



Construction of a radiomics-based model for predicting the efficacy of radiotherapy and chemotherapy for non-small cell lung cancer

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ABSTRACT

Objective: Pre-treatment enhanced CT image data were used to train and build models to predict the efficacy of non-small cell lung cancer after conventional radiotherapy and chemotherapy using two classification algorithms, Logistic Regression (LR) and Gaussian Naive Baye (GNB).

Methods: In this study, we used pre-treatment enhanced CT image data for region of interest (ROI) sketching and feature extraction. We utilized the least absolute shrinkage and selection operator (LASSO) mutual confidence method for feature screening. We pre-screened logistic regression (LR) and Gaussian naive Bayes (GNB) classification algorithms and trained and modeled the screened features. We plotted 5-fold and 10-fold cross-validated receiver operating characteristic (ROC) curves to calculate the area under the curve (AUC). We performed DeLong's test for validation and plotted calibration curves and decision curves to assess model performance.

Results: A total of 102 patients were included in this study, and after a comparative analysis of the two models, LR had only slightly lower specificity than GNB, and higher sensitivity, accuracy, AUC value, precision, and F1 value than GNB (training set accuracy: 0.787, AUC value: 0.851; test set accuracy: 0.772, AUC value: 0.849), and the LR model has better performance in both the decision curve and the calibration curve.

Conclusion: CT can be used for efficacy prediction after radiotherapy and chemotherapy in NSCLC patients. LR is more suitable for predicting whether NSCLC prognosis is in remission without considering the computing speed.

1. Introduction

According to data published in the Global Cancer Statistics report, lung cancer is currently the malignancy with the highest risk of morbidity and mortality [1]; 85 % of them are non-small cell lung cancer (NSCLC), mainly adenocarcinoma and squamous carcinoma [2]. The National Comprehensive Cancer Network (NCCN) practice guideline 2022.5 for NSCLC recommends that the primary treatment options for patients with NSCLC include surgery, radiotherapy therapy, chemotherapy, molecular targeted therapy, and immunotherapy [3], and the current treatment modality for all patients with NSCLC for whom radical surgery is not preferred is often a combination of radiation therapy and chemotherapy. However, in clinical practice, due to the insidious nature of NSCLC, most patients have missed the opportunity of radical surgery at the time of diagnosis, or are unable to undergo surgery due to the location of the tumor close to the vital organs or main bronchus, so a large proportion of patients take combined radiotherapy and chemotherapy.

In clinical practice, even at the same disease stage, heterogeneity exists in different individuals, and there is also heterogeneity within the tumor, which does not manifest as homogeneous. Current diagnostic modalities, such as pathological diagnosis and imaging

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diagnosis, have difficulty in reflecting the heterogeneity within the tumor that is not visible to the naked eye, so the treatment outcome after radiotherapy and chemotherapy may also vary considerably [4]. Suppose reliable efficacy predictors are available to classify patients before treatment or early in the disease. In that case, they can help physicians develop individualized treatment plans, provide a basis for decision-making on drug therapy, adjust the dose of radiotherapy and chemotherapy regimens, or extend the duration of treatment appropriately, which can be used to improve patient prognosis [5].

Radiomics mines the vast amount of information inherent in CT images and performs quantitative analysis to explore potential diagnostic, therapeutic, and prognostic applications, which have the advantages of non-invasiveness, reproducibility, convenience, and ease of operation occupy an important position in precision medicine. Currently, with the continuous development and improvement of chest Computed Tomography (CT), CT is the most commonly used noninvasive examination modality in lung cancer diagnosis and efficacy assessment, and CT has the advantages of a lower price, fewer scanning parameters, more stable image quality, and more standard image data [6]. Therefore, this study was conducted to predict and validate the prognosis of NSCLC patients after conventional radiotherapy and chemotherapy by radiomics based on the enhanced CT images before treatment.

2. Materials and methods

This study was approved by the Ethics Review Committee of the Hospital of Chuanbei Medical College, File No. 2022ER581-1, and the patient gave consent for his images and data to be used for the experiment and study. The study was conducted in accordance with established ethical guidelines and complied with all regulations.

3. Data and methods

3.1. Study population

Inclusion criteria: ① pathological histological puncture biopsy confirmed the diagnosis of NSCLC (including squamous cell carcinoma, adenocarcinoma, and other types); ② stage II and III patients who were inoperable; ③ stage IV patients without multiple metastases and distant organ metastases; ④ receiving radical radiotherapy and 4–6 cycles of platinum-containing chemotherapy; ⑤ patients in good general condition with a Karnofsky (KPS) score ≥ 70 scores (Table 1).

Exclusion criteria: (i) patients with incomplete clinical or image information of low quality; (ii) patients without efficacy assessment information after treatment; (iii) those with other primary tumors; (iv) those who did not follow up regularly according to the follow-up requirements.

The data of patients diagnosed with NSCLC in our hospital between December 2018 and July 2022 and meeting the above criteria were retrospectively analyzed, and 102 cases were finally included.

3.1.1. Radiotherapy protocols

The total dose was 40–60 Gy, with single irradiation of 2.0 Gy, once a day, five days a week treatment.

3.1.2. Chemotherapy regimen

Four to six treatment cycles of platinum-based chemotherapy drugs were used. The drug doses were calculated based on the patient's body surface area, dosing regimen, and liver and kidney function.

Table 1
Karnofsky (kps, percentage method) functional status rating scale.

Physical status	Score*
Normal, no signs and symptoms	100
Able to perform normal activities with minor signs and symptoms	90
Barely able to perform normal activities with some signs and symptoms	80
Self-care, but unable to maintain normal life and work	70
Mostly self-care, but occasionally needs help	60
Needs frequent care	50
Cannot take care of himself/herself, needs special care and assistance.	40
Severely unable to care for self	30
Severely ill, requiring hospitalization and active, supportive care	20
Critically ill, near death	10
Death	0

* : The higher the score, the better the health status, the more tolerable the side effects of the treatment, and therefore the more likely it is that the treatment will be complete. The lower the score, the worse the health status; if the score is below 60, many effective anti-tumor treatments cannot be implemented.

