full names of the abbreviations in this table can be found in Table VII.

has increased by 8% and 50% respectively. Finally, as shown in Fig. 8(e), the use of inter-modality similarity loss further enhances the overall performance in all categories, demonstrating the effectiveness of the proposed multi-modality method.

D. Attribution Evalation

Understanding a deep learning model's decision is not only an important aspect to improve the transparency, persuasiveness and trustworthiness for medical practitioners but also a way to reveal bias or other unexpected patterns of a certain disease. This subsection interprets the proposed model from an attribution perspective. We also explain the meaning of some highly-focused feature interactions from a clinical point of view.

To evaluate how a patient's EMR data contribute to the model's prediction, we utilized a simple but widely-adopted gradient-based saliency method called saliency score (Gradient \odot Input) [52], [53]. In a formal setup, a model describes a function $S : \mathbb{R}^d \to \mathbb{R}^C$, where *C* is the number of classes. The

TABLE VI Appendix.A: Major Types of Lung Lesions Caused by COVID-19 Desease

Lesion Type	Appearance	
No visible abnormalities	The lung lobes are healthy and has	
No visible abilitinanties	no visible lesion.	
	An area of hazy increased lung opacity through	
Ground-glass opacity [56]	where vessels and bronchial structures may	
	still be seen. Less opaque than consolidation	
	A circumference of a nodule or mass surrounded	
Halo sign [57]	by GGO. Representing a focus of pulmonary	
	infarction surrounded by alveolar hemorrhage.	
	A distinct radiological sign representing a	
Reversed halo sign [57]	focal rounded area of GGO surrounded by a	
	crescent or complete ring of consolidation.	
	The lesion is distributed in a grid pattern,	
Septal thickening [58]	which is a characteristic manifestation of	
	local tissue fibrosis.	
Cancelidation [50]	A region of normally compressible lung tissue	
Consolidation [59]	that has filled with liquid instead of air.	
Fibrosis [60]	A lung disease that occurs when lung tissue	
1.1010818 [00]	becomes damaged and scarred.	

Gradient \odot Input method computes the element-wise product of the input and the gradient, denoted $x \odot \frac{\partial S}{\partial x}$, which has been applied to address gradient saturation issue and highlight the more relevant features from the rest. Since it has been widely-used to explain models in visual or sequential data classification tasks [54], [55], we adopt this method to help us understand the contribution of each medical feature in our model. Furthermore, to show the impact of the interactions, interaction feature with the highest weighting is displayed in Table V. Meanwhile, Fig 9 displays the saliency map from each severity level.

For the mild and moderate severety level, Fig. 9(a) shows the attribution for the proposed model to classify a patient's severity level as mild. Features with high importance score include cough & fatigue, body temperature, age, and lymphocyte (%). Among them, the main reason for the model's decision is that the patient has no symptoms of cough &

