Classification of COVID-19 from tuberculosis and pneumonia using deep learning techniques

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Abstract

Deep learning provides the healthcare industry with the ability to analyse data at exceptional speeds without compromising on accuracy. These techniques are applicable to healthcare domain for accurate and timely prediction. Convolutional neural network is a class of deep learning methods which has become dominant in various computer vision tasks and is attracting interest across a variety of domains, including radiology. Lung diseases such as tuberculosis (TB), bacterial and viral pneumonias, and COVID-19 are not predicted accurately due to availability of very few samples for either of the lung diseases. The disease could be easily diagnosed using X-ray or CT scan images. But the number of images available for each of the disease is not as equally as other resulting in imbalance nature of input data. Conventional supervised machine learning methods do not achieve higher accuracy when trained using a lesser amount of COVID-19 data samples. Image data augmentation is a technique that can be used to artificially expand the size of a training dataset by creating modified versions of images in the dataset. Data augmentation helped reduce overfitting when training a deep neural network. The SMOTE (Synthetic Minority Oversampling Technique) algorithm is used for the purpose of balancing the classes. The novelty in this research work is to apply combined data augmentation and class balance techniques before classification of tuberculosis, pneumonia, and COVID-19. The classification accuracy obtained with the proposed multi-level classification after training the model is recorded as 97.4% for TB and pneumonia and 88% for bacterial, viral, and COVID-19 classifications. The proposed multi-level classification method produced is ~8 to ~10% improvement in classification accuracy when compared with the existing methods in this area of research. The results reveal the fact that the proposed system is scalable to growing medical data and classifies lung diseases and its sub-types in less time with higher accuracy.

Keywords Data augmentation · Normalization · Data balancing · Multi-level classification · Convolutional neural network · Feature selection

1 Introduction

Increasing medical abnormalities has led to life insecurity. Detection and treatment of such abnormalities at an early stage can help save lives. When highly dangerous pathogens like viruses reach the lungs, ground-glass opacity in the chest X-rays is observed due to fibrosis in the lungs. Due to the significant differences between X-ray images of an infected and non-infected person, artificial intelligence techniques can be used to identify the presence and severity of the infection. Authors propose a deep learning classification model using convolutional neural networks (CNN) that can analyse the chest X-rays and diagnose deadly lung diseases like pneumonia and tuberculosis and ultimately identify the victims of COVID-19. Motivation
behind this work involves two tasks. (1) Most classification methods tend to perform poorly on minority class samples [1]. This is because they aim to optimize the overall accuracy without considering the relative distribution of each class. Hence, abalanced dataset is important for creating a good prediction model. (2) Conventional supervised machine learning methods do not achieve higher accuracy when trained using lesser amount of COVID-19 data samples [2]. A deep neural network achieves improved accuracy compared to other conventional methods. The objectives of this research work are as follows: (1) to solve the class imbalance problem caused by minimal number of COVID-19 images by oversampling using SMOTE, (2) to increase the efficiency and accuracy of diagnosis using deep learning models to classify the chest X-ray images, (3) to help the physicians make chest X-ray diagnosis more accurately and avoid misdiagnosis of TB and pneumonia using computer-aided diagnosis, and (4) to implement multi-level classification, firstly to classify TB and pneumonia, and secondly to classify COVID-19 and bacterial and viral pneumonias.

The novelty of this research work is to perform multi-stage classification of chest X-ray images, to help the physicians make chest X-ray diagnosis more accurately and avoid misdiagnosis of TB and pneumonia using computer-aided diagnosis.

At the first stage, a binary classification of TB and pneumonia was performed. As COVID-19 is a type of pneumonia, it is further categorized as COVID-19 and viral and bacterial pneumonias in the second stage.

COVID-19-positive images are very few (minority class) compared to TB and pneumonia. The classification will be biased towards the class which has more number of images (called majority class). To deal with this problem, SMOTE technique was employed to oversample the minority class images and generate more images for COVID-19. This could avoid bias nature of learning algorithm towards the class having more images.

Deep learning architectures were applied to classify the type of chest disease more accurately and efficiently.