3.2. Efficacy assessment and grouping

3.2.1. Efficacy assessment

All cases were rated according to the WHO Response Evaluation Criteria In Solid Tumors [7] (RECIST1.1) as complete response (CR), partial response (PR), stable disease (SD), and progressive disease (PD).

3.2.2. Case grouping

Patients were divided into the remission group (including CR and PR) and the non-remission group (including PD and SD).

3.3. Image Acquisition and processing

3.3.1. Acquisition of images

In order to ensure the timeliness of obtaining image information that can reflect the closest treatment phase, all images were obtained from the radiotherapy center's images of the radiation target area plan developed within 1–3 days prior to radiotherapy. For patients who could cooperate with breathing instructions, we instructed patients to hold their breath during the scanning period to minimize the impact of respiratory motion. Specifically, patients held their breath at maximal inspiration upon hearing the breath-hold command from the CT technologist prior to initiating non-table motion scanning. All patients were trained to familiarize themselves with the breath-holding process before the scan. For patients with potential inadequate breath-hold times, scanning was performed in segments. Depending on patient positioning, the scanning direction was adjusted to minimize respiratory motion of the tumor. Scanning equipment: CT (Siemens Emotion 16-slice configuration) Scanning parameters: Tube Voltage: 120 kV; Tube Current: 200 mA; Layer Thickness: 3 mm; Pitch: 0.625 mm; FOV = 700 mm × 700 mm. In this study, we used an iodixanol contrast agent manufactured by Yangzijiang Pharmaceutical Group Co., Ltd. with a specification of 74.1 g of iodine in 100 ml, i.e., 350 mg of iodine per 100 ml.

3.3.2. Image preprocessing

in order to provide a high level of detail, we resample the image using a 3d slicer, the resampled voxel size is set to $1 \times 1 \times 1$ mm. reconstruction matrix size: 64×256 , reconstruction kernel size: $2 \times 3 \times 4$.

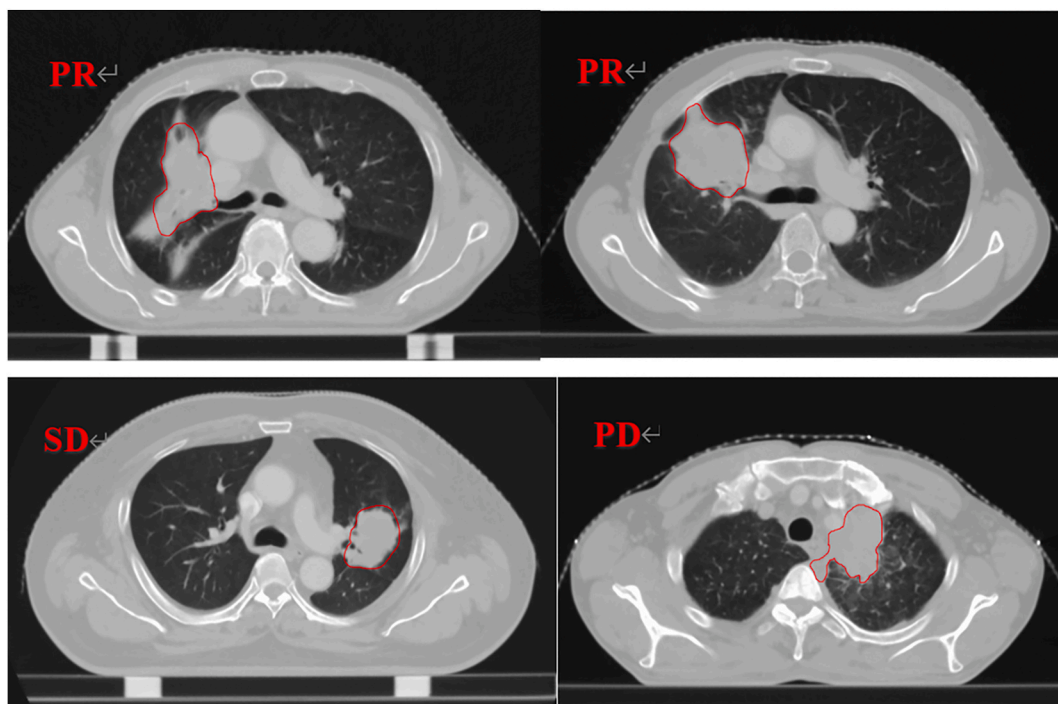


Fig. 1. Illustration of ROI Contour

PR, SD and PD respectively represent three types of therapeutic effects after evaluation based on RECIST 1.1 criteria; the contour in the figure represents GTV, where A corresponds to PR (Partial Response), B corresponds to SD (Stable Disease), and C corresponds to PD (Progressive Disease) in terms of total tumor volume. ROI refers to Region of Interest.

3.3.3. Image segmentation

Export the images in DICOM format and upload them to the Monaco system for image segmentation. When uploading the images, the images were renamed to achieve the effect of removing the blended names. In this study, the Gross Tumor Volume (GTV) was manually segmented layer by layer using 3D Slicer software by two radiation therapists with more than five years of experience in thoracic tumor therapy. After the initial segmentation, the results were reviewed and modified by a radiation therapist with more than ten years of experience in thoracic tumor radiotherapy. The whole segmentation process strictly followed the recommendations of the National Comprehensive Cancer Network (NCCN) [8] Guidelines for the Treatment of Non-Small Cell Lung Cancer (NCCN Guidelines Version 3.2022) (Fig. 1). [9].