TABLE VII

A

APPENDIX.B: FEATURE'S NAME AND ABBREVIATI	ON
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	N	Normal Reference
Abbreviation		Value Range
CK	Creatine Kinase	$18.0 \sim 198.0 U/L$
LD or LDH	Lactate Dehydrogenase	$200 \sim 380 U/L$
ADA	Adenosine Deaminase	$4 \sim 22U/L$
TP	Total Protein	$60 \sim 80 g/L$
AL	Albumin	$35 \sim 51 q/L$
ALT	Alanine Aminotransferase	$0 \sim 40 U/L$
		Female: $50 \sim 135U/L$
ALP	Alkaline Ahosphatase	Male: $45 \sim 125U/L$
		Female: $120 \sim 165 g/L$
HC	Haemoglobin Concentration	Male: $130 \sim 185g/L$
		Female: $4 \sim 5 \times 10^6 / mm^3$
RBCs	Red Blood Cells	Male: $4.5 \sim 5.5 \times 10^6 / mm^3$
WDC		$3.5 \sim 9.5 \times 10^3 / mm^3$
WBCs	White Blood Cells	$3.5 \sim 9.5 \times 10^{\circ}/mm^{\circ}$
LYMPH	Lymphocyte (%)	$20\% \sim 40\%$
MC	Monocytes(%)	Female: $3 \sim 10\%$
	•	Male: $3 \sim 10\%$
Hs-cTnI	High-sensitivity Cardiac Troponins	
Mb	Myoglobin	$29 \pm 16.3 g/L$
Hs-CRP	Hypersensitive C-reactive protein	< 1.0 mg/L
IgG	Immunoglobulin G	$9.5 \sim 12.5 mg/mL$
IgM	Immunoglobulin M	$0.6 \sim 2.5 g/L$
AST	Aspartate Aminotransferase	$0 \sim 40 U/L$
PCT	Procalcitonin	< 0.1 ng/ml
TCHO	Total Cholesterol	$2.83 \sim 5.20 mmol/L$
DD	D-Dime	< 0.5 mg/L
LIDO	TT 1'1'	Female: $0 \sim 2.64 \mu mol/L$
URO	Urobilinogen	Male: $0.30 \sim 3.55 \mu mol/L$
DBIL	Direct Bilirubin	$0 \sim 6.8 \mu mol/L$
CKI	Creatine Kinase Isoenzymes	< 5%
BT	Body Temperature	$36.3 \sim 37.2$ °
HR	Heart Rate	$60 \sim 100/min$
SaO2	Oxygen Saturation	> 94%
Age	Age	_
Cp-IgM	Chlamydia pneumoniae-IgM	_
ADV	Adenoviridae	
CoxB	Coxsackievirus-B Type	
OfR	Q fever Rickettsia	_
HPIV	Human Parainfluenza Virus	_
IAV		_
	Influenza A Virus	_
IBV	Influenza B Virus	-
LP	Legionella Pneumophila	_
MP	Mycoplasma Pneumonia	-
RSV	Respiratory Syncytial Virus	-
HBP	High Blood Pressure	-
DM	Diabetes Mellitus	-
CIS	Cerebral Ischemic Stroke	-
HS	Hepatorenal Syndrome	-
CVD	Cardiovascular Disease	-
HL	Hyperlipidemia	_
RT-PCR	Real Time Reverse Transcription	_
	-Polymerase Chain Reaction.	
COUGH	Cough	_
TS	Thick Sputum	-
	51 101	
PI	Pulmonary Infections	—

fatigue. From a pathological point of view, although this patient has positive RT-PRC test result and increasement in percentage of lymphocytes (indicating a certain degree of inflammation), other indicators are basically in the normal range. Besides, since the patient's body temperature is below 37 °C, and he has no respiratory disease (i.e., cough & fatigue), it is reasonable to assign the patient to the mild level. Likewise, Fig. 9(b) shows the importance score for a patient of moderate severity level. Different from the mild case, the model focuses heavily on cough & fatigue and SAO2. Taking the patient's actual condition into consideration, we could summarize that the model assigns the patient to the moderate level mainly because the patient does not have too many abnormalities

except coughing and fatigue while his blood oxygen saturation remains within the normal range.

For the severe and fatal severety level, a person at a severe level is displayed in Fig. 9(c). Judging from his original EMR data, this confirmed COVID-19 case already has comorbidities such as hypertension and diabetes before admission. More severely, he has a hepatorenal syndrome and high D-dimer value which indicate he is likely or has been suffering from vital organ damage. In correspondence, the importance score of the abovementioned indicators is higher than others. Compared with the mild and moderate cases, high body temperature has also become a feature of high importance. In contrast, some relatively normal indicators such as total protein and albumin have little effect on making the decisions. Finally, the importance score shown in Fig. 9(d) is from a fatal case. The model focuses on comorbidities (diabetes and cardiopathy), and a set of extremely high blood test values including creatine kinase, lactate dehydrogenase, alanine aminotransferase, and alkaline phosphatase.

Table. V demonstrates the most frequently-appeared high-weighted interactions seen in severe and critical-level cases. Because of our model's capacity to explicitly model high-dimensional feature interactions and learn features from multiple inputs, the combination of some variables which are statistically insignificant if they are treated as independent variables can also become a very influential decision basis. In addition, through statistical and analytical approach, we could find the potential patterns of a diseases after the proposed method have been fully-trained on a large dataset, which is conducive to more efficient judgment of the actual or possible conditions of patients.