2 Related work

Chest X-ray classification using deep learning for automated COVID-19 screening [3] The authors have discussed a classification model that can analyse the chest X-rays and help in the accurate diagnosis of COVID-19. The methodology classifies the chest X-rays into 4 classes normal, pneumonia, tuberculosis (TB), and COVID-19. Furthermore, the X-rays indicating COVID-19 are classified on severity basis into mild, medium, and severe. The deep learning model used for the classification of pneumonia, TB, and normal is VGG16. For the segregation of normal pneumonia and COVID-19, the DenseNet-161 was used. ResNet-18 worked best for severity classification. Their approach allows mass screening of the people using X-rays as a primary validation for COVID-19 [3–5].

The input chest X-ray image is passed through a VGG-16 model and labeled as normal, tuberculosis, or pneumonia. Furthermore, the pneumonia images are passed through a DenseNet-161 model and categorized as normal pneumonia or COVID-19. The COVID-19 images are passed through a ResNet18 model and classified as severe, medium, or mild [6, 7].

The input size of the image for the VGG-16 model is $64 \times 64 \times 1$. After this, the image passes through a 2D convolution layer with dimensions $224 \times 224$. Then, it passes through the next layer with dimensions $112 \times 112$. After this, it passes to consecutive convolution layers along with MaxPooling with dimension changes $56 \times 56$, $28 \times 28$, and $14 \times 14$. Then, it goes to a MaxPooling layer with dimension change $7 \times 7$. After a series of $2500$, $4096$, and $4096$ fully connected layers, the X-rays are classified into normal X-ray, tuberculosis-affected X-ray, and pneumonia-affected X-Ray [8].

The input size of the image for the DenseNet-161 model is $64 \times 64 \times 1$. After which, the image passes through a 2D convolutional layer with dimensions $112 \times 112$. Then, it passes through the next layer with dimensions $56 \times 56$. After passing through the consecutive layers, the dimensions change from $56 \times 56$ to $28 \times 28$, $14 \times 14$, and finally, $7 \times 7$. After 1000 fully connected layers, the X-rays are classified into normal pneumonia and COVID-19.

The input size of the image for the ResNet-18 model is $64 \times 64 \times 1$. After this, the image passes through a 2D convolution layer with dimensions $112 \times 112 \times 64$. Then, it passes through the next layer with dimensions $56 \times 56 \times 64$. After this, it passes to consecutive 2D convolution layers with dimension changes $28 \times 28 \times 128$, $14 \times 14 \times 256$, and $7 \times 7 \times 512$. After 1000 fully connected layers, the X-rays are classified into severe, medium, and mild [9, 10].

A multi-level classification model was introduced in previous work [3]. They did separate models for separate classification tasks, so that the misclassification of the lung diseases tuberculosis and pneumonia can be eliminated first and pneumonia-classified patients only can be concentrated for further classifications. So, the multi-level classification model was proposed for the classification of lung diseases.

Pneumonia diagnosis using deep learning [11] The authors have discussed about classification of chest X-ray images to bacterial pneumonia, viral pneumonia, and normal using the VGG pretrained model which has 16 convolution layers followed by a connected hidden layer and dropout layers in between. Dropout layers help to regularize the model to
prevent the model from overfitting. A dataset has more than nine thousand chest X-ray images for the classes bacterial pneumonia, viral pneumonia, and normal chest X-rays. This particular model which has CNN layers is able to perform better mainly due to the deep structure of CNN that uses the power of extracting different level features, which resulted in a better generalization capability.

**Cascaded deep learning classifiers for computer-aided diagnosis of COVID-19 and pneumonia diseases in X-ray scans** [12] This study includes eleven pretrained convolutional neural network models, such as Visual Geometry Group (VGG) network and residual neural network (ResNet). They have been successfully tested and evaluated on a public X-ray image dataset for normal and three diseases cases. The results of proposed cascaded classifiers showed that VGG16, ResNet50V2, and dense neural network (DenseNet169) models achieved the best detection accuracy of COVID-19, viral (non-COVID-19) pneumonia, and bacterial pneumonia images, respectively [12].

**Viral and bacterial pneumonia detection using artificial intelligence in the era of COVID-19** [13] The authors have discussed a classification model to classify COVID-19, non-COVID, bacterial pneumonia, viral pneumonia, and normal. The paper proposed that pretrained AlexNet model is employed due to its high accuracy in carrying out feature extraction and image classification. With regard to the classification of COVID-19 and normal CXR images, it was observed that the model provided significantly better performance compared to studies that utilized small amounts of dataset such as Mahmud et al. (2020) and models developed from scratch.