3.4. Feature extraction

The image and the outlined ROI structure are imported into 3D Slicer (4.11) for feature extraction. The features extracted by 3D Slicer are mainly performed through the open-source plug-in (Slicer Radiomics). The extracted features include the mean, minimum, maximum, standard deviation, skewness, kurtosis, etc., of the first-order statistical features of the original image; the surface area, volume, surface area to volume ratio, sphericity, compactness, and 3D diameter of the morphological features and five types of texture features [10]: Gray Level Cooccurrence Matrix (GLCM), Gray Level Run Length Matrix (GLRLM), Gray Level Size Zone Matrix (GLSZM), Neighboring Gray Tone Difference Matrix (NGTDM), Gray Level Dependence Matrix (GLDM); and the features after filtering and transforming the images using Laplacian of Gaussian (LoG) with LoG-sigma sizes of 4.0 mm, 5.0 mm, and 6.0 mm, respectively, and the eight filter functions of LLL, LLH, LHL, LHH, HLL, HHL, HLH, HHH, and HHH in the wavelet transform for a total of 11 filtering methods, each first-order, second-order and texture features mentioned above after the transformation [11]. In this case, the wavelet transform image was created using the default parameters of 3D Slicer. The wavelet sequence used was Daubechies (Db4). A 3-layer decomposition was performed. The fill mode was symmetric filling. Quantization of 64 bins was applied. The sampling mode was custom sampling, with specific sampling densities of x-, y- and z-axis sampling intervals of 4, 5, and 6 voxels, respectively.

3.5. Model training and validation

3.5.1. Screening of features

Radiomic features were extracted using the Python module in 3D Slicer 4.10.2 with Anaconda Python 3.7. Downstream data processing and analysis were performed in R 4.2.2. Patients were categorized into mitigation and non-mitigation groups, labeled as 0 and 1 respectively. The data were then normalized and standardized. Features with confidence coefficients greater than 0.1 were filtered out using the confidence method. The mutual information coefficient threshold was set to 0.1 for relatively loose initial filtering, which retained more potential predictive features. This prepared for the next stage of more stringent LASSO regression selection. Finally, useful features were filtered again using LASSO. Next, the data were randomized and randomly split into training (70 %) and test sets (30 %).

3.5.2. Model selection and training

The training set data were input into two classification algorithms - Logistic Regression (LR), Gaussian Naive Bayes (GNB), Support Vector Machine (SVM) and K-Nearest Neighbors (KNN). The above models are commonly used in the binary classification problem of lung cancer and have been supported by studies to have better prediction effects [12–14]. The training set was further randomly divided into a training subset (70 %) and validation subset (30 %). The training subset data were fed into models for training. During the model training process, the training set data are randomly divided into a training set and validation set [15], brought into the model for training and adjusting the model, and then substituted into the test set data and plotted the training curve together. Pre-screening based on AUC results to exclude SVM and KNN models. In this study, we set the parameters of LR and GNB models as follows. For the LR model, we use LogisticRegression in sklearn, and the main parameters are: penalty = 'l1'; C = 0.1; solver = 'liblinear'; max_iter = 1000. For the GNB model, we use GaussianNB in sklearn, and the main parameters are: var_smoothing = 1e-9; priors are empty; fit_prior = False. The model training dynamics are observed for overfitting or underfitting. The training curve is used to select the best training depth of the model, i.e., the best model for that data for that algorithm. After the model is trained and brought into the test set data, the corresponding evaluation index values are generated by (i) bringing in the data and then calculating the average value after getting some evaluation index values, such as the accuracy of the model by ten times five-fold cross-validation, (i) bringing in the data and then calculating the average value after getting some evaluation index values, such as the accuracy of the model by ten times five-fold cross-validation. In this study, 5-fold cross-validation was performed 10 times to evaluate model performance. In each cross-validation iteration, the training data was randomly split into 5 equal folds. 4 folds were used for model training and the remaining fold was used for validation. This process was repeated 10 times so that each fold served as the validation set once. The cross-validation accuracy was recorded for each iteration. The final reported cross-validation accuracy is the average of the $5 \times 10 = 50$ accuracy values obtained during cross-validation.; (ii) building the confusion matrix and the classification report of the model with the model prediction results to evaluate the model; (iii) then plotting the ROC curve with each model training and prediction results and calculating the AUC values [16]. We adhered to the Image Biomarker Standardization Initiative (IBSI) [17] guidelines during image processing procedures to ensure standardization.

3.5.3. Validation of the model

For the LR and GNB models constructed in this paper, the calibration degree of model classification was assessed by calibration

curves on the training set and test set, respectively. The calibration curves indicate the extent to which the predicted probabilities match the actual observed probabilities. Decision curves were plotted to assess the performance of model classification on the training set and test set, respectively. The threshold was set to 0.5, reflecting the model's false positive rate versus the true positive rate, and thus indicating the net benefit rate.

3.6. Statistical analysis

The *t*-test was used for the measurement data; the χ^2 test or Fisher's exact test was used for the one-way analysis of the count data. The test levels $\alpha = 0.05$ and $P < 0.05$ were considered statistically significant differences.

4. Results

4.1. General information of patients

Following the flow of Fig. 2, a total of 102 patients were included in this study, according to the clinical efficacy evaluation index of solid tumors, including 67 cases of PR, 33 cases of SD, 2 cases of PD, 67 cases in the remission group, and 35 cases in the non-remission group. The overall mean age was (62.18 ± 8.50) years, the mean age of the influential group was (62.46 ± 8.47) years, and the mean age of the non-remission group was (61.63 ± 8.57) years.

4.2. Statistical analysis of patients' clinical data and efficacy

The efficacy of 102 patients was evaluated according to the RECIST 1.1 evaluation index. The results showed that the current remission rate of patients who received 4–6 cycles of chemotherapy and radical radiotherapy was 65.69 % (67/102). The general clinical data of the patients are described in Table 2; the measurement data are expressed as "x \pm s," and the counting data are expressed as [n(%)]. Except for the age of the patients, the pairwise results between the groups showed ($p > 0.05$), demonstrating no significant difference between the two groups of patients, which was beneficial for the following analysis.

4.3. Results of radiomics feature screening

The general clinical data were converted into digital form and normalized, and after standardization, the features were screened together with radiomics features. The features with confidence coefficients greater than 0.1 were screened by the confidence method, and then eight useful features were finally screened by LASSO, at which time the best $\lambda = 0.0693$ (as in Fig. 3). It can be seen that all are filtered features: two features after filtering with an LoG-sigma kernel size of 4.0 mm, one feature after filtering with 6.0 mm, and the filtered features of LLH, HLH, and HHH filter functions in wavelet transform. Among the five selected features, only the features of the LLH filter function in wavelet transform are first-order features. The rest are texture features, of which three texture features of the wavelet-HHH filter function in wavelet transform are selected, and the specific features are named in Table 3.