V. CONCLUSION AND FUTURE WORK

This paper puts forward a novel multi-modality feature learning and fusion model for COVID-19 patient severity prediction. To cope with categorical and numerical features extracted from a patient's EMR, the HoFN is proposed to explicitly learn the high-order and low-order interactions of a patient's condition. To locate and analyze the patient's lung infection situation, a pre-trained CNN with spatial and channel-wise attention is used to generate discriminative visual features. Finally, the IMSLoss is applied to help the model learn modality-shared features which is beneficial for modality fusion. To facilitate the evaluation of the proposed model, a dataset containing COVID-19 inpatients is collected from a medical institution.

From the experimental results and comparisons in Section IV, it can be concluded that the proposed multimodality method is very competitive in real-life conditions. The use of HoFN and pre-trained SCResNet are essential to learning discriminative features to evaluate the patient's severity level while the inter-modality loss is able to improve the performance by introducing cross-modality subspace similarity. However, the proposed method failed to make use of any temporal features of an impatient which could be an important cue to tell whether the situation of a patient is getting better or worse. The exploration of novel methods to incorporate temporal features and new design of optimization models for more effective heterogeneous modality fusion further explored in our future work.

REFERENCES

- W. H. Organization. (2021). Weekly Epidemiological Update—23 February 2021. [Online]. Available: https://www.who.int/publications/m/ item/weekly-epidemiological-update—%-23-february-2021
- [2] C. Sohrabi et al., "World health organization declares global emergency: A review of the 2019 novel coronavirus (COVID-19)," Int. J. Surgery, vol. 76, pp. 71–76, Apr. 2020.
- [3] M. Barstugan, U. Ozkaya, and S. Ozturk, "Coronavirus (COVID-19) classification using CT images by machine learning methods," 2020, arXiv:2003.09424. [Online]. Available: http://arxiv.org/abs/2003.09424
- [4] C. Galván Casas *et al.*, "Classification of the cutaneous manifestations of COVID-19: A rapid prospective nationwide consensus study in Spain with 375 cases," *Brit. J. Dermatology*, vol. 183, no. 1, pp. 71–77, 2020.
- [5] S. Hu *et al.*, "Weakly supervised deep learning for COVID-19 infection detection and classification from CT images," *IEEE Access*, vol. 8, pp. 118869–118883, 2020.
- [6] L. Sun, G. W. DePuy, and G. W. Evans, "Multi-objective optimization models for patient allocation during a pandemic influenza outbreak," *Comput. Oper. Res.*, vol. 51, pp. 350–359, Nov. 2014.
- [7] S.-H. Kim, C. W. Chan, M. Olivares, and G. Escobar, "ICU admission control: An empirical study of capacity allocation and its implication for patient outcomes," *Manage. Sci.*, vol. 61, no. 1, pp. 19–38, Jan. 2015.