**A deep learning approach to detect COVID-19 coronavirus with X-ray images** [14] The authors have discussed about classifying chest X-ray images into viral pneumonia, bacterial pneumonia, and normal cases using ResNet50 deep network architecture at the first stage, and, furthermore, viral pneumonia samples were classified into COVID-19 induced pneumonia and other viral pneumonias with ResNet101 architecture. The limited number of COVID-19 images makes it a bit difficult to train the deep learning models from scratch, which is overcome by using deep transfer learning methods in the current study.

### 3 Proposed multi-level classification model for classification of lung diseases

The proposed system has two models (multi-level classification): first model for classifying tuberculosis (TB) and pneumonia and the second model for classifying the types of pneumonia such as bacterial pneumonia, viral pneumonia, and COVID-19 from taking the pneumonia-classified X-ray images (output of the first model) as depicted in Fig. 1.

Since the dataset for the TB, pneumonia, and pneumonia types (first, second levels) are less when compared with datasets for other classification tasks, to improve the classification performance of the model, data augmentation and oversampling techniques are applied to the training set. In the dataset considered under current work [15–17], the data is biased for the pneumonia class; to solve this bias problem, authors have compared with class balancing techniques and have found that SMOTE performs well for class balancing, so authors have used SMOTE to overcome the problem from our dataset. For the accurate classification of lung diseases, authors have proposed this method of multi-level classification.

- In first level classification — classification of images into pneumonia and tuberculosis.
- In second level classification — further classification into viral and bacterial pneumonias and COVID-19.

This is the novel idea authors have proposed in this model.

Data augmentation techniques such as image shifts — *width_shift_range* and *height_shift_range*; image flips — *horizontal_flip* and *vertical_flip*; image rotations — *rotation_range*; and image zoom — *zoom_range* are done for the training set for both the models [18, 19]. The data augmentation and data splitting are depicted in Fig. 2.

Since the dataset is unbalanced, some classes are having a smaller number of images compared to other classes. For balancing the dataset, oversampling techniques such as SMOTE and Borderline-SMOTE SVM (SVM SMOTE) are used for increasing the number of samples in the minority classes [20–24].

After balancing the dataset, a convolutional neural network model was built and trained for the pneumonia and tuberculosis datasets. The model was tested against a test set. The images which are classified as “pneumonia” were saved and are to be used as a test set for the second model.

Another convolutional neural network was built and trained using the bacterial pneumonia, viral pneumonia, and COVID-19 for the second-level classification. And this model was tested by the pneumonia-classified images (from the first model). The proposed system is shown in Fig. 3.

#### 3.1 SMOTE

Synthetic Minority Oversampling Technique (SMOTE) is an oversampling approach in which the minority class is oversampled by creating “synthetic” examples rather than by oversampling with replacement [25, 26].
Algorithm

1. Setting the minority class set A, for each \( x \) in \( A \), the k-nearest neighbours of \( x \) are obtained by calculating the Euclidean distance between \( x \) and every other sample in set \( A \).
2. The sampling rate \( N \) is set according to the imbalanced proportion. For each \( x \) in \( A \), \( N \) examples (i.e., \( x_1, x_2, \ldots, x_N \)) are randomly selected from its k-nearest neighbours and construct the set \( A_1 \).

3. For each example \( x_k \) in \( A_1 \) (\( k = 1, 2, 3 \ldots N \)), the following formula is used to generate a new example: \( x' = x + \text{rand}(0,1) \times |x - x_K| \) in which \( \text{rand}(0,1) \) represents the random number between 0 and 1.

Often, real-world datasets are predominantly composed of “normal” examples with only a small percentage of “abnormal” or “interesting” examples. With SMOTE, one can determine the ratio of the oversampling or specifies the class to be oversampled. We can also control the
randomization of the algorithm by specifying it in the function’s argument. Other optional parameters like number of nearest neighbours can be specified. SMOTE synthesizes new minority instances between existing minority instances. SMOTE algorithm is performed here which oversamples the specified class and produces synthetic samples. Sampling strategy is specified which determines the ratio of the oversampling or specifies the class to be oversampled. The randomization of the algorithm can be controlled by specifying it in the function’s argument.