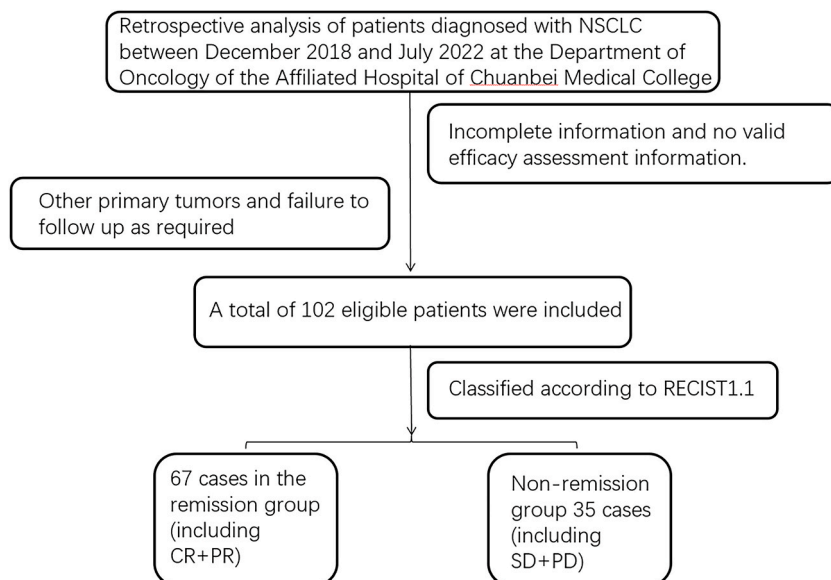


Fig. 2. Inclusion of exclusion flow chart CR,complete response; PR,partial response; SD,stable disease; PD,progressive disease; NSCLC, non-small cell lung cancer.

Table 2
General clinical data non-remission group remission group and non-remission group description.

Group	remission group	non-remission group	P
Quantity	67(65.69 %)	35(34.31 %)	
Age (mean ± standard deviation)	62.46 ± 8.47	61.63 ± 8.57	0.2961
Gender	60(89.55 %)		0.0062
Man	7(10.45 %)	23(65.71 %)	
Woman		12(34.29 %)	
Smoke	48(71.64 %)		0.0515
Yes	19(28.36 %)	18(51.43 %)	
No		17(48.57 %)	
History of chronic lung disease	8(11.94 %)		>0.9999
Yes	59(88.06 %)	4(11.43 %)	
No		31(88.57 %)	
Pathological pattern	39(58.21 %)		0.0971
squamous carcinoma	28(41.79 %)	14(40.00 %)	
Adenocarcinoma and others		21(60.00 %)	
Staging	2(2.99 %)		0.6590
II	44(65.67 %)	2(5.71 %)	
III	21(31.34 %)	20(57.14 %)	
IV		13(37.14 %)	

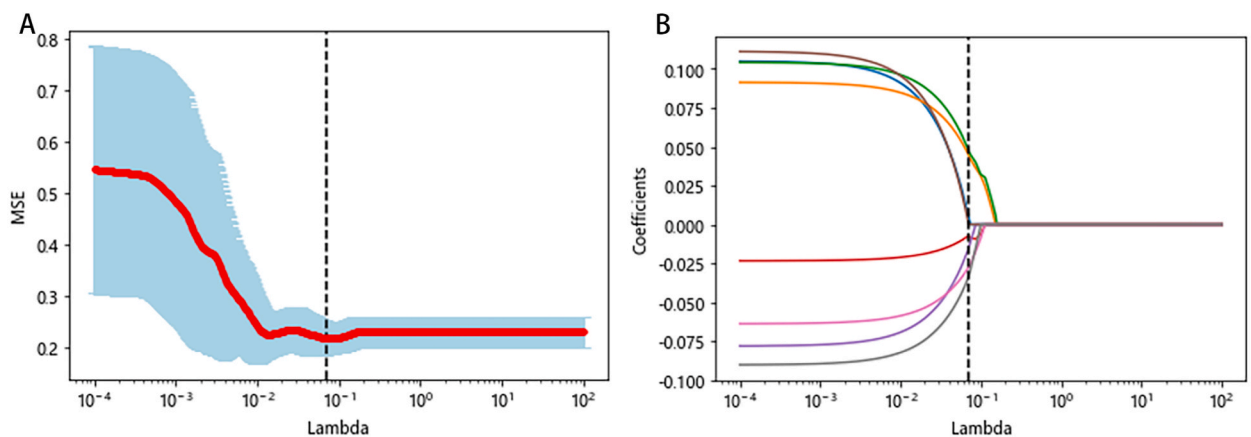


Fig. 3. LASSO Feature Selection Results ($\lambda = 0.0693$)

LASSO regression was employed for feature selection. The parameter (λ) was adjusted using ten rounds of five-fold cross-validation, and bias curves were plotted. The dashed lines indicate the minimum criterion (B) and the 1-SE of the minimum criterion (A). Applying the 1-SE criterion resulted in the selection of eight features, with the optimal value of λ being 0.0693.

Table 3
Radiomics feature screening.