- [8] L. A. Goncalves et al., "Nursing allocation and adverse events/incidents in intensive care units," Revista da Escola de Enfermagem da USP, vol. 46, pp. 71–77, Oct. 2012.
- [9] B. Khaleghi, A. Khamis, F. O. Karray, and S. N. Razavi, "Multisensor data fusion: A review of the state-of-the-art," *Inf. Fusion*, vol. 14, no. 1, pp. 28–44, Jan. 2013.
- [10] D. Lahat, T. Adali, and C. Jutten, "Multimodal data fusion: An overview of methods, challenges, and prospects," *Proc. IEEE*, vol. 103, no. 9, pp. 1449–1477, Sep. 2015.
- [11] J. Gao, P. Li, Z. Chen, and J. Zhang, "A survey on deep learning for multimodal data fusion," *Neural Comput.*, vol. 32, no. 5, pp. 829–864, May 2020.
- [12] S. Antol et al., "VQA: Visual question answering," in Proc. IEEE Int. Conf. Comput. Vis. (ICCV), Dec. 2015, pp. 2425–2433.
- [13] L. Yang, H. Sahli, X. Xia, E. Pei, M. C. Oveneke, and D. Jiang, "Hybrid depression classification and estimation from audio video and text information," in *Proc. 7th Annu. Workshop Audio/Visual Emotion Challenge*, Oct. 2017, pp. 45–51.
- [14] J. Chen, B. Sun, H. Li, H. Lu, and X.-S. Hua, "Deep CTR prediction in display advertising," in *Proc. 24th ACM Int. Conf. Multimedia*, Oct. 2016, pp. 811–820, doi: 10.1145/2964284.2964325.
- [15] D. Wang and K. Mao, "Learning semantic text features for Web textaided image classification," *IEEE Trans. Multimedia*, vol. 21, no. 12, pp. 2985–2996, Dec. 2019.
- [16] D. Wang and K. Mao, "Task-generic semantic convolutional neural network for Web text-aided image classification," *Neurocomputing*, vol. 329, pp. 103–115, Feb. 2019.
- [17] A. Samareh, Y. Jin, Z. Wang, X. Chang, and S. Huang, "Predicting depression severity by multi-modal feature engineering and fusion," in *Proc. AAAI Conf. Artif. Intell.*, vol. 32, no. 1, 2018, pp. 1–2.
- [18] A. Samareh, Y. Jin, Z. Wang, X. Chang, and S. Huang, "Detect depression from communication: How computer vision, signal processing, and sentiment analysis join forces," *IISE Trans. Healthcare Syst. Eng.*, vol. 8, no. 3, pp. 196–208, Jul. 2018.
- [19] P. Anderson *et al.*, "Bottom-up and top-down attention for image captioning and visual question answering," in *Proc. IEEE/CVF Conf. Comput. Vis. Pattern Recognit.*, Jun. 2018, pp. 6077–6086.
- [20] J. Lu, J. Yang, D. Batra, and D. Parikh, "Hierarchical question-image co-attention for visual question answering," in *Proc. Adv. Neural Inf. Process. Syst.*, 2016, pp. 289–297.
- [21] COVID-19 Map—Johns Hopkins Coronavirus Resource Center. Accessed: Nov. 7, 2020. [Online]. Available: https://coronavirus.jhu.edu/ map.html
- [22] A. Salcedo-Bernal, M. P. Villamil-Giraldo, and A. D. Moreno-Barbosa, "Clinical data analysis: An opportunity to compare machine learning methods," *Procedia Comput. Sci.*, vol. 100, pp. 731–738, Jan. 2016.
- [23] W. Liang *et al.*, "Development and validation of a clinical risk score to predict the occurrence of critical illness in hospitalized patients with COVID-19," *JAMA Internal Med.*, vol. 180, no. 8, p. 1081, Aug. 2020.