The undersampling will reduce the number of images which would affect the model accuracy.

Bootstrapping would select samples in random which would result in probability of selecting the samples from the class where many images will be high compared to selecting images from class with very few images.

It is necessary to deal with the class which has very few images and remove the bias towards classes during training the model.

Other optional parameters like number of nearest neighbours can be specified. SMOTE synthesizes new minority instances between existing minority instances [27]. It generates the virtual training records by linear interpolation for the minority class. Data balancing using SMOTE is depicted in Fig. 4. Table 1 shows the class distribution of images for tuberculosis and pneumonia.

3.2 Borderline-SMOTE SVM or SVM SMOTE

Borderline-SMOTE SVM is a variant of SMOTE. In this SMOTE type, Borderline-SMOTE, support vector machine (SVM) algorithm is used instead of a KNN to identify misclassified examples on the decision boundary. An SVM is used to locate the decision boundary defined by the support vectors and examples in the minority class that close to the support vectors becomes the focus for generating synthetic examples.

The main difference between SVM-SMOTE and the other SMOTE is that instead of using K-nearest neighbours to identify the misclassification in the Borderline-SMOTE, the technique would incorporate the SVM algorithm [21].

In the SVM-SMOTE, the borderline area is approximated by the support vectors after training SVMs classifier on the original training set. Synthetic data will be randomly created along the lines joining each minority class support vector with a number of its nearest neighbours [22].

In addition to using an SVM, the technique attempts to select regions where there are fewer examples of the minority class and tries to extrapolate towards the class boundary. This variation can be implemented via the SVM SMOTE class from the imbalanced-learn library. Table 2 shows the class distribution of types of pneumonia.

3.3 Construction of DNN model

A convolutional deep neural network model is built in which we have used 1 input layer, 1 output layer with Softmax activation layer, and 5 hidden layers adding totally to 7 layers. The hidden layers are:

- Conv2D layer with ReLu activation layer
Table 1 Class distribution of TB and pneumonia dataset

| Class label | No. of samples before applying SMOTE | No. of samples after applying SMOTE |
|-------------|-------------------------------------|------------------------------------|
| TB          | 2520                                | 3780                               |
| Pneumonia   | 3780                                | 3780                               |

Table 2 Class distribution of types of pneumonia dataset

| Class label            | No. of samples before applying SMOTE | No. of samples after applying SMOTE |
|------------------------|-------------------------------------|------------------------------------|
| Bacterial pneumonia    | 2469                                | 2469                               |
| Viral pneumonia         | 1332                                | 2469                               |
| COVID-19                | 1013                                | 2469                               |

The number of neurons used in the input layer and the output layer and the number of hidden layers deployed in different datasets are tabulated correspondingly in Table 4.

4 Materials

The datasets, namely the TB and pneumonia dataset, and the types of pneumonia dataset are used for the implementation. In the class distribution, the number of features in each
The dataset is tabulated in Table 5. From Table 5, it can be seen that the first dataset has a moderate class imbalance and the second dataset has a high class imbalance. The number of sample and class distribution of images is tabulated in Table 5. The chest X-ray images were extracted from Kaggle database, chest X-ray images (pneumonia) [1], tuberculosis (TB) chest X-ray database [28], and curated chest X-ray image dataset for COVID-19 [2].

Table 5 Dataset description

| Dataset                          | No. of samples | No. of classes | Class distribution             |
|----------------------------------|----------------|----------------|--------------------------------|
| TB and pneumonia dataset (2 classes) | 8755           | 2              | TB — 3500                       |
|                                  |                |                | Pneumonia — 5255               |
| Types of pneumonia dataset (3 classes) | 5938           | 3              | Bacterial — 3001               |
|                                  |                |                | Viral — 1656                    |
|                                  |                |                | COVID — 1281                   |

(i) Tuberculosis and pneumonia classification

The dataset was split into 70% for training and 30% for testing. After applying image augmentation and oversampling techniques (SMOTE), a convolutional neural network was built and trained for this dataset and achieved 97.4% accuracy.

(ii) Bacterial pneumonia, viral pneumonia, and COVID-19 classification

The dataset was split into 70% for training and 30% for testing. After applying image augmentation and oversampling techniques (SMOTE), a convolutional neural network was built and trained for this dataset and achieved 88% accuracy. The hyperparameter tuning is mentioned as follows.