Feature name	Source	Type
log-sigma-4-0-mm-3D:glzm:SmallAreaLowGrayLevelEmphasis	Gauss-Laplace transform	Texture features
log-sigma-6-0-mm-3D:firstorder:RootMeanSquared	Wavelet Transform	Texture features
log-sigma-6-0-mm-3D:glcm:Imc1	Wavelet Transform	Texture features
wavelet-LLH:firstorder:Median	Wavelet Transform	Texture features
wavelet-HLH:glzm:SmallAreaEmphasis	Wavelet Transform	First-order features
wavelet-HHH:firstorder:Kurtosis	Wavelet Transform	Texture features
wavelet-HHH:glcm:ClusterShade	Wavelet Transform	Texture features
wavelet-HHH:glzm:LowGrayLevelZoneEmphasis	Wavelet Transform	Texture features

4.4. Model tuning

4.4.1. Model screening

Four classifiers - logistic regression (LR), support vector machine (SVM), K-nearest neighbors (KNN), and Gaussian naive Bayes (GNB) - were trained on the dataset using eight selected feature values. The AUC value for each model was calculated. The AUC values for SVM and KNN models were relatively low at 0.62 and 0.59, respectively. Comparatively, the LR and GNB models achieved higher AUC values of 0.85 and 0.79. Based on these results, the LR and GNB classifiers were selected for further analysis in this study due to

their better performance.

4.4.2. Training of the model

The selected eight features and the corresponding labeled training set data are brought into the LR, GNB two classifiers to train the tuning model, and the training curves of the corresponding models are obtained (as in Fig. 4).

From the above training curves, we can see that when the training rate increases, i.e., the input training data keeps increasing, the training set scores of both models show a decreasing trend. The test set scores show an increasing trend with some fluctuations, and GNB and LR have a similar trend. When the data are fully input, i.e., up to 1.0 LR training set and test set scores are closest, the test set score is slightly higher than the training set.

4.5. Model test results

Table 4. The remission and non-remission groups were tested using both models, and all results were arithmetically averaged as the corresponding final values after ten times five-fold cross-validation. The corresponding data were entered into the model to generate a confusion matrix to calculate further the sensitivity, specificity, precision, and F1 values.

The model was brought into the test set data to generate the ROC curve (Fig. 5), and its AUC value was measured after ten times five-fold cross-validation. The arithmetic mean of the AUC value was taken as the AUC value of the ROC curve of this model (Table 4). The two algorithms were subjected to Delong' test corresponding to the model. The AUC difference between the two algorithms was statistically significant when comparing $P = 0.045 < 0.05$ between the training set ROC and $P = 0.012 < 0.05$ between the test set ROC.

From the accuracy and AUC values, we can see that for the LR model, the values of the training set and the test set are very close to each other, regardless of the accuracy or AUC values, and the accuracy, AUC values, sensitivity, precision, and F1 values are higher than those of the GNB model; for the specificity, the GNB model is higher than the LR model.

4.6. Model validation results

The calibration curves for the training set and test set of the LR and GNB models are shown in Fig. 6. The curves indicate that both models have large fluctuations in calibration stability. However, the LR model demonstrates better calibration than the GNB model on both the training set and test set. Additionally, the difference between the LR calibration curves on the training and test sets is smaller compared to the GNB model. This suggests the LR model has less variance in performance across datasets.

The decision curves for the training and test sets of the LR and GNB models are presented in Fig. 7. The curves show superior performance of the LR model over GNB on both datasets, with higher net benefit across threshold probabilities.

In summary, the LR model achieves better calibration and clinical utility compared to the GNB model based on the calibration and decision curve analysis.

5. Discussion

The prerequisite for achieving precision medicine is the ability to further classify similar patients based on the current

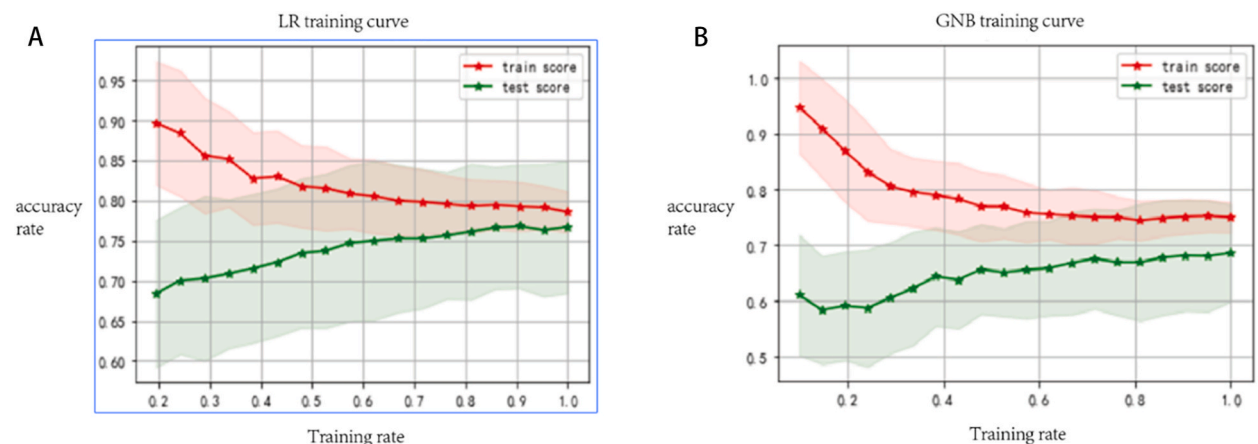


Fig. 4. Training Curves of LR and GNB Algorithms for Mitigation and Non-Mitigation Groups (horizontal axis represents training rates, vertical axis represents the model's accuracy on the training and testing sets at that moment, solid line denotes the average accuracy, and shaded region indicates the standard deviation generated by validation). A corresponds to the training curve of the LR model, and B corresponds to the training curve of the GNB model.