- [24] D. Chen *et al.*, "Deep learning and alternative learning strategies for retrospective real-world clinical data," *NPJ Digit. Med.*, vol. 2, no. 1, pp. 1–5, Dec. 2019.
- [25] H.-T. Cheng et al., "Wide & deep learning for recommender systems," in Proc. 1st Workshop Deep Learn. Recommender Syst., Sep. 2016, pp. 7–10, doi: 10.1145/2988450.2988454.
- [26] C. Chen, P. Zhao, L. Li, J. Zhou, X. Li, and M. Qiu, "Locally connected deep learning framework for industrial-scale recommender systems," in *Proc. 26th Int. Conf. World Wide Web Companion - WWW Companion*, 2017, pp. 769–770.
- [27] H. Guo, R. Tang, Y. Ye, Z. Li, and X. He, "DeepFM: A factorizationmachine based neural network for CTR prediction," in *Proc. 26th Int. Joint Conf. Artif. Intell.*, Aug. 2017, pp. 1725–1731.
- [28] J. Zhou, X. Zhang, Y. Liu, and X. Lan, "Facial expression recognition using spatial-temporal semantic graph network," in *Proc. IEEE Int. Conf. Image Process. (ICIP)*, Oct. 2020, pp. 1961–1965.
- [29] Y. Ye, X. Zhang, Y. Lin, and H. Wang, "Facial expression recognition via region-based convolutional fusion network," J. Vis. Commun. Image Represent., vol. 62, pp. 1–11, Jul. 2019.
- [30] O. Ronneberger, P. Fischer, and T. Brox, "U-Net: Convolutional networks for biomedical image segmentation," in *Medical Image Computing and Computer-Assisted Intervention—MICCAI.* Cham, Switzerland: Springer, 2015, pp. 234–241.
- [31] K. Kamnitsas *et al.*, "Efficient multi-scale 3D CNN with fully connected CRF for accurate brain lesion segmentation," *Med. Image Anal.*, vol. 36, pp. 61–78, Feb. 2017.
- [32] G. Litjens et al., "A survey on deep learning in medical image analysis," Med. Image Anal., vol. 42, pp. 60–88, Dec. 2017.
- [33] S. Wang et al., "A deep learning algorithm using ct images to screen for corona virus disease (COVID-19)," MedRxiv, pp. 1–9, Feb. 2021.
- [34] Y. Song *et al.*, "Deep learning enables accurate diagnosis of novel coronavirus (COVID-19) with CT images," *IEEE/ACM Trans. Comput. Biol. Bioinf.*, p. 1, Mar. 2021, doi: 10.1109/TCBB.2021.3065361.
- [35] L. Wang and A. Wong, "COVID-net: A tailored deep convolutional neural network design for detection of COVID-19 cases from chest Xray images," 2020, arXiv:2003.09871. [Online]. Available: http://arxiv. org/abs/2003.09871
- [36] J. Deng, W. Dong, R. Socher, L.-J. Li, K. Li, and L. Fei-Fei, "ImageNet: A large-scale hierarchical image database," in *Proc. IEEE Conf. Comput. Vis. Pattern Recognit.*, Jun. 2009, pp. 248–255.
- [37] J. Dolz, K. Gopinath, J. Yuan, H. Lombaert, C. Desrosiers, and I. Ben Ayed, "HyperDense-net: A hyper-densely connected CNN for multimodal image segmentation," *IEEE Trans. Med. Imag.*, vol. 38, no. 5, pp. 1116–1126, May 2019.
- [38] D. Nie, L. Wang, Y. Gao, and D. Shen, "Fully convolutional networks for multi-modality isointense infant brain image segmentation," in *Proc. IEEE 13th Int. Symp. Biomed. Imag. (ISBI)*, Apr. 2016, pp. 1342–1345.
- [39] L. Chen, Y. Wu, A. M. DSouza, A. Z. Abidin, A. Wismüller, and C. Xu, "MRI tumor segmentation with densely connected 3D CNN," *Proc. SPIE*, vol. 10574, Mar. 2018, Art. no. 105741F
- [40] M. Liang, Z. Li, T. Chen, and J. Zeng, "Integrative data analysis of multi-platform cancer data with a multimodal deep learning approach," *IEEE/ACM Trans. Comput. Biol. Bioinf.*, vol. 12, no. 4, pp. 928–937, Jul. 2015.
- [41] N. H. Commission *et al.*, "Diagnosis and treatment protocol for novel coronavirus pneumonia (trial version 7)," *Chin. Med. J.*, vol. 133, no. 9, pp. 1087–1095, 2020.
- [42] A. Bhandary *et al.*, "Deep-learning framework to detect lung abnormality—A study with chest X-ray and lung CT scan images," *Pattern Recognit. Lett.*, vol. 129, pp. 271–278, Jan. 2020.
- [43] M. S. Al-Tarawneh, "Lung cancer detection using image processing techniques," *Leonardo Electron. J. Practices Technol.*, vol. 11, no. 21, pp. 147–158, 2012.
- [44] M. A. Babyak, "What you see may not be what you get: A brief, nontechnical introduction to overfitting in regression-type models," *Psychosomatic Med.*, vol. 66, no. 3, pp. 411–421, May 2004.
- [45] M. Blondel, M. Ishihata, A. Fujino, and N. Ueda, "Polynomial networks and factorization machines: New insights and efficient training algorithms," in *Proc. Int. Conf. Mach. Learn.*, 2016, pp. 850–858.
- [46] M. Blondel, A. Fujino, N. Ueda, and M. Ishihata, "Higher-order factorization machines," in *Proc. NIPS*, 2016, pp. 3359–3367.
- [47] S. Woo, J. Park, J.-Y. Lee, and I. So Kweon, "CBAM: Convolutional block attention module," in *Proc. Eur. Conf. Comput. Vis. (ECCV)*, Sep. 2018, pp. 3–19.