Initial learning rate: 0.1

Optimizers used are as follows:
At the hidden layers: ReLu and Softmax
At the output layer: Adam
Loss function: Categorical cross-entropy
Number of epochs: 50 to 200

After applying oversampling techniques, misclassification was reduced and the accuracy was improved for both the classification models. The performance metrics of both levels of classification is tabulated in Tables 6 and 7.

The increase in images will improve accuracy of the model as the model gets trained with more images. But class imbalance will overfit the model to the class which has more number of images. To overcome these bias, SMOTE is applied to balance the number of images in each class so that the bias is removed during classification.

Table 6 shows 3% increase in performance for the first level classification of TB and pneumonia.

Table 7 shows 10 to 15% increase in performance for the second level of classification between viral and bacterial pneumonias and COVID-19.

The TB and pneumonia classification model (first model) was tested against 1646 images. Out of 1646 images, 988 images were classified as “pneumonia”, 658 images were classified as “tuberculosis.” The images that are classified as “pneumonia” from the first model were used as the testing set for the second model (types of pneumonia classification). Out of 988 images, the model classified 541 images as “bacterial pneumonia” and 175 images as “viral pneumonia” and 272 images as “COVID-19.”
Data augmentation and SMOTE techniques have improved the multi-class classification accuracy. Prior to data balancing, the accuracy obtained was 76.3% and post-data balancing the accuracy was 87.1%. This accuracy is with respect to the three-class classification of COVID-19 and bacterial and viral pneumonias. Nevertheless, the binary classification of these three types of chest diseases yielded classification accuracy ~95 to ~98% which was detailed in Section 6 in Table 9.

5.1 Comparison of existing models vs proposed model

The efficiency of existing models was compared with our proposed model’s efficiency in terms of the performance metric accuracy in percentage.

From the above Table 8, it is evident that our proposed model has higher efficiency of 8–11% when compared to the performance of existing models. The framework mentioned in the paper [31] was designed, trained, and validated to identify four classes of CXR images, namely healthy, bacterial pneumonia, viral pneumonia, and COVID-19. The experimental results pose the proposed framework as a potential candidate for COVID-19 disease diagnosis using CXR images, with training, validation, and testing accuracy of 92.4%, 88.24%, and 87.13%, respectively, in four-class classification.

6 Discussions

The other methods to improve classification accuracy are feature selection and autoencoders which are discussed below.

6.1 Feature selection

Irrelevant or partially relevant features can negatively impact model performance. Feature selection is the process where you automatically or manually select those features which contribute most to your prediction variable or output in which you are interested in. Having irrelevant features in your data can decrease the accuracy of the models and make your model learn based on irrelevant features. Some of the advantages of feature selection are,

- Reduces overfitting: Less redundant data means less opportunity to make decisions based on noise.
- Improves accuracy: Less misleading data means modeling accuracy improves.
- Reduces training time: Fewer data points reduce algorithm complexity and algorithms train faster.

6.2 Autoencoders

The current research presents a computer-aided detection model utilizing chest X-ray images for combating the pandemic. The research work in [32] uses features extracted from pretrained networks along with sparse autoencoder for dimensionality reduction and a feed forward neural network (FFNN) for the detection of COVID-19. Autoencoders are proposed as a solution for feature selection. DL techniques used in COVID-19 have also been categorized into seven main distinct categories as long short-term memory networks (LSTM), self-organizing maps (SOMs), conventional neural networks (CNNs), generative adversarial networks (GANs), recurrent neural networks (RNNs), autoencoders, and hybrid approaches [33, 34]. The research work in [35] introduces a novel unsupervised DL-based variational autoencoder (UDL-VAE) model for COVID-19 detection and classification. Autoencoders are used for feature selection to uncover existing nonlinear relationships between

| Metrics       | Before balancing using SMOTE | After balancing using SMOTE |
|---------------|-----------------------------|----------------------------|
| Accuracy (%)  | 76.3                        | 87.1                       |
| Precision (%) | 80.6                        | 89.3                       |
| Recall (%)    | 73.9                        | 88                         |
| Specificity (%) | 75.4                      | 87.4                       |
| F1-Score (%)  | 74.7                        | 88.5                       |