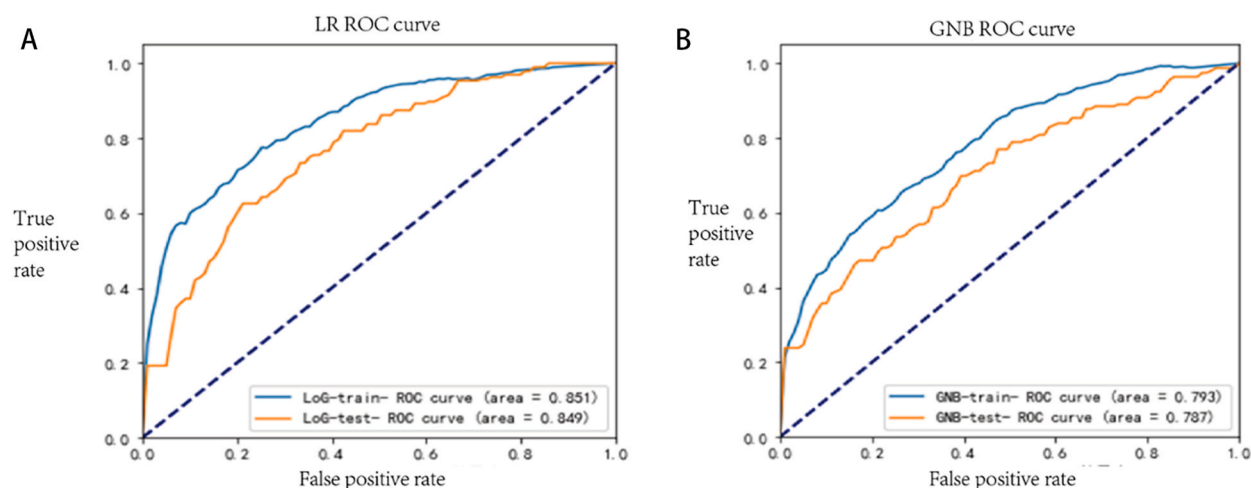


Fig. 5. AUC Values of ROC Curves for Two Algorithms in Mitigation and Non-Mitigation Groups, where Panel A represents the LR model and Panel B represents the GNB model.

Table 4

Two algorithm model results.

model	accuracy rate		AUC		sensitivity	specificity	precision	F1	P*	
	training group	test group	training group	test group						
LR	0.787	0.772	0.851	0.849	0.909-0.667	0.542	0.861	0.686	0.570	0.012
GNB	0.745	0.712	0.793	0.787		0.449	0.902	0.708	0.549	

P*: P-value resulting from the Delong' test performed on the test set ROC of the corresponding model for both algorithms. AUC, Area Under Curve; LR, Logistic Regression; GNB, Gaussian Naive Bayes.

classification. Patients with driver-negative inoperable NSCLC are currently treated with radical radiotherapy combined with platinum-containing chemotherapy as the treatment of choice, but this modality lacks individuality. Conventional pathology and classification factors such as clinical staging do not allow further exploration of patient tumor heterogeneity. They can lead to different prognostic outcomes in patients with seemingly identical pathology and consistent staging, especially in more severe cases. Therefore, non-invasive and easily accessible predictors are needed to delineate patient populations further and are used to estimate the remission rate of NSCLC [18]. With the increasing development of big data and artificial intelligence, various histology concepts have been proposed one after another, and radiomics technology has emerged, accompanied by the mission to address precision medicine (also known as personalized medicine) [19]. Radiomics has been a hotspot and focus of medical research recently and was first conceptualized by Dutch scholar Lambin et al., in 2012. Defined as the high-throughput extraction of a large amount of image information through medical images, which enables image segmentation, feature extraction, and model building and can quantify image heterogeneity caused by changes that are not observed by the human eye, and then through deep data mining, prediction, and analysis thus used to analyze specific information [20–22]. Radiomics aims to break through the limits of the application value of traditional imaging examinations in early diagnosis, efficacy assessment, and prognosis prediction of tumors [23], to provide individualized treatment plans, and to achieve precision medicine. Radiomics has been applied to many types of tumors, but lung cancer is one of the most widely studied and applied malignancies through radiomics [24,25].

In the image segmentation, the ROI area were sketched using 3d slicers software, and two radiation therapists semi-automatically outlined the GTV. After initial segmentation, a chief physician experienced in radiation therapy for thoracic tumors reviewed and modified the results. Multiple physicians checked each other, so the accuracy of the outline could be guaranteed, and the error caused by manual outline could be effectively controlled [26]. we resampled all images to $1 \times 1 \times 1$ mm cubic voxels using trilinear interpolation in 3D Slicer software prior to feature extraction. This voxel size was chosen to achieve isotropic voxels in order to enable standard 3D wavelet-based feature computation across patients. The trilinear interpolation method utilizes neighboring voxels in three dimensions to perform interpolation, which helps minimize resampling artifacts and maintain image continuity compared to lower order interpolation methods. Resampling to isotropic cubic voxels using trilinear interpolation allowed consistent wavelet feature calculation in 3D while preserving image information. For radiomics feature extraction, this study extracts first-order, second-order, and high-order texture features of the original image but also transforms first-order, second-order, and high-order texture features by including wavelet and LoG filter functions. The number of features extracted is as high as 1130, which increases selection possibilities and improves prediction accuracy in feature selection and phase relationship studies. In most articles, the Gaussian filtering parameters of CT images of lung cancer were chosen in the range of 2–6 mm [27,28], and 4–6 mm standard deviation was matched with it to extract details and global features. Fixed three standard deviation parameters can avoid the risk of data overfitting, which is conducive