- [48] Y. Zhu, Z. Yang, L. Wang, S. Zhao, X. Hu, and D. Tao, "Heterocenter loss for cross-modality person re-identification," *Neurocomputing*, vol. 386, pp. 97–109, Apr. 2020.
- [49] J. Duchi, E. Hazan, and Y. Singer, "Adaptive subgradient methods for online learning and stochastic optimization," J. Mach. Learn. Res., vol. 12, pp. 2121–2159, Feb. 2011. [Online]. Available: http://jmlr. org/papers/v12/duchi11a.html
- [50] M. A. Turk, S. D. Landes, M. K. Formica, and K. D. Goss, "Intellectual and developmental disability and COVID-19 case-fatality trends: TriNetX analysis," *Disability Health J.*, vol. 13, no. 3, Jul. 2020, Art. no. 100942.
- [51] S. L. Harrison, E. Fazio-Eynullayeva, D. A. Lane, P. Underhill, and G. Y. H. Lip, "Comorbidities associated with mortality in 31,461 adults with COVID-19 in the united states: A federated electronic medical record analysis," *PLOS Med.*, vol. 17, no. 9, Sep. 2020, Art. no. e1003321.
- [52] M. Denil, A. Demiraj, and N. de Freitas, "Extraction of salient sentences from labelled documents," 2014, arXiv:1412.6815. [Online]. Available: http://arxiv.org/abs/1412.6815
- [53] A. Shrikumar, P. Greenside, and A. Kundaje, "Learning important features through propagating activation differences," in *Proc. Int. Conf. Mach. Learn.*, 2017, pp. 3145–3153.
- [54] A. Shrikumar, P. Greenside, A. Shcherbina, and A. Kundaje, "Not just a black box: Learning important features through propagating activation differences," 2016, arXiv:1605.01713. [Online]. Available: http://arxiv.org/abs/1605.01713
- [55] J. Adebayo, J. Gilmer, M. Muelly, I. Goodfellow, M. Hardt, and B. Kim, "Sanity checks for saliency maps," 2018, arXiv:1810.03292. [Online]. Available: http://arxiv.org/abs/1810.03292
- [56] M. Infante *et al.*, "Differential diagnosis and management of focal ground-glass opacities," *Eur. Respiratory J.*, vol. 33, no. 4, pp. 821–827, Apr. 2009.
- [57] S. P. Georgiadou, N. V. Sipsas, E. M. Marom, and D. P. Kontoyiannis, "The diagnostic value of halo and reversed halo signs for invasive mold infections in compromised hosts," *Clin. Infectious Diseases*, vol. 52, no. 9, pp. 1144–1155, May 2011.
- [58] Z. Ye, Y. Zhang, Y. Wang, Z. Huang, and B. Song, "Chest CT manifestations of new coronavirus disease 2019 (COVID-19): A pictorial review," *Eur. Radiol.*, vol. 30, no. 8, pp. 4381–4389, Aug. 2020.
- [59] Q. Yu *et al.*, "Multicenter cohort study demonstrates more consolidation in upper lungs on initial ct increases the risk of adverse clinical outcome in COVID-19 patients," *Theranostics*, vol. 10, no. 12, p. 5641, 2020.
- [60] P. Spagnolo et al., "Pulmonary fibrosis secondary to COVID-19: A call to arms," *Lancet Respiratory Med.*, vol. 8, no. 8, pp. 750–752, Aug. 2020.



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