| Research method               | Second level of classification (COVID diagnosis) accuracy (in %) |
|------------------------------|---------------------------------------------------------------|
| Existing work 1 [29]         | 80                                                           |
| Existing work 2 [30]         | 78.9                                                         |
| Existing work 3 [31]         | 87                                                           |
| Our proposed model           | 88                                                           |

| Chest disease type              | Second level of classification (COVID diagnosis) accuracy (in %) |
|---------------------------------|---------------------------------------------------------------|
| Viral pneumonia vs COVID-19     | 95                                                           |
| Bacterial pneumonia vs COVID-19 | 98                                                           |
| Bacterial pneumonia vs viral pneumonia | 97                                                             |
features. Autoencoders are unsupervised learners that can learn to compress and encode data and then learn how to reconstruct the data back from the reduced encoded representation. The resulting representation is as close as possible to the original input. In other words, we are taking the feature space and reducing its dimensionality and then reconstructing it from its reduced format. The principle of autoencoders helps us by reducing the data dimensions, learning how to ignore the noise in them, and moreover, knowing which features can best help us reconstruct the data. So, after reconstructing the data, first use the weight matrix of the autoencoder that connects the input feature layer to the reduced layer:

- If a feature contributes little to the representation of others, it means that the corresponding weight squared ($w^2$) is close to zero.
- On the other hand, if the feature plays an important role in the representation of other features, the corresponding weight must be significant.

One of the main drawbacks of this method arises from the simplicity of the autoencoder, where a simple single-layer autoencoder cannot model complex non-linear feature dependencies. Feature selection was tried before a multi-level classification module for experimental purposes just to compare the results and see if the efficiency (accuracy) of the model gets better. But feature selection did not work well for our model. The accuracy was achieved only about 77% after applying the feature selection step, whereas without feature selection, our model achieved a higher accuracy of 87%.

Furthermore, the binary classification on two sets of disease types was performed and the results obtained are tabulated in Table 9.

As COVID-19 is a type of viral pneumonia, the accurate categorization was bit low as 95% while bacterial pneumonia vs COVID-19 yielded 98% and bacterial vs viral pneumonia yielded 97%.

7 Conclusion

In our proposed model, the class imbalance problem was handled with the help of SMOTE and SVM-SMOTE, an oversampling algorithm to even out the class distribution. SMOTE performance was evaluated on two datasets: (1) TB and pneumonia multi-class dataset which was moderately imbalanced but highly dimensional and (2) types of pneumonia multi-class dataset which was highly imbalanced. The evaluated results after applying SMOTE were compared with results obtained from before and after applying SMOTE using deep neural network as the classifier. The comparison showed that SMOTE performs significantly better using imbalanced data as it is as there was a 10% increase in the accuracy of the model. The algorithm effectively handled the class imbalance problem. The explanation for this improvement in performance relates to the ability of SVM-SMOTE to generate artificial data in safe areas of the input space, while, at the same time, aggressively increasing the diversity of the generated instances. SVM-SMOTE can be a useful tool for researchers and practitioners since it results in the generation of high-quality artificial data and only requires the tuning of a small number of parameters.

Deep learning can be a very useful tool in the medical industry for the detection of diseases just by processing the images of the chest X-rays and feeding it as an input to the model. The chest X-rays are classified into different classes such as pneumonia, tuberculosis, and COVID-19. Mass screening of people for detection of COVID-19 can be done effectively with our proposed model. It will help yield faster and accurate results and will be cost-effective as compared to the conventional RT-PCR method. This approach can be implemented at the local level and in the rural areas where adequate facilities are absent. The limitation of the current work is that the features used for classifying the chest diseases were not investigated. The feature engineering has to be performed for obtained in the biological feature relevance for identifying the type of chest disease.

Future research directions will include the exploration of image data augmentation techniques to improve accuracy even more, while avoiding overfitting. To expand the size of the middle convolutional layers and making the stride and filter size on the first layer smaller. Use GoogLeNet to reduce the number of parameters in the network. Add more data augmentation techniques to increase the efficiency of training. Design an online application to diagnose pneumonia in chest X-ray images, to serve the largest possible number of slums who lack health care and could not afford for the diagnosis.

Declarations

Ethical approval This article does not contain any studies with human participants or animals performed by any of the authors.

Informed consent Informed consent is not necessary as this article does not involve human or animal participants.
Conflict of interest The authors declare no competing interests.

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