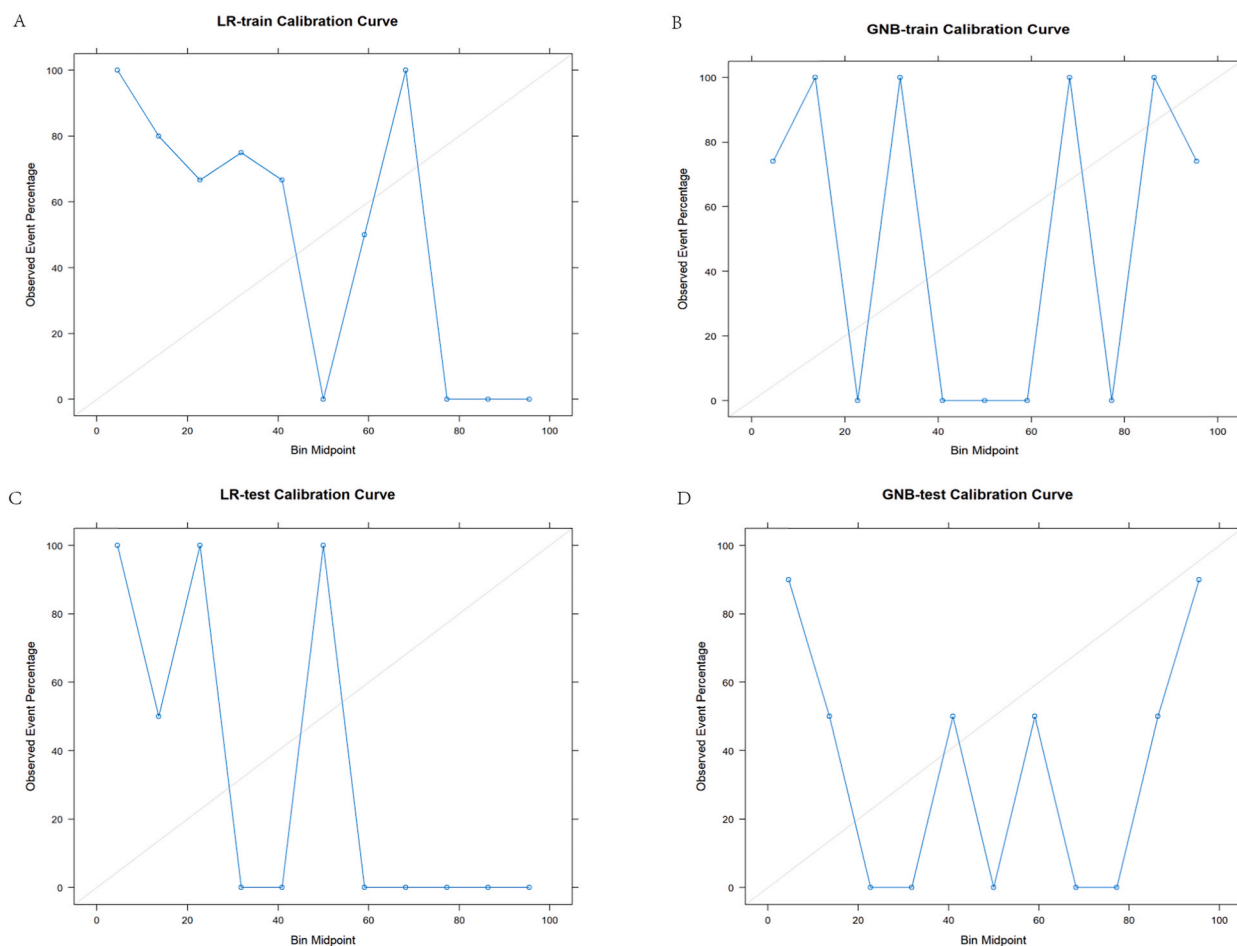


Fig. 6. These are the calibration curves for the logistic regression (LR) and Gaussian naive Bayes (GNB) models, where the closer the curve is to the diagonal line, the more accurate the model is. A is the calibration curve for the training set of the LR model; B is the calibration curve for the test set of the LR model; C is the calibration curve for the training set of the GNB model; And D is the calibration curve for the test set of the GNB model.

to model generalization. The interval between the three parameters is 1 mm, which can balance the feature richness and the computational amount.

For data processing, this study standardized and normalized the data. The study showed that normalized and normalized processing could eliminate the differences in magnitudes between different data and make the prediction results show a better performance [29]. In the feature screening, this study applied the multiple screening of mutual trust and LASSO regression [30,31] in turn to screen the features required for the model, and according to the "one-tenth" principle proposed by Babyak [32], each feature requires at least 10–15 patients, so finally eight features with high correlation coefficients were selected for model training to reduce the risk of overfitting. In this study, we selected 8 radiomic features through multivariate statistical analysis. These features reflect information regarding gray level distribution, texture, and spatial distribution of the lesions, providing an effective feature basis for further pathological grading modeling. In summary, the selected radiomic features comprehensively reflect the imaging characteristics of the lesions, and provide a key basis of feature selection for pathological grading modeling based on CT images. In model construction, both LR and GNB algorithms were used in this study, and the ROC curves of both models were subjected to Delong's test [33] with $P < 0.05$. The AUC differences between them were significantly different. All evaluation indexes were averaged after ten times of five-fold cross-validation as the final. This avoids over- or under-scores of the model due to a random assignment of the training set test set [34]. The 5×10 cross-validation provides a reliable estimate of model performance by reducing variability while avoiding excessive computation compared to other methods like leave-one-out cross-validation. The multiple repetitions help average out randomness in the fold splits. 5 folds was chosen as it offers a good balance between bias and variance. The results showed that LR had only slightly lower specificity than GNB. The sensitivity, accuracy, AUC value, precision, and F1 value were all higher than GNB. After combining the above evaluation indexes, it can be seen that LR is more suitable for predicting whether NSCLC prognosis is in remission without considering the speed of computing (training set accuracy: 0.787, AUC value: 0.851; test set accuracy: 0.772, (AUC value: 0.849). We also plotted calibration curves and decision curves for both the logistic regression (LR) and Gaussian naive Bayes (GNB) models on both the test and validation sets to assess the calibration and net benefit of the comparative models. The results showed that LR performed

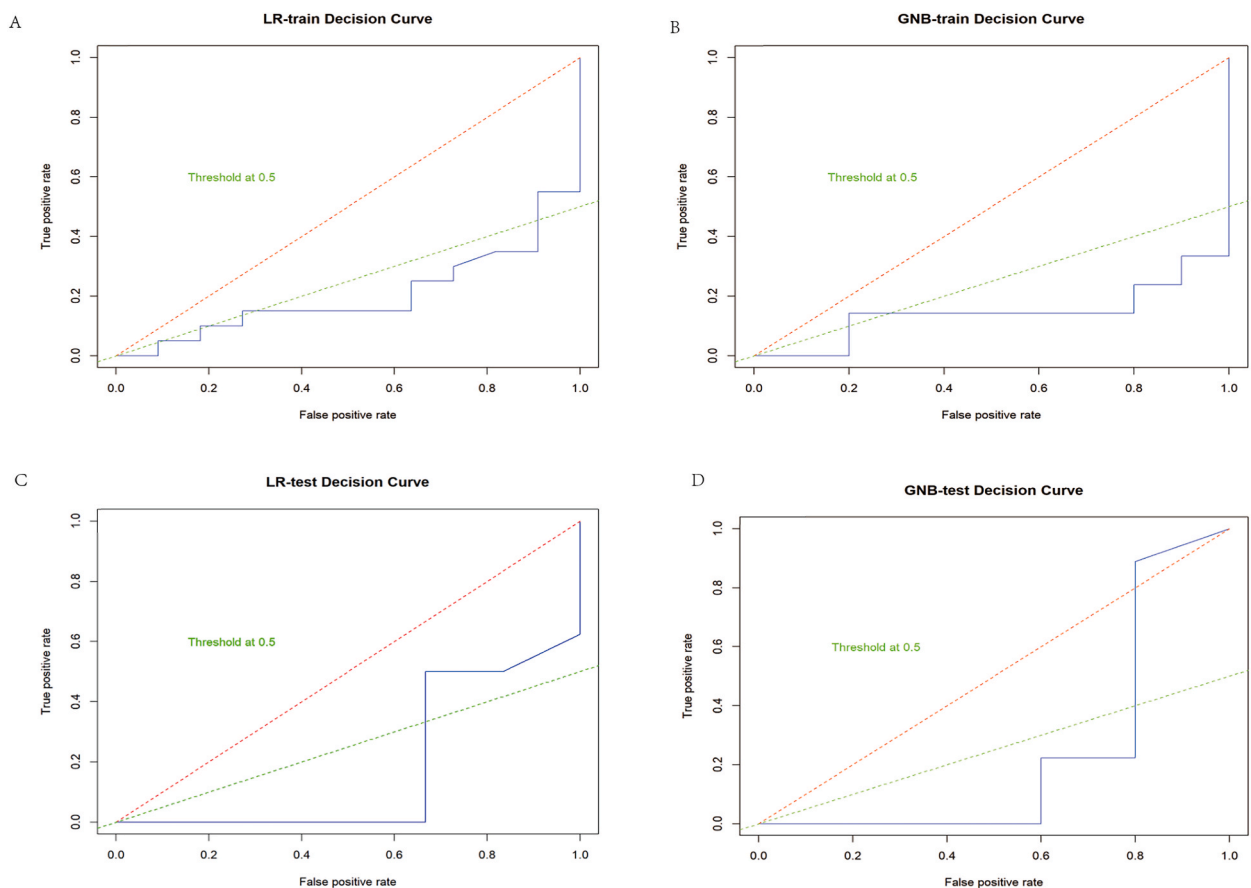


Fig. 7. These are the decision curves for the logistic regression (LR) and Gaussian naive Bayes (GNB) models. The area below the decision curves represents all the net benefits brought by the model when combining different thresholds. The larger the area is, the stronger the model's ability to improve the overall decision-making effect. The green dashed line is the 0.5 threshold line. A is the decision curve for the training set of the LR model; B is the decision curve for the test set of the LR model; C is the decision curve for the training set of the GNB model; D is the decision curve for the test set of the GNB model.

better in terms of model consistency, calibration, and net benefit. Yuto Sugai [35] et al. used COX to construct a prediction model based on CT images by LASSO screening features and c-dex and Kaplan-Meier curves to model and evaluate the prognosis of NSCLC patients, which is similar to the present study. However, the study used one physician to perform manual segmentation, making reducing the impact generated by subjective errors challenging. The study used COX to construct a prediction model, which often produces overfitting due to the Cox model that directly predicts the time to event with a simple regression equation. This study established two arithmetic prediction models, LR and GNB, using multi-person outlining and checking. ROC curves were drawn based on the enhanced CT of patients before treatment, and the sensitivity, specificity, accuracy, and F1 values of the two models were calculated, which can help reduce the errors generated by manual outlining and adopt a more stable prediction model with a more complex evaluation mechanism and a better evaluation effect, which can be used before patient treatment. The prediction model is used to screen out patients with less than ideal treatment effects and guide the clinic to add subsequent treatment to this part of patients based on the current radiotherapy and chemotherapy mode of treatment.

However, there are still some limitations in this study. First, this study uses a dichotomous classification method, which is more limited [36,37], with a small sample size and a single-center research scheme. It is intended to add multicenter data in later studies to increase the sample size and improve the model's accuracy.

Based on this study, the next intention is to add several algorithms currently commonly used at home and abroad for multi-classification comparison, and also to introduce deep learning methods for comparison, and then expand to multimodality (e.g., adding MRI or PET/CT) [38–41] for a more comprehensive and in-depth study, in anticipation of developing clinical translation tools with better results.

6. Conclusion

In conclusion, predicting the efficacy of radiotherapy and chemotherapy in NSCLC patients based on pre-treatment-enhanced CT radiomics has some predictive value. This study successfully constructed a radiomics-based efficacy prediction model through which the efficacy of NSCLC patients can be predicted before radiotherapy and chemotherapy. If patients were predicted to be in non-remission by the prediction model, the current radiotherapy and chemotherapy modalities were considered to have a limited therapeutic effect, and possible further treatment modalities could be explored to improve the outcome.

Data availability

Given that the data in this study originate from a hospital, and in accordance with relevant laws and hospital policies, we are obliged to protect the privacy and confidentiality of patient data. Consequently, these data cannot be deposited into a publicly accessible repository. However, we have made every effort to ensure that the study results and analyses described in the article are understandable and verifiable by readers, and that all results are supported by the references listed.

CRedit authorship contribution statement

Hanjing Zhang: Writing - review & editing, Writing - original draft, Validation, Software, Resources, Investigation, Data curation, Conceptualization. **Yu Deng:** Writing - original draft, Investigation, Data curation, Conceptualization. **M.A. Xiaojie:** Supervision. **Qian Zou:** Software, Data curation. **Huanhui Liu:** Data curation. **Ni Tang:** Data curation. **Yuanyuan Luo:** Data curation. **Xuejing Xiang:** Data curation.

Declaration of competing interest

